# Joint Estimation and Inference for Multiple Multi-layered Gaussian Graphical Models

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Abstract: The rapid development of high-throughput technologies has enabled generation of data from biological processes that span multiple layers, like genomic, proteomic or metabolomic data; and pertain to multiple sources, like disease subtypes or experimental conditions. In this work we propose a general statistical framework based on graphical models for horizontal (i.e. across conditions or subtypes) and vertical (i.e. across different layers containing data on molecular compartments) integration of information in such datasets. We start with decomposing the multi-layer problem into a series of two-layer problems. For each two-layer problem, we model the outcomes at a node in the lower layer as dependent on those of other nodes in that layer, as well as all nodes in the upper layer. Following the biconvexity of our objective function, this estimation problem decomposes into two parts, where we use neighborhood selection and subsequent refitting of the precision matrix to quantify the dependency of two nodes in a single layer, and use grouppenalized least square estimation to quantify the directional dependency of two nodes in different layers. Finally, to test for differences in these directional dependencies across multiple sources, we devise a hypothesis testing procedure that utilizes already computed neighborhood selection coefficients for nodes in the upper layer. We establish theoretical results for the validity of this testing procedure and the consistency of our estimates, and also evaluate their performance through simulations and a real data application.

**Keywords**: Data integration; Gaussian Graphical Models; Neighborhood selection; Group lasso

#### 1 Introduction

#### 1.1 Notations

We denote scalars by small letters, vectors by bold small letters and matrices by bold capital letters. For any matrix  $\mathbf{A}$ ,  $(\mathbf{A})_{ij}$  denote its element in the  $(i,j)^{\text{th}}$  position. For  $a,b \in \mathbb{N}$ , we denote the set of all  $a \times b$  real matrices by  $\mathbb{M}(a,b)$ . For any positive integer c, define  $\mathcal{I}_c = \{1,\ldots,c\}$ .

# 2 The Joint Multiple Multilevel Estimation Framework

#### 2.1 Formulation

Suppose there are K independent datasets, each pertaining to an M-layered Gaussian Graphical Model (GGM). The k<sup>th</sup> model has the following structure:

Layer 1- 
$$\mathbb{D}_1^k = (D_{11}^k, \dots, D_{1p_1}^k) \sim \mathcal{N}(0, \Sigma_1^k); \quad k \in \mathcal{I}_K,$$
 Layer  $m \ (1 < m \le M)$ - 
$$\mathbb{D}_m^k = \mathbb{D}_{m-1}^k \mathbf{B}_m^k + \mathbb{E}_m^k, \text{ with } \mathbf{B}_m^k \in \mathbb{M}(p_{m-1}, p_m)$$
 and 
$$\mathbb{E}_m^k = (E_{m1}^k, \dots, E_{mp_m}^k) \sim \mathcal{N}(0, \Sigma_m^k); \quad k \in \mathcal{I}_K.$$

We assume known structured sparsity patterns, denoted by  $\mathcal{G}_m$  and  $\mathcal{H}_m$ , for the parameters of interest in the above model, i.e. the precision matrices  $\Omega_m^k := (\Sigma_m^k)^{-1}$  and the regression coefficient matrices  $\mathbf{B}_m^k$ , respectively. These patterns provide information on horizontal dependencies across k for the corresponding parameters, and our goal here is to leverage them to estimate the full hierarchical structure of the network- specifically to obtain the undirected edges inside nodes of a single layer, and the directed edges between two successive layers through jointly estimating  $\{\Omega_m^k\}$  and  $\{\mathbf{B}_m^k\}$ .

Consider the following two-layer model, which is a special case of the above model with M=2:

$$\mathbb{X}^k = (X_1^k, \dots, X_p^k)^T \sim \mathcal{N}(0, \Sigma_x^k); \tag{2.1}$$

$$\mathbb{Y}^k = \mathbb{X}^k \mathbf{B}^k + \mathbb{E}^k; \quad \mathbb{E}^k = (E_1^k, \dots, E_p^k)^T \sim \mathcal{N}(0, \Sigma_y^k); \tag{2.2}$$

$$\mathbf{B}^k \in \mathbb{M}(p,q), \quad \Omega_x^k = (\Sigma_x^k)^{-1}; \quad \Omega_y^k = (\Sigma_y^k)^{-1}; \tag{2.3}$$

where we want to estimate  $\{(\Omega_x^k, \Omega_y^k, \mathbf{B}^k); k \in \mathcal{I}_K \text{ from data } \mathcal{Z}^k = \{(\mathbf{Y}^k, \mathbf{X}^k); \mathbf{Y}^k \in \mathbb{M}(n,q), \mathbf{X}^k \in \mathbb{M}(n,p), k \in \mathcal{I}_K\}$ . in presence of known grouping structures  $\mathcal{G}_x, \mathcal{G}_y, \mathcal{H}_x$  respectively. We focus most of the theoretical discussion in the rest of the paper on jointly estimating  $\Omega_y := \{\Omega_y^k\}$  and  $\mathcal{B} := \{\mathbf{B}^k\}$ . This is because the parameters  $\Omega_x := \{\Omega_x^k\}$  are only dependent on  $\{\mathbf{X}^k\}$ , so any method for joint estimation of multiple graphical models can be used to estimate them (e.g. Ma and Michailidis (2016); ?). On the other hand for M > 2, xyz. Thus a joint estimation method for  $\Omega_y$  and  $\mathcal{B}$  provides all building blocks for estimating the full hierarchical structure of our M-layered multiple GGMs.

#### 2.2 Algorithm

Estimation of  $\{\Omega_x^k\}$  done using JSEM. For the other part, we use the following two-step procedure:

1. Run neighborhood selection on y-network incorporating effects of x-data and an additional blockwise group penalty:

$$\min_{\mathcal{B},\Theta} \left\{ \sum_{j=1}^{q} \frac{1}{n_k} \left[ \sum_{k=1}^{K} \|\mathbf{Y}_j^k - (\mathbf{Y}_{-j}^k - \mathbf{X}^k \mathbf{B}_{-j}^k) \boldsymbol{\theta}_j^k - \mathbf{X}^k \mathbf{B}_j^k \|^2 + \sum_{j \neq i} \sum_{g \in \mathcal{G}_y^{ij}} \lambda_{ij}^g \|\boldsymbol{\theta}_{ij}^{[g]}\| \right] + \sum_{b \in \mathcal{G}_x \times \mathcal{G}_y \times \mathcal{H}} \eta^b \|\mathbf{B}^{[b]}\| \right\}$$

$$= \min \left\{ f(\mathcal{Y}, \mathcal{X}, \mathcal{B}, \Theta) + P(\Theta) + Q(\mathcal{B}) \right\}$$

$$(2.5)$$

where 
$$\Theta = {\Theta_i}, \mathcal{B} = {\mathbf{B}^k}, \mathcal{Y} = {\mathbf{Y}^k}, \mathcal{X} = {\mathbf{X}^k}, \mathcal{E} = {\mathbf{E}^k}.$$

This estimates  $\mathcal{B}$  (possibly refit and/or within-group threshold).

2. Step I part 2 and step II of JSEM (see 15-656 pg 6) follows to estimate  $\{\Omega_v^k\}$ .

The objective function is bi-convex, so we are going to do the following in step 1-

- Start with initial estimates of  $\mathcal{B}$  and  $\Theta$ , say  $\mathcal{B}^{(0)}$ ,  $\Theta^{(0)}$ .
- Iterate:

$$\Theta^{(t+1)} = \arg\min\left\{f(\mathcal{Y}, \mathcal{X}, \mathcal{B}^{(t)}, \Theta^{(t)}) + P(\Theta^{(t)})\right\}$$
(2.6)

$$\mathcal{B}^{(t+1)} = \arg\min\left\{f(\mathcal{Y}, \mathcal{X}, \mathcal{B}^{(t)}, \Theta^{(t+1)}) + Q(\mathcal{B}^{(t)})\right\}$$
(2.7)

• Continue till convergence.

#### 2.3 Theoretical properties

#### 2.3.1 Conditions

Conditions A1 from JSEM paper holds for  $\mathcal{X}$  and  $\mathcal{E}$ . Also A2, A3 from JSEM paper.

To prove the results in this section, we use a reparametrization of the neighborhood coefficients at the lower level. Specifically, notice that for  $j \in \mathcal{I}_q, k \in \mathcal{I}_K$ , the corresponding summand in  $f(\mathcal{Y}, \mathcal{X}, \mathcal{B}, \Theta)$  can be rearranged as

$$\begin{aligned} \|\mathbf{Y}_{j}^{k} - \mathbf{X}^{k}\mathbf{B}_{j}^{k} - (\mathbf{Y}_{-j}^{k} - \mathbf{X}^{k}\mathbf{B}_{-j}^{k})\boldsymbol{\theta}_{j}^{k}\|^{2} &= \|\mathbf{Y}_{j}^{k} - \mathbf{Y}_{-j}^{k}\boldsymbol{\theta}_{j}^{k} - (\mathbf{X}^{k}\mathbf{B}_{j}^{k} - \mathbf{X}^{k}\mathbf{B}_{-j}^{k}\boldsymbol{\theta}_{j}^{k})\|^{2} \\ &= \|(\mathbf{Y} - \mathbf{X}\mathbf{B})\mathbf{T}_{j}^{k}\|^{2} \end{aligned}$$

where

$$T_{jj'}^k = \begin{cases} 1 \text{ if } j = j' \\ -\theta_{jj'}^k \text{ if } j \neq j' \end{cases}$$

Thus, with  $\mathbf{T}^k := (\mathbf{T}_j^k)_{j \in \mathcal{I}_q}$ , we have

$$f(\mathcal{Y}, \mathcal{X}, \mathcal{B}, \Theta) = \frac{1}{n} \sum_{i=1}^{p} \sum_{k=1}^{K} \| (\mathbf{Y}^k - \mathbf{X}^k \mathbf{B}^k) \mathbf{T}_j^k \|^2 = \frac{1}{n} \sum_{k=1}^{K} \| \mathbf{Y}^k - \mathbf{X}^k \mathbf{B}^k) \mathbf{T}^k \|_F^2 = \sum_{k=1}^{K} \operatorname{Tr}(\mathbf{S}^k (\mathbf{T}^k)^2)$$

where  $\mathbf{S}^k = (1/n)(\mathbf{Y}^k - \mathbf{X}^k \mathbf{B}^k)(\mathbf{Y}^k - \mathbf{X}^k \mathbf{B}^k)^T$  is the sample covariance matrix.

Now suppose  $\beta = \text{vec}(\mathbf{B})$ , and any subscript or superscript on **B** is passed on to  $\beta$ . Denote by  $\hat{\boldsymbol{\beta}}$  and  $\hat{\boldsymbol{\Theta}}$  the generic estimators given by

$$\widehat{\boldsymbol{\beta}} = \underset{\boldsymbol{\beta} \in \mathbb{R}^{pq}}{\min} \left\{ -2\boldsymbol{\beta}^T \widehat{\boldsymbol{\gamma}} + \boldsymbol{\beta}^T \widehat{\boldsymbol{\Gamma}} \boldsymbol{\beta} + \lambda_n \sum_{h \in \mathcal{H}} \|\boldsymbol{\beta}^{[h]}\| \right\}$$
(2.8)

$$\widehat{\Theta}_{j} = \underset{\Theta_{j} \in \mathbb{M}(q-1,K)}{\operatorname{arg\,min}} \left\{ \frac{1}{n} \sum_{k=1}^{K} \|\mathbf{Y}_{j}^{k} - \mathbf{X}^{k} \widehat{\mathbf{B}}_{j}^{k} - (\mathbf{Y}_{-j}^{k} - \mathbf{X}^{k} \widehat{\mathbf{B}}_{-j}^{k}) \boldsymbol{\theta}_{j}^{k} \|^{2} + \gamma_{n} \sum_{j \neq j'} \sum_{g \in \mathcal{G}_{y}^{jj'}} \|\boldsymbol{\theta}_{jj'}^{[g]}\| \right\}$$
(2.9)

where

$$\widehat{\boldsymbol{\Gamma}} = \begin{bmatrix} (\widehat{\mathbf{T}}^1)^2 \otimes \frac{(\mathbf{X}^1)^T \mathbf{X}^1}{n} & & \\ & \ddots & \\ & & (\widehat{\mathbf{T}}^K)^2 \otimes \frac{(\mathbf{X}^K)^T \mathbf{X}^K}{n} \end{bmatrix}; \quad \widehat{\boldsymbol{\gamma}} = \begin{bmatrix} (\widehat{\mathbf{T}}^1)^2 \otimes \frac{(\mathbf{X}^1)^T}{n} \\ \vdots \\ (\widehat{\mathbf{T}}^K)^2 \otimes \frac{(\mathbf{X}^K)^T}{n} \end{bmatrix} \begin{bmatrix} \operatorname{vec}(\mathbf{Y}^1) \\ \vdots \\ \operatorname{vec}(\mathbf{Y}^K) \end{bmatrix}$$

with  $\widehat{\mathbf{T}}^k$  defined the same way using  $\widehat{\boldsymbol{\theta}}_i^k$  as we defined  $\mathbf{T}^k$  using  $\boldsymbol{\theta}_i^k$ .

**Theorem 2.1.** Assume fixed  $\mathcal{X}, \mathcal{E}$  and deterministic  $\widehat{\mathcal{B}} = {\{\widehat{\mathbf{B}}^k\}}$ . Also for k = 1, ..., K,

(T1) 
$$\|\widehat{\mathbf{B}}^k - \mathbf{B}_0^k\|_1 \le v_\beta$$
, where  $v_\beta = \eta_\beta \sqrt{\frac{\log(pq)}{n}}$  with  $\eta_\beta \ge 0$  depending on  $\mathcal{B}$  only;

(T2) Denote 
$$\hat{\mathbf{E}}^k = \mathbf{Y}^k - \mathbf{X}^k \hat{\mathbf{B}}^k, k \in \mathcal{I}_K$$
. Then for all  $j \in \mathcal{I}_q$ ,

$$\frac{1}{n} \left\| (\widehat{\mathbf{E}}_{-j}^k)^T \widehat{\mathbf{E}}^k \mathbf{T}_{0,j}^k \right\|_{\infty} \leq \mathbb{Q} \left( v_{\beta}, \Sigma_x^k, \Sigma_y^k \right)$$

where  $\mathbb{Q}\left(v_{\beta}, \Sigma_{x}^{k}, \Sigma_{y}^{k}\right)$  is a  $O(\sqrt{\log(pq)/n})$  deterministic function of  $\mathcal{B}, \Sigma_{x}^{k}$  and  $\Sigma_{y}^{k}$ . (T3) Denote  $\widehat{\mathbf{S}}^{k} = (\widehat{\mathbf{E}}^{k})^{T} \widehat{\mathbf{E}}^{k}/n$ . Then  $\widehat{\mathbf{S}}^{k} \sim RE(\psi^{k}, \phi^{k})$  with  $Kq\phi \leq \psi/2$  where  $\psi = (\mathbf{E}^{k})^{T} \widehat{\mathbf{E}}^{k}/n$ .  $\min_k \psi^k, \phi = \max_k \phi^k;$ 

(T4) Assumption (A2) holds for  $\Sigma_n^k$ 

Then, given the choice of tuning parameter

$$\gamma_n = 4\sqrt{|g_{\max}|}\mathbb{Q}_0; \quad \mathbb{Q}_0 := \max_{k \in \mathcal{I}_K} \mathbb{Q}\left(v_\beta, \Sigma_x^k, \Sigma_y^k\right)$$

the following holds

$$\frac{1}{K} \sum_{k=1}^{K} \|\widehat{\Omega}_{y}^{k} - \Omega_{y}^{k}\|_{F} \le O\left(\mathbb{Q}_{0} \sqrt{\frac{|g_{\max}|S}{K}}\right)$$

where  $|g_{\text{max}}|$  is the maximum group size.

When  $\mathcal{X}$  and  $\mathcal{E}$  are random, the following propositions ensures that conditions (T2) and (T3) hold with probabilities approaching to 1.

**Proposition 2.2.** Consider deterministic  $\widehat{\mathcal{B}}$  satisfying assumption (T1). Then for sample size  $n \succsim \log(pq)$  and  $k \in \mathcal{I}_K$ ,

1.  $\widehat{\mathbf{S}}^k$  satisfies the RE condition:  $\widehat{\mathbf{S}}^k \sim RE(\psi^k, \phi^k)$ , where

$$\psi^k = \frac{\Lambda_{\min}(\Sigma_x^k)}{2}; \quad \phi^k = \frac{\psi^k \log p}{n} + 2v_\beta c_2 [\Lambda_{\max}(\Sigma_x^k) \Lambda_{\max}(\Sigma_y^k)]^{1/2} \sqrt{\frac{\log(pq)}{n}}$$

with probability  $\geq 1 - 6c_1 \exp[-(c_2^2 - 1)\log(pq)] - 2\exp(-c_3n), c_1, c_3 > 0, c_2 > 1.$ 

2. The following deviation bound is satisfied for any  $j \in \mathcal{I}_q$ 

$$\left\| \frac{1}{n} (\widehat{\mathbf{E}}_{-j}^k)^T \widehat{\mathbf{E}}^k \mathbf{T}_{0,j}^k \right\|_{\infty} \leq \mathbb{Q} \left( v_{\beta}, \Sigma_x^k, \Sigma_y^k \right)$$

with probability  $\geq 1 - 1/p^{\tau_1 - 2} - 6c_1 \exp[-(c_2^2 - 1)\log(pq)] - 6c_4 \exp[-(c_5^2 - 1)\log(pq)], c_4 > 0, c_5 > 1, \text{ where}$ 

$$\mathbb{Q}\left(v_{\beta}, \Sigma_{x}^{k}, \Sigma_{y}^{k}\right) = 2v_{\beta}^{2} V_{x}^{k} + 4v_{\beta} c_{2} \left[\Lambda_{\max}(\Sigma_{x}^{k}) \Lambda_{\max}(\Sigma_{y}^{k})\right]^{1/2} \sqrt{\frac{\log(pq)}{n}} + c_{5} \left[\Lambda_{\max}(\Sigma_{y,-j}^{k}) \sigma_{y,j,-j}^{k}\right]^{1/2} \sqrt{\frac{\log q}{n}}$$

with  $\sigma_{y,j,-j}^k = \mathbb{V}(E_j - \mathbb{E}_{-j}\boldsymbol{\theta}_{0,j})$ , and

$$V_x^k = \sqrt{\frac{\log 4 + \tau_1 \log p}{c_x^k n}} + \max_i \sigma_{x,ii}^k; \quad c_x^k = \left[128(1 + 4\Lambda_{\max}(\Sigma_x))^2 \max_i (\sigma_{x,ii})^2\right]^{-1}$$

The error bounds for  $\widehat{\Omega}_{y}^{k}$ ,  $k \in \mathcal{I}_{K}$  follow immediately from the above two results.

Corollary 2.3. Consider any deterministic  $\widehat{\mathcal{B}}$  that satisfy the following bound

$$\|\widehat{\mathbf{B}}^k - \mathbf{B}_0^k\|_1 \le v_\beta = \eta_\beta \sqrt{\frac{\log(pq)}{n}}$$

Then, for sample size  $n \gtrsim \log(pq)$  and choice of tuning parameter  $\gamma_n = 4\sqrt{|g_{\text{max}}|}\mathbb{Q}_0$ , there exist constants  $c_1, c_3, c_4 > 0, c_2, c_5 > 1$  such that the following holds

$$\frac{1}{K} \sum_{k=1}^{K} \|\widehat{\Omega}_y^k - \Omega_y^k\|_F \le O\left(\mathbb{Q}_0 \sqrt{\frac{|g_{\text{max}}|S}{K}}\right) \tag{2.10}$$

with probability  $\geq 1 - 1/p^{\tau_1 - 2} - 6c_1 \exp[-(c_2^2 - 1)\log(pq)] - 2\exp(-c_3n) - 6c_4 \exp[-(c_5^2 - 1)\log(pq)].$ 

#### Discuss tighter bound compared to vanilla JSEM

After providing the error bounds for solutions to the subproblem (2.9), we concentrate on the subproblem (2.8). Following a similar strategy, we first get error bounds for  $\hat{\beta}$  assuming everything else fixed.

**Theorem 2.4.** Assume fixed  $\mathcal{X}, \mathcal{E}$ , and deterministic  $\widehat{\Theta} = {\{\widehat{\Theta}_j\}}$ , so that for  $j \in \mathcal{I}_q$ ,

**(B1)**  $\|\widehat{\Theta}_j - \Theta_{0,j}\|_F \le v_{\Theta} \sqrt{\frac{\log q}{n}}$  for some  $v_{\Theta}$  dependent on  $\Theta$ .

**(B2)** Denote  $\widehat{\mathbf{\Gamma}}^k = (\widehat{\mathbf{T}}^k)^2 \otimes (\mathbf{X}^k)^T \mathbf{X}^k / n$ ,  $\widehat{\boldsymbol{\gamma}}^k = (\widehat{\mathbf{T}}^k)^2 \otimes (\mathbf{X}^k)^T \mathbf{Y}^k / n$ . Then the deviation bound holds:

$$\left\| \widehat{\boldsymbol{\gamma}}^k - \widehat{\boldsymbol{\Gamma}}^k \boldsymbol{\beta}_0 \right\|_{\infty} \leq \mathbb{R}(v_{\Theta}, \Sigma_x^k, \Sigma_y^k) \sqrt{\frac{\log(pq)}{n}}$$

where  $\mathbb{R}\left(v_{\Theta}, \Sigma_{x}^{k}, \Sigma_{y}^{k}\right)$  is a O(1) deterministic function of  $\Theta, \Sigma_{x}^{k}$  and  $\Sigma_{y}^{k}$ .

**(B3)**  $\widehat{\Gamma} \sim RE(\psi_*, \phi_*)$  with  $Kpq\phi_* \leq \psi_*/2$ .

Then, given the choice of tuning parameter

$$\lambda_n \ge 4\sqrt{|h_{\max}|} \mathbb{R}_0 \sqrt{\frac{\log(pq)}{n}}; \quad \mathbb{R}_0 := \max_{k \in \mathcal{I}_K} \mathbb{R}\left(v_{\Theta}, \Sigma_x^k, \Sigma_y^k\right)$$

the following holds

$$\|\widehat{\boldsymbol{\beta}} - \boldsymbol{\beta}_0\|_1 \le 48\sqrt{|h_{\text{max}}|} s_{\beta} \lambda_n / \psi^* \tag{2.11}$$

$$\|\widehat{\boldsymbol{\beta}} - \boldsymbol{\beta}_0\| \le 12\sqrt{s_\beta}\lambda_n/\psi^* \tag{2.12}$$

$$\sum_{h \in \mathcal{H}} \|\boldsymbol{\beta}^{[h]} - \boldsymbol{\beta}_0^{[h]}\| \le 48s_{\beta} \lambda_n / \psi^* \tag{2.13}$$

$$(\widehat{\boldsymbol{\beta}} - \boldsymbol{\beta}_0)^T \widehat{\boldsymbol{\Gamma}}(\widehat{\boldsymbol{\beta}} - \boldsymbol{\beta}_0) \le 72s_{\beta} \lambda_n^2 / \psi^* \tag{2.14}$$

Next we verify that conditions (B2) and (B3) hold with high probability given fixed  $\widehat{\Theta}$ .

**Proposition 2.5.** Consider deterministic  $\widehat{\Theta}$  satisfying assumption (B1). Assume that the matrices  $(\widehat{\mathbf{T}}^k)^2, k \in \mathcal{I}_K$  are diagonally dominant. Then for sample size  $n \succeq \log(pq)$ ,

1.  $\widehat{\Gamma}$  satisfies the RE condition:  $\widehat{\Gamma} \sim RE(\psi_*, \phi_*)$ , where

$$\psi_* = \min_k \psi^k \left( \min_i \psi_t^i - dv_\Theta \right), \phi_* = \max_k \phi^k \left( \min_i \phi_t^i + dv_\Theta \right)$$

with probability  $\geq 1 - 2\exp(c_3 n), c_3 > 0$ .

2. The deviation bound in (B2) is satisfied with probability  $\geq 1-12c_1 \exp[(c_2^2-1)\log(pq)], c_1 > 0, c_2 > 1$ , where

$$\mathbb{R}\left(v_{\Theta}, \Sigma_{x}^{k}, \Sigma_{y}^{k}\right) = c_{2}\sqrt{\Lambda_{\max}(\Sigma_{x}^{k})}\left(dv_{\Theta}\Lambda_{\min}(\Sigma_{y}^{k}) + \frac{1}{\Lambda_{\min}(\Sigma_{y}^{k})}\right)$$

We now put both the pieces together, and prove that our alternating algorithm results in a solution sequence  $\{\widehat{\mathcal{B}}^{(r)}, \widehat{\Theta}^{(r)}\}, r = 1, 2, \ldots$  that lies uniformly within a non-expanding ball around the true parameter values.

#### $\mathbf{3}$ Two-sample testing

Suppose there are two disease subtypes: k = 1, 2, and we are interested in testing whether the downstream effect of a predictor in X-data is same across both subtypes, i.e. if  $\mathbf{b}_{0i}^1 = \mathbf{b}_{0i}^2$  for some fixed  $i \in \mathcal{I}_p$ . For this we consider the modified optimization problem:

$$\min_{\mathcal{B},\Theta} \frac{1}{n} \left\{ \sum_{j=1}^{q} \sum_{k=1}^{2} \|\mathbf{Y}_{j}^{k} - \mathbf{Y}_{-j}^{k} \boldsymbol{\theta}_{j}^{k} - \mathbf{X}^{k} \mathbf{B}_{j}^{k} \|^{2} + \sum_{j \neq j'} \lambda_{jj'} \|\boldsymbol{\theta}_{jj'}^{*}\| + \sum_{i=1}^{p} \eta_{i} \|\mathbf{B}_{i*}^{*}\| \right\}$$

$$= \min \left\{ f(\mathcal{Y}, \mathcal{X}, \mathcal{B}, \Theta) + P(\Theta) + Q(\mathcal{B}) \right\} \tag{3.1}$$

with  $n_1 = n_2 = n$  for simplicity; and  $\mathbf{B}^k = (\mathbf{b}_1^k, \dots, \mathbf{b}_q^k), (\mathbf{B}_{i*}^*) \in \mathbb{R}^{q \times K}$ 

In this setup, define the desparsified estimate of  $\mathbf{b}_{0i}^{k}$  as

$$\widehat{\mathbf{c}}_{i}^{k} = \widehat{\mathbf{b}}_{i}^{k} + \frac{1}{nt_{i}^{k}} \left( \mathbf{X}_{i}^{k} - \mathbf{X}_{-i}^{k} \widehat{\boldsymbol{\zeta}}_{i}^{k} \right)^{T} \left( \mathbf{Y}^{k} - \mathbf{X}^{k} \widehat{\mathbf{B}}^{k} \right)$$
(3.2)

for k = 1, 2, where  $t_i^k = (\mathbf{X}_i^k - \mathbf{X}_{-i}^k \hat{\boldsymbol{\zeta}}_i^k)^T \mathbf{X}_{-i}^k / n$ . Then we have the asymptotic joint distribution of a scaled version of the debiased coefficients for the  $i^{th}$  predictor effect.

**Theorem 3.1.** Define  $\hat{s}_i^k = \sqrt{\|\mathbf{X}_i^k - \mathbf{X}_{-i}^k \hat{\zeta}_i^k\|^2/n}$ , and  $m_i^k = \sqrt{n} t_i^k / \hat{s}_i^k$ . Assume the following lowing:

- (C1) For the X-neighborhood estimators we have  $\|\hat{\zeta}^k \zeta_0^k\|_1 \le v_\zeta = \eta_\zeta \sqrt{\frac{\log p}{n}}$ .
- (C2) The precision matrix estimators satisfy

$$\left\| (\widehat{\Omega}_y^k)^{1/2} - (\Omega_y^k)^{1/2} \right\|_{\infty} \le v_{\Omega} = \eta_{\Omega} \sqrt{\frac{\log q}{n}}$$

(C3)  $\|\widehat{\mathbf{B}}^k - \mathbf{B}_0^k\|_1 \leq v_{\beta}$ , where  $v_{\beta} = \eta_{\beta} \sqrt{\frac{\log(pq)}{n}}$  with  $\eta_{\beta}$  depending on  $\mathcal{B}$  only. Then for the debiased estimators in (3.2) and sample size satisfying  $n \gtrsim \log(pq), \log p = 1$ 

 $o(n^{1/2}), \log q = o(n^{1/2})$  we have

$$\begin{bmatrix} \widehat{\Omega}_y^1 & \\ & \widehat{\Omega}_y^2 \end{bmatrix}^{1/2} \begin{bmatrix} m_i^1 (\widehat{\mathbf{c}}_i^1 - \mathbf{b}_{0i}^1) & \\ & m_i^2 (\widehat{\mathbf{c}}_i^2 - \mathbf{b}_{0i}^2) \end{bmatrix} \sim \mathcal{N}_{2q}(\mathbf{0}, \mathbf{I}) + o_P(1)$$
(3.3)

Based on the theorem, we have the global test for the effect of the  $i^{th}$  X-covariate.

**Algorithm 1.** (Global test for  $H_0^i: \mathbf{b}_{0i}^1 = \mathbf{b}_{0i}^2$  at level  $\alpha, 0 < \alpha < 1$ )

- 1. Obtain the debiased estimators  $\hat{\mathbf{c}}_i^1, \hat{\mathbf{c}}_i^2$  using (3.2).
- 2. Calculate the test statistic

$$D_i = (\widehat{\mathbf{c}}_i^1 - \widehat{\mathbf{c}}_i^2)^T \left( \frac{(\widehat{\Omega}_y^1)^{-1}}{(m_i^1)^2} + \frac{(\widehat{\Omega}_y^2)^{-1}}{(m_i^2)^2} \right)^{-1} (\widehat{\mathbf{c}}_i^1 - \widehat{\mathbf{c}}_i^2)$$

3. Reject  $H_0^i$  if  $D_i \geq \chi_{q,1-\alpha}^2$ .

Given the null hypothesis is rejected, we now consider the multiple testing problem of simultaneously testing for all entrywise differences, i.e. testing

$$H_0^{ij}: b_{0ij}^1 = b_{0ij}^2$$
 vs.  $H_1^{ij}: b_{0ij}^1 \neq b_{0ij}^2$ 

for  $j \in \mathcal{I}_q$ . Here we use the test statistic

$$d_{ij} = \frac{\hat{c}_{ij}^1 - \hat{c}_{ij}^2}{\sqrt{\tau_{ij}^1/(m_i^1)^2 + \tau_{ij}^2/(m_i^2)^2}}$$
(3.4)

with  $\tau_{ij}^k$  being the  $(i,j)^{\text{th}}$  element of  $(\widehat{\Omega}_y^k)^{-1}$ , for k=1,2.

Now consider tests where  $H_0^{ij}$  is rejected if  $|d_{ij}| > \tau$ . We denote  $\mathcal{H}_0^i = \{j: b_{ij}^1 = b_{ij}^2\}$ and define the false discovery proportion (FDP) and false discovery rate (FDR) for these tests as follows:

$$FDP(\tau) = \frac{\sum_{j \in \mathcal{H}_0^i} \mathbb{I}(|d_{ij}| \ge \tau)}{\max \left\{ \sum_{j \in \mathcal{I}_q} \mathbb{I}(|d_{ij}| \ge \tau), 1 \right\}} \quad FDR(\tau) = \mathbb{E}[FDP(\tau)]$$

For a pre-specified level  $\alpha$ , we choose a threshold that ensures both FDP and FDR  $< \alpha$ using the Benjamini-Hochberg (BH) procedure. The procedure for FDR control is now given by Algorithm 2.

**Algorithm 2.** (Simultaneous tests for  $H_0^{ij}: b_{0ij}^1 = b_{0ij}^2$  at level  $\alpha, 0 < \alpha < 1$ ) 1. Calculate the pairwise test statistics  $d_{ij}$  using (2) for  $j \in \mathcal{I}_q$ .

- 2. Obtain the threshold

$$\hat{\tau} = \inf \left\{ \tau \in \mathbb{R} : 1 - \Phi(\tau) \le \frac{\alpha}{2q} \max \left( \sum_{j \in \mathcal{I}_q} \mathbb{I}(|d_{ij}| \ge \tau), 1 \right) \right\}$$

3. For  $j \in \mathcal{I}_q$ , reject  $H_0^{ij}$  if  $|d_{ij}| \geq \hat{\tau}$ .

This procedure maintains FDR and FDP asymptotically at a pre-specified level  $\alpha \in$ (0,1) under weak dependence conditions.

**Theorem 3.2.** Suppose  $\mu_j = b_{0,ij}^1 - b_{0,ij}^2$ ,  $\sigma_j^2 = n\mathbb{E}(\tau_{ij}^1/(m_i^1)^2 + \tau_{ij}^2/(m_i^2)^2)$ . Assume the following holds as  $(n,q) \to \infty$ 

$$\left| \left\{ j \in \mathcal{I}_q : |\mu_j / \sigma_j| \ge 4\sqrt{\log q / n} \right\} \right| \to \infty \tag{3.5}$$

Now Consider the conditions C1, C1\*

If (C1) is satisfied, then the following holds when  $\log q = O(n^{\xi}), 0 < \xi < 3/23$ :

$$\frac{FDP(\hat{\tau})}{(|\mathcal{H}_0^i|/q)\alpha} \xrightarrow{P} 1; \quad \lim_{n,q \to \infty} \frac{FDR(\hat{\tau})}{(|\mathcal{H}_0^i|/q)\alpha} = 1 \tag{3.6}$$

Further, if  $(C1^*)$  is satisfied, then (3.6) holds for  $\log q = o(n^{1/3})$ .

The condition (3.5) is essential for FDR control in a diverging parameter space (Liu, 2017; Liu and Shao, 2014), while the dependence conditions (C1) and (C1\*) control the amount of correlation between variables in the Y-layer.

Remark 1. Following Liu and Shao (2014), a version of Algorithm 2, where the null distribution is calibrated using bootstrap instead of normal approximation, gives asymptotic FDR control under (C1\*) and  $\log q = o(n^{1/2})$ . We believe it is possible to obtain (3.6) under the weaker condition (C1) for  $\log q = o(n^{1/2})$  by extending the framework of Liu (2017) that performs multiple testing in multiple (single layer) GGMs, with the added advantage of being generalizable to the case of K > 2. However, this requires a significant amount of theoretical analysis, and we leave it for future research.

#### Remark 2. does within-group thresholding with FDR control for K=1

### A Proofs

Proof of Theorem 2.1. The proof has three parts, where we prove the consistency of the neighborhood regression coefficients, selection of edge sets, and finally the refitting step, respectively. This is the same structure as the proof of Theorem 1 in Ma and Michailidis (2016), where they prove consistency of the (single layer) JSEM estimates. The derivation of the first part is different from that in the JSEM proof, which we shall show in detail (in the proof of Proposition A.1). The second and third parts follow similar lines, incorporating the updated quantities from the first part. For these we provide outlines and leave the details to the reader.

Step 1: consistency of neighborhood regression. The following proposition establishes error bounds for estimated neighborhood coefficients in the Y-network.

**Proposition A.1.** Consider the estimation problem in (2.9) and take  $\gamma_n \geq 4\sqrt{|g_{\text{max}}|}\mathbb{Q}_0$ . Given the conditions (T2) and (T3) hold, for any solution of (2.9) we shall have

$$\|\widehat{\Theta}_j - \Theta_{0,j}\|_F \le 12\sqrt{s_j}\gamma_n/\psi \tag{A.1}$$

$$\sum_{j \neq j', g \in \mathcal{G}_y^{jj'}} \|\hat{\boldsymbol{\theta}}_{jj'}^{[g]} - \boldsymbol{\theta}_{0,jj'}^{[g]}\| \le 48s_j \gamma_n / \psi \tag{A.2}$$

Also denote the non-zero support of  $\widehat{\Theta}_j$  by  $\widehat{\mathcal{S}}_j$ , i.e.  $\widehat{\mathcal{S}}_j = \{(j',g) : \widehat{\boldsymbol{\theta}}_{jj'}^{[g]} \neq \mathbf{0}\}$ . Then

$$|\widehat{\mathcal{S}}_i| \le 128s_i/\psi \tag{A.3}$$

Step 2: Edge set selection. We denote the selected edge set for the  $k^{\text{th}}$  Y-network by  $\hat{E}^k$ . Denote its population version by  $E_0^k$ . Further, let

$$\tilde{\Omega}_y^k = \operatorname{diag}(\Omega_y^k) + \Omega_{y, E_0^k \cap \hat{E}^k}^k$$

With similar derivations to the proof of Corollary A.1 in Ma and Michailidis (2016), The following two upper bounds can be established:

$$|\hat{E}^k| \le \frac{128S}{\psi} \tag{A.4}$$

$$\frac{1}{K} \sum_{k=1}^{K} \|\tilde{\Omega}_{y}^{k} - \Omega_{y}^{k}\|_{F} \le \frac{12c_{0}\sqrt{S}\gamma_{n}}{\sqrt{K}\psi}$$
(A.5)

following which, taking  $\gamma_n = 4\sqrt{|g_{\text{max}}|}\mathbb{Q}_0$ ,

$$\Lambda_{\min}(\tilde{\Omega}_y^k) \ge d_0 - \frac{48c_0 \mathbb{Q}_0 \sqrt{|g_{\max}|S}}{\psi} \ge (1 - t_1)d_0 > 0 \tag{A.6}$$

$$\Lambda_{\max}(\tilde{\Omega}_y^k) \le c_0 + \frac{48c_0 \mathbb{Q}_0 \sqrt{|g_{\max}|S}}{\psi} \le c_0 + t_1 d_0 < \infty \tag{A.7}$$

with  $0 < t_1 < 1$ , and the sample size n satisfying

$$n \ge |g_{\text{max}}| S \left[ \frac{48c_0 Q_0}{\psi t_1 d_0} \right]^2; \quad Q_0 := \sqrt{n} \mathbb{Q}_0$$

Step 3: Refitting. Following the same steps as part A.3 in the proof of Theorem 4.1 in Ma and Michailidis (2016), it can be proven using (A.4)–(A.7) that

$$\sum_{k=1}^{K} \|\widehat{\Omega}_{y}^{k} - \widetilde{\Omega}_{y}^{k}\|^{2} \le O\left(\mathbb{Q}_{0}^{2} |g_{\max}|S\right)$$

The proof is now complete by combining this with (A.5) then applying Cauchy-Schwarz inequality and triangle inequality.

Proof of Proposition 2.2. We drop the superscript k since there is no scope of ambiguity. For part 1, we start with an auxiliary lemma:

**Lemma A.2.** For a sub-gaussian design matrix  $\mathbf{X} \in \mathbb{M}(n,p)$  with columns having mean  $\mathbf{0}_p$  and covariance matrix  $\Sigma_x$ , the sample covariance matrix  $\widehat{\Sigma}_x = \mathbf{X}^T \mathbf{X}/n$  satisfies the RE condition

$$\widehat{\Sigma}_x \sim RE\left(\frac{\Lambda_{\min}(\Sigma_x)}{2}, \frac{\Lambda_{\min}(\Sigma_x)\log p}{2n}\right)$$

with probability  $\geq 1 - 2 \exp(-c_3 n)$  for some  $c_3 > 0$ .

Now denote  $\hat{\mathbf{E}} = \mathbf{Y} - \mathbf{X}\hat{\mathbf{B}}$ . For  $\mathbf{v} \in \mathbb{R}^q$ , we have

$$\mathbf{v}^{T} \widehat{\mathbf{S}} \mathbf{v} = \frac{1}{n} \|\widehat{\mathbf{E}} \mathbf{v}\|^{2}$$

$$= \frac{1}{n} \|(\mathbf{E} + \mathbf{X}(\mathbf{B}_{0} - \widehat{\mathbf{B}})) \mathbf{v}\|^{2}$$

$$= \mathbf{v}^{T} \mathbf{S} \mathbf{v} + \frac{1}{n} \|\mathbf{X}(\mathbf{B}_{0} - \widehat{\mathbf{B}}) \mathbf{v}\|^{2} + 2 \mathbf{v}^{T} (\mathbf{B}_{0} - \widehat{\mathbf{B}})^{T} \left(\frac{(\mathbf{X})^{T} \mathbf{E}}{n}\right) \mathbf{v}$$
(A.8)

For the first summand,  $\mathbf{v}^T \mathbf{S}^k \mathbf{v} \ge \psi_y \|\mathbf{v}\|^2 - \phi_y \|\mathbf{v}\|_1^2$  with  $\psi_y = \Lambda_{\min}(\Sigma_y)/2$ ,  $\phi_y = \psi_y \log p/n$  by applying Lemma A.2 on S. The second summand is greater than or equal to 0. For the third summand,

$$2\mathbf{v}^{T}(\mathbf{B}_{0} - \widehat{\mathbf{B}})^{T} \left(\frac{(\mathbf{X})^{T} \mathbf{E}}{n}\right) \mathbf{v} \geq -2v_{\beta} \left\|\frac{(\mathbf{X})^{T} \mathbf{E}}{n}\right\|_{\infty} \|\mathbf{v}\|_{1}^{2}$$

by assumption (T1). Now we use another lemma:

**Lemma A.3.** For zero-mean independent sub-gaussian matrices  $\mathbf{X} \in \mathbb{M}(n,p)$ ,  $\mathbf{E} \in \mathbb{M}(n,q)$  with parameters  $(\Sigma_x, \sigma_x^2)$  and  $(\Sigma_e, \sigma_e^2)$  respectively, given that  $n \succeq \log(pq)$  the following holds with probability  $\geq 1 - 6c_1 \exp[-(c_2^2 - 1)\log(pq)]$  for some  $c_1 > 0, c_2 > 1$ :

$$\frac{1}{n} \|\mathbf{X}^T \mathbf{E}\|_{\infty} \le c_2 [\Lambda_{\max}(\Sigma_x) \Lambda_{\max}(\Sigma_e)]^{1/2} \sqrt{\frac{\log(pq)}{n}}$$

Subsequently we collect all summands in (A.8) and get

$$\mathbf{v}^T \widehat{\mathbf{S}} \mathbf{v} \ge \psi_y \|\mathbf{v}\|^2 - \left(\phi_y + 2v_\beta c_2 [\Lambda_{\max}(\Sigma_x) \Lambda_{\max}(\Sigma_y)]^{1/2} \sqrt{\frac{\log(pq)}{n}}\right) \|\mathbf{v}\|_1^2$$

with probability  $\geq 1 - 2\exp(-c_3n) - 6c_1\exp[-(c_2^2 - 1)\log(pq)]$ . This concludes the proof of part 1.

To prove part 2, we decompose the quantity in question:

$$\left\| \frac{1}{n} \widehat{\mathbf{E}}_{-j}^{T} \widehat{\mathbf{E}} \mathbf{T}_{0,j} \right\|_{\infty} = \left\| \frac{1}{n} \left[ \mathbf{E}_{-j} + \mathbf{X} (\mathbf{B}_{0,j} - \widehat{\mathbf{B}}_{j}) \right]^{T} \left[ \mathbf{E} + \mathbf{X} (\mathbf{B}_{0} - \widehat{\mathbf{B}}) \right] \mathbf{T}_{0,j} \right\|_{\infty} 
\leq \left\| \frac{1}{n} \mathbf{E}_{-j}^{T} \mathbf{E} \mathbf{T}_{0,j} \right\|_{\infty} + \left\| \frac{1}{n} \mathbf{E}_{-j}^{T} \mathbf{X} (\mathbf{B}_{0} - \widehat{\mathbf{B}}) \mathbf{T}_{0,j} \right\|_{\infty} 
+ \left\| \frac{1}{n} (\mathbf{B}_{0,j} - \widehat{\mathbf{B}}_{j})^{T} \mathbf{X}^{T} \mathbf{X} (\mathbf{B}_{0} - \widehat{\mathbf{B}}) \mathbf{T}_{0,j} \right\|_{\infty} + \left\| \frac{1}{n} (\mathbf{B}_{0,j} - \widehat{\mathbf{B}}_{j})^{T} \mathbf{X}^{T} \mathbf{E} \mathbf{T}_{0,j} \right\|_{\infty} 
= \| \mathbf{W}_{1} \|_{\infty} + \| \mathbf{W}_{2} \|_{\infty} + \| \mathbf{W}_{3} \|_{\infty} + \| \mathbf{W}_{4} \|_{\infty}$$
(A.9)

Now

$$\mathbf{W}_1 = \frac{1}{n} \mathbf{E}_{-j}^T (\mathbf{E}_j - \mathbf{E}_{-j} \boldsymbol{\theta}_{0,j})$$

For node j in the y-network,  $\mathbb{E}_{-j}$  and  $E_j - \mathbb{E}_{-j}\boldsymbol{\theta}_{0,j}$  are the neighborhood regression coefficients and residuals, respectively. Thus they are orthogonal, so we can apply Lemma A.3 on  $\mathbf{E}_{-j}$  and  $\mathbf{E}_j - \mathbf{E}_{-j}\boldsymbol{\theta}_{0,j}$  to obtain that for  $n \succeq \log(q-1)$ ,

$$\|\mathbf{W}_1\|_{\infty} \le c_5 \left[\Lambda_{\max}(\Sigma_{y,-j})\sigma_{y,j,-j}\right]^{1/2} \sqrt{\frac{\log(q-1)}{n}}$$
 (A.10)

holds with probability  $\geq 1 - 6c_4 \exp[-(c_5^2 - 1)\log(pq)]$  for some  $c_4 > 0, c_5 > 1$ .

The same bounds hold for  $W_2$  and  $W_4$ :

$$\|\mathbf{W}_2\|_{\infty} \leq \left\| \frac{1}{n} \mathbf{E}_{-j}^T \mathbf{X} (\mathbf{B}_0 - \widehat{\mathbf{B}}) \right\|_{\infty} \|\mathbf{T}_{0,j}\|_1 \leq \left\| \frac{1}{n} \mathbf{E}^T \mathbf{X} \right\|_{\infty} \|\mathbf{B}_0 - \widehat{\mathbf{B}}\|_1 \|\mathbf{T}_{0,j}\|_1$$
$$\|\mathbf{W}_4\|_{\infty} \leq \left\| \frac{1}{n} (\mathbf{B}_{0,j} - \widehat{\mathbf{B}}_j)^T \mathbf{X}^T \mathbf{E} \right\|_{\infty} \|\mathbf{T}_{0,j}\|_1 \leq \left\| \frac{1}{n} \mathbf{E}^T \mathbf{X} \right\|_{\infty} \|\mathbf{B}_0 - \widehat{\mathbf{B}}\|_1 \|\mathbf{T}_{0,j}\|_1$$

Now since  $\Omega_y$  is diagonally dominant,  $|\omega_{y,jj}| \geq \sum_{j \neq j'} |\omega_{y,jj'}|$  for any  $j \in \mathcal{I}_q$ . Hence

$$\|\mathbf{T}_{0,j}\|_1 = \sum_{j'=1}^q |T_{jj'}| = 1 + \sum_{j \neq j'} |\theta_{jj'}| = 1 + \frac{1}{\omega_{y,jj}} \sum_{j \neq j'} |\omega_{y,jj'}| \le 2$$

so that for  $n \gtrsim \log(pq)$ ,

$$\|\mathbf{W}_2\|_{\infty} + \|\mathbf{W}_4\|_{\infty} \le 4v_{\beta}c_2[\Lambda_{\max}(\Sigma_x)\Lambda_{\max}(\Sigma_y)]^{1/2}\sqrt{\frac{\log(pq)}{n}}$$
(A.11)

with probability  $\geq 1 - 6c_1 \exp[-(c_2^2 - 1) \log(pq)]$  by applying Lemma A.3 and assumption (T1).

Finally for  $W_3$ , we apply Lemma 8 of Ravikumar et al. (2011) on the (sub-gaussian) design matrix X to obtain that for sample size

$$n \ge 512(1 + 4\Lambda_{\max}(\Sigma_x^k))^4 \max_i (\sigma_{x,ii}^k)^4 \log(4p^{\tau_1})$$
(A.12)

we get that with probability  $\geq 1 - 1/p^{\tau_1 - 2}, \tau_1 > 2$ 

$$\left\| \frac{\mathbf{X}^T \mathbf{X}}{n} \right\|_{\infty} \leq \sqrt{\frac{\log 4 + \tau_1 \log p}{c_x n}} + \max_i \sigma_{x,ii} = V_x; \quad c_x = \left[ 128(1 + 4\Lambda_{\max}(\Sigma_x))^2 \max_i (\sigma_{x,ii})^2 \right]^{-1}$$

Thus with the same probability,

$$\|\mathbf{W}_4\|_{\infty} \le \left\| \frac{\mathbf{X}^T \mathbf{X}}{n} \right\|_{\infty} \|\widehat{\mathbf{B}} - \mathbf{B}_0\|_1^2 \|\mathbf{T}_{0,j}\|_1 \le 2v_{\beta}^2 V_x$$
 (A.13)

We now bound the right hand side of (A.9) using (A.10), (A.11) and (A.13) to complete the proof, with the leading term of the sample size requirement being  $n \gtrsim \log(pq)$ .

*Proof of Theorem 2.4.* The proof follows that of Proposition A.1, with a different group norm structure. We only point out the differences.

Putting  $\beta = \beta_0$  in (2.8) we get

$$-2\widehat{\boldsymbol{\beta}}^T\widehat{\boldsymbol{\gamma}} + \boldsymbol{\beta}^T\widehat{\boldsymbol{\Gamma}}\widehat{\boldsymbol{\beta}} + \lambda_n \sum_{h \in \mathcal{H}} \|\widehat{\boldsymbol{\beta}}^{[h]}\| \leq -2\boldsymbol{\beta}_0^T\widehat{\boldsymbol{\gamma}} + \boldsymbol{\beta}_0^T\widehat{\boldsymbol{\Gamma}}\boldsymbol{\beta}_0 + \lambda_n \sum_{h \in \mathcal{H}} \|\boldsymbol{\beta}_0^{[h]}\|$$

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Denote  $\mathbf{b} = \hat{\boldsymbol{\beta}} - \boldsymbol{\beta}_0$ . Then we have

$$\mathbf{b}^T \widehat{\mathbf{\Gamma}} \mathbf{b} \leq 2 \mathbf{b}^T (\widehat{\boldsymbol{\gamma}} - \widehat{\mathbf{\Gamma}} \boldsymbol{\beta}_0) + \lambda_n \sum_{h \in \mathcal{H}} (\|\boldsymbol{\beta}_0^{[h]}\| - \|\boldsymbol{\beta}_0^{[h]} + \mathbf{b}^{[h]}\|)$$

Proceeding similarly as the proof of Proposition A.1, with a different deviation bound and choice of  $\lambda_n$ , we get expressions equivalent to (B.3) and (B.4) respectively:

$$\mathbf{b}^T \widehat{\mathbf{\Gamma}} \mathbf{b} \le \frac{3}{2} \sum_{h \in \mathcal{H}} \|\mathbf{b}^{[h]}\| \tag{A.14}$$

$$\frac{\psi^*}{3} \|\mathbf{b}\|^2 \le \lambda_n \sum_{h \in \mathcal{H}} \|\mathbf{b}^{[h]}\| \le 4\lambda_n \sqrt{s_\beta} \|\mathbf{b}\|$$
(A.15)

Furthermore,  $\|\mathbf{b}\|_1 \leq \sqrt{|h_{\max}|} \sum_{h \in \mathcal{H}} \|\mathbf{b}^{[h]}\|$ . The bounds in (2.11), (2.12), (2.13) and (2.14) now follow.

Proof of Proposition 2.5. For part 1 it is enough to prove that with  $\widehat{\Sigma}_x^k := (\mathbf{X}^k)^T \mathbf{X}^k / n$ ,

$$\widehat{\mathbf{T}}_k^2 \otimes \widehat{\Sigma}_x^k \sim RE(\psi_*^k, \phi_*^k) \tag{A.16}$$

with high enough probability. because then we can take  $\psi_* = \min_k \psi_*^k$ ,  $\phi_* = \max_k \phi_*^k$ . The proof of (A.16) follows similar lines of the proof of Proposition 1 in Lin et al. (2016), only replacing  $\Theta_{\epsilon}$ ,  $\widehat{\Theta}_{\epsilon}$ ,  $\mathbf{X}$  therein with  $(\mathbf{T}^k)^2$ ,  $(\widehat{\mathbf{T}}^k)^2$ ,  $\mathbf{X}^k$ , respectively. We omit the details. Part 2 follows the proof of Proposition 2 in Lin et al. (2016).

*Proof of Theorem 3.1.* Let us define the following:

$$\begin{split} \widehat{\Omega}_y &= \operatorname{diag}(\widehat{\Omega}_y^1, \widehat{\Omega}_y^2) \\ \mathbf{M}_i &= \operatorname{diag}(m_i^1, m_i^2) \\ \widehat{\mathbf{C}}_i &= \operatorname{diag}(\widehat{\mathbf{c}}_i^1, \widehat{\mathbf{c}}_i^2) \\ \widehat{\mathbf{D}}_i &= \operatorname{diag}(\widehat{\mathbf{b}}_i^1, \widehat{\mathbf{b}}_i^2) \\ \mathbf{D}_i &= \operatorname{diag}(\mathbf{b}_{0,i}^1, \mathbf{b}_{0,i}^2) \\ \mathbf{R}_i^k &= \mathbf{X}_i^k - \mathbf{X}_{-i}^k \widehat{\boldsymbol{\zeta}}_i^k; k = 1, 2 \end{split}$$

Then from (3.2) we have

$$\mathbf{M}_{i}(\widehat{\mathbf{C}}_{i} - \widehat{\mathbf{D}}_{i})^{T} = \frac{1}{\sqrt{n}} \begin{bmatrix} \frac{1}{\widehat{s}_{i}^{1}} (\mathbf{R}_{i}^{1})^{T} \widehat{\mathbf{E}}^{1} \\ \frac{1}{\widehat{s}_{i}^{2}} (\mathbf{R}_{i}^{2})^{T} \widehat{\mathbf{E}}^{2} \end{bmatrix}$$
(A.17)

We now decompose  $\widehat{\mathbf{E}}^k$ :

$$\begin{split} \widehat{\mathbf{E}}^k &= \mathbf{Y}^k - \mathbf{X}^k \widehat{\mathbf{B}}^k \\ &= \mathbf{E}^k + \mathbf{X}^k (\mathbf{B}_0^k - \widehat{\mathbf{B}}^k) \\ &= \mathbf{E}^k + \mathbf{X}_i^k (\mathbf{b}_{0,i}^k - \widehat{\mathbf{b}}_i^k) + \mathbf{X}_{-i}^k (\mathbf{B}_{0-i}^k - \widehat{\mathbf{B}}_{-i}^k) \end{split}$$

Putting them back in (A.17) and using  $t^k = (\mathbf{R}^k)^T \mathbf{X}^k / n$ ,

$$\mathbf{M}_{i}(\widehat{\mathbf{C}}_{i} - \widehat{\mathbf{D}}_{i})^{T} = \frac{1}{\sqrt{n}} \begin{bmatrix} \frac{1}{\widehat{s}_{i}^{T}} (\mathbf{R}_{i}^{1})^{T} \mathbf{E}^{1} \\ \frac{1}{\widehat{s}_{i}^{T}} (\mathbf{R}_{i}^{2})^{T} \mathbf{E}^{2} \end{bmatrix} + \mathbf{M}_{i} (\mathbf{D}_{i} - \widehat{\mathbf{D}}_{i})^{T} + \frac{1}{\sqrt{n}} \begin{bmatrix} \frac{1}{\widehat{s}_{i}^{T}} (\mathbf{R}_{i}^{1})^{T} \mathbf{X}_{-i}^{1} (\mathbf{B}_{0,-i}^{1} - \widehat{\mathbf{B}}_{-i}^{1}) \\ \frac{1}{\widehat{s}_{i}^{T}} (\mathbf{R}_{i}^{1})^{T} \mathbf{X}_{-i}^{T} (\mathbf{B}_{0,-i}^{1} - \widehat{\mathbf{B}}_{-i}^{2}) \end{bmatrix}$$

$$\Rightarrow \widehat{\Omega}_{y}^{1/2} \mathbf{M}_{i} (\widehat{\mathbf{C}}_{i} - \mathbf{D}_{i})^{T} = \frac{\widehat{\Omega}_{y}^{1/2}}{\sqrt{n}} \begin{bmatrix} \frac{1}{\widehat{s}_{i}^{T}} (\mathbf{R}_{i}^{1})^{T} \mathbf{E}^{1} \\ \frac{1}{\widehat{s}_{i}^{T}} (\mathbf{R}_{i}^{1})^{T} \mathbf{X}_{-i}^{T} (\mathbf{B}_{0,-i}^{1} - \widehat{\mathbf{B}}_{-i}^{1}) \\ \frac{1}{\widehat{s}_{i}^{T}} (\mathbf{R}_{i}^{2})^{T} \mathbf{X}_{-i}^{T} (\mathbf{B}_{0,-i}^{1} - \widehat{\mathbf{B}}_{-i}^{2}) \end{bmatrix}$$

$$(A.18)$$

At this point, we drop k in the subscripts, and prove the following:

**Lemma A.4.** Given conditions (C1) and (C2), the following holds for sample size n such that  $n \gtrsim \log(pq)$  and  $\sigma_{x,i,-i} - n^{-1/4} - v_{\zeta}\sqrt{V_x} > 0$ :

$$\frac{1}{\sqrt{n}\widehat{s}_i}\widehat{\Omega}_y^{1/2}\mathbf{E}^T\mathbf{R}_i \sim \mathcal{N}_q(\mathbf{0}, \mathbf{I}) + \mathbf{S}_{1n};$$

$$\|\mathbf{S}_{1n}\|_{\infty} \le \frac{v_{\Omega}(2 + v_{\zeta})c_{2}[\Lambda_{\max}(\Sigma_{x})\Lambda_{\max}(\Sigma_{e})]^{1/2}\sqrt{\log(pq)}}{\sigma_{x,i,-i} - n^{-1/4} - v_{\zeta}\sqrt{V_{x}}} = O\left(\frac{\log(pq)}{\sqrt{n}}\right)$$
(A.19)

with probability larger than or equal to

$$1 - 6c_1 e^{-(c_2^2 - 1)\log pq} - \frac{1}{p^{\tau_1 - 2}} - \frac{\kappa_i}{\sqrt{n}}$$
(A.20)

for some  $c_1, c_4 > 0, c_2, c_5 > 1$ , and  $\kappa_i := \mathbb{V}[(X_i - \mathbb{X}_{-i} \boldsymbol{\zeta}_{0,-i})^2]$ . Additionally, given condition (C3)

$$\left\| \frac{1}{\sqrt{n}\widehat{s}_{i}} \mathbf{R}_{i}^{T} \mathbf{X}_{-i} (\mathbf{B}_{0,-i} - \widehat{\mathbf{B}}_{-i}) \widehat{\Omega}_{y}^{1/2} \right\|_{\infty} \\
\leq \frac{v_{\beta} (\Lambda_{\min}(\Sigma_{y})^{1/2} + v_{\Omega})}{\sigma_{x,i,-i} - n^{-1/2} - v_{\zeta} \sqrt{V_{x}}} \left[ c_{7} \sqrt{(\sigma_{x,i,-i}\Lambda_{\max}(\Sigma_{x,-i})) \log p} + \sqrt{n} v_{\zeta} V_{x} \right] = O\left(\frac{\log(pq)}{\sqrt{n}}\right) \tag{A.21}$$

with probability condition (A.20).

Given Lemma A.4, the first and second summands on the right hand side of (A.18) are bounded above by applying each of (A.19) and (A.21) twice, respectively. This completes our proof.

Proof of Theorem 
$$3.2$$
.

# B Proof of auxiliary results

Proof of Proposition A.1. In its reparametrized version, (2.9) becomes

$$\widehat{\mathbf{T}}_{j} = \operatorname*{arg\,min}_{\mathbf{T}_{j}} \left\{ \frac{1}{n} \sum_{k=1}^{K} \| (\mathbf{Y}^{k} - \mathbf{X}^{k} \widehat{\mathbf{B}}^{k}) \mathbf{T}_{j}^{k} \|^{2} + \gamma_{n} \sum_{j \neq j', g \in \mathcal{G}_{y}^{jj'}} \| \mathbf{T}_{jj'}^{[g]} \| \right\}$$
(B.1)

with  $\mathbf{T}_{jj'}^{[g]} := (T_{jj'}^k)_{k \in g}$ . Now for any  $\mathbf{T}_j \in \mathbb{M}(q,K)$  we have

$$\frac{1}{n} \sum_{k=1}^{K} \| (\mathbf{Y}^k - \mathbf{X}^k \widehat{\mathbf{B}}^k) \widehat{\mathbf{T}}_j^k \|^2 + \gamma_n \sum_{j \neq j', g \in \mathcal{G}_y^{jj'}} \| \widehat{\mathbf{T}}_{jj'}^{[g]} \| \leq \frac{1}{n} \sum_{k=1}^{K} \| (\mathbf{Y}^k - \mathbf{X}^k \widehat{\mathbf{B}}^k) \mathbf{T}_j^k \|^2 + \gamma_n \sum_{j \neq j', g \in \mathcal{G}_y^{jj'}} \| \mathbf{T}_{jj'}^{[g]} \|$$

For  $\mathbf{T}_j = \mathbf{T}_{0,j}$  this reduces to

$$\sum_{k=1}^{K} (\mathbf{d}_{j}^{k})^{T} \widehat{\mathbf{S}}^{k} \mathbf{d}_{j}^{k} \leq -2 \sum_{k=1}^{K} (\mathbf{d}_{j}^{k})^{T} \widehat{\mathbf{S}}^{k} \mathbf{T}_{0,j}^{k} + \gamma_{n} \sum_{j \neq j', g \in \mathcal{G}_{y}^{jj'}} \left( \|\mathbf{T}_{jj'}^{[g]}\| - \|\mathbf{T}_{jj'}^{[g]}\| + \mathbf{d}_{jj'}^{[g]}\| \right)$$
(B.2)

with  $\mathbf{d}_{j}^{k} := \widehat{\mathbf{T}}_{j}^{k} - \mathbf{T}_{0,j}^{k}$  etc. For the  $k^{\text{th}}$  summand in the first term on the right hand side, since  $d_{jj}^{k} = 0$ ,  $\widehat{\mathbf{E}}^{k} \mathbf{d}_{j}^{k} = \widehat{\mathbf{E}}_{-j}^{k} \mathbf{d}_{-j}^{k}$ . Thus

$$\begin{split} \sum_{k=1}^{K} \left| (\mathbf{d}_{j}^{k})^{T} \widehat{\mathbf{S}}^{k} \mathbf{T}_{0,j}^{k} \right| &= \sum_{k=1}^{K} \left| \mathbf{d}_{j}^{k} \cdot \frac{1}{n} (\widehat{\mathbf{E}}^{k})^{T} \widehat{\mathbf{E}}^{k} \mathbf{T}_{0,j}^{k} \right| \\ &\leq \sum_{k=1}^{K} \left\| \frac{1}{n} (\widehat{\mathbf{E}}_{-j}^{k})^{T} \widehat{\mathbf{E}}^{k} \mathbf{T}_{0,j}^{k} \right\|_{\infty} \|\mathbf{d}_{-j}^{k}\|_{1} \\ &\leq \mathbb{Q}_{0} \sqrt{|g_{\max}|} \sum_{j \neq j', g \in \mathcal{G}_{y}^{jj'}} \|\mathbf{d}_{jj'}^{[g]}\| \end{split}$$

by assumption (T2). For the second term, suppose  $S_{0,j}$  is the support of  $\Theta_{0,j}$ , i.e.  $S_{0,j} = \{(j',g) : \boldsymbol{\theta}_{jj'}^{[g]} \neq 0\}$ . Then

$$\begin{split} \sum_{j \neq j', g \in \mathcal{G}_{y}^{jj'}} \left( \| \mathbf{T}_{jj'}^{[g]} \| - \| \mathbf{T}_{jj'}^{[g]} + \mathbf{d}_{jj'}^{[g]} \| \right) &\leq \sum_{(j', g) \in \mathcal{S}_{0, j}} \left( \| \mathbf{T}_{jj'}^{[g]} \| - \| \mathbf{T}_{jj'}^{[g]} + \mathbf{d}_{jj'}^{[g]} \| \right) - \sum_{(j', g) \notin \mathcal{S}_{0, j}} \| \mathbf{d}_{jj'}^{[g]} \| \\ &\leq \sum_{(j', g) \in \mathcal{S}_{0, j}} \| \mathbf{d}_{jj'}^{[g]} \| - \sum_{(j', g) \notin \mathcal{S}_{0, j}} \| \mathbf{d}_{jj'}^{[g]} \| \end{split}$$

so that by choice of  $\gamma_n$  (B.2) reduces to

$$\sum_{k=1}^{K} (\mathbf{d}_{j}^{k})^{T} \widehat{\mathbf{S}}^{k} \mathbf{d}_{j}^{k} \leq \frac{\gamma_{n}}{2} \left[ \sum_{(j',g) \in \mathcal{S}_{0,j}} \|\mathbf{d}_{jj'}^{[g]}\| + \sum_{(j',g) \notin \mathcal{S}_{0,j}} \|\mathbf{d}_{jj'}^{[g]}\| \right] + \gamma_{n} \left[ \sum_{(j',g) \in \mathcal{S}_{0,j}} \|\mathbf{d}_{jj'}^{[g]}\| - \sum_{(j',g) \notin \mathcal{S}_{0,j}} \|\mathbf{d}_{jj'}^{[g]}\| \right] \\
= \frac{3\gamma_{n}}{2} \sum_{(j',g) \in \mathcal{S}_{0,j}} \|\mathbf{d}_{jj'}^{[g]}\| - \frac{\gamma_{n}}{2} \sum_{(j',g) \notin \mathcal{S}_{0,j}} \|\mathbf{d}_{jj'}^{[g]}\| \\
\leq \frac{3\gamma_{n}}{2} \sum_{j \neq j', g \in \mathcal{G}_{y}^{jj'}} \|\mathbf{d}_{jj'}^{[g]}\| \tag{B.3}$$

Since the left hand side is  $\geq 0$ , this also implies

$$\sum_{(j',g)\notin\mathcal{S}_{0,j}} \|\mathbf{d}_{jj'}^{[g]}\| \leq 3 \sum_{(j',g)\in\mathcal{S}_{0,j}} \|\mathbf{d}_{jj'}^{[g]}\| \quad \Rightarrow \sum_{j\neq j',g\in\mathcal{G}_y^{jj'}} \|\mathbf{d}_{jj'}^{[g]}\| \leq 4 \sum_{(j',g)\in\mathcal{S}_{0,j}} \|\mathbf{d}_{jj'}^{[g]}\| \leq 4\sqrt{s_j} \|\mathbf{D}_j\|_F$$

with  $\mathbf{D}_j = (\mathbf{d}_j^k)_{k \in \mathcal{I}_K}$ . Now the RE condition on  $\widehat{\mathbf{S}}^k$  means that

$$\sum_{k=1}^{K} (\mathbf{d}_{j}^{k})^{T} \widehat{\mathbf{S}}^{k} \mathbf{d}_{j}^{k} \geq \sum_{k=1}^{K} \left( \psi_{k} \|\mathbf{d}_{j}^{k}\|^{2} - \phi_{k} \|\mathbf{d}_{j}^{k}\|_{1}^{2} \right) \geq \psi \|\mathbf{D}_{j}\|_{F}^{2} - \phi \|\mathbf{D}_{j}\|_{1}^{2} \geq (\psi - Kq\phi) \|\mathbf{D}_{j}\|_{F}^{2} \geq \frac{\psi}{2} \|\mathbf{D}_{j}\|_{F}^{2}$$

by assumption (T3).

Thus we finally have

$$\frac{\psi}{3} \|\mathbf{D}_j\|_F^2 \le \gamma_n \sum_{j \ne j', g \in \mathcal{G}_y^{jj'}} \|\mathbf{d}_{jj'}^{[g]}\| \le 4\gamma_n \sqrt{s_j} \|\mathbf{D}_j\|_F \tag{B.4}$$

Since

$$(\mathbf{D}_{j})_{j',k} = \widehat{T}_{jj'}^{k} - T_{0,jj'}^{k} = \begin{cases} 0 \text{ if } j = j' \\ -(\widehat{\theta}_{jj'}^{k} - \theta_{0,jj'}^{k}) \text{ if } j \neq j' \end{cases}$$

The bounds in (A.1) and (A.2) are obtained by replacing the corresponding elements in (B.4).

For the bound on  $|\widehat{\mathcal{S}}_j|$ , notice that if  $\hat{\theta}_{jj'}^{[g]} \neq 0$  for some (j', g),

$$\begin{split} \frac{1}{n} \sum_{k \in g} \left| ((\widehat{\mathbf{E}}_{-j}^k)^T \widehat{\mathbf{E}}^k (\widehat{\mathbf{T}}_j^k - \mathbf{T}_{0,j}^k))^{j'} \right| &\geq \frac{1}{n} \sum_{k \in g} \left| ((\widehat{\mathbf{E}}_{-j}^k)^T \widehat{\mathbf{E}}^k \widehat{\mathbf{T}}_j^k)^{j'} \right| - \frac{1}{n} \sum_{k \in g} \left| ((\widehat{\mathbf{E}}_{-j}^k)^T \widehat{\mathbf{E}}^k \mathbf{T}_{0,j}^k)^{j'} \right| \\ &\geq |g| \gamma_n - \sum_{k \in g} \mathbb{Q}(v_\beta, \Sigma_x^k, \Sigma_y^k) \end{split}$$

using the KKT condition for (2.9) and assumption (T2). The choice of  $\gamma_n$  now ensures

that the right hand side is  $\geq 3|g|\gamma_n/4$ . Hence

$$\begin{aligned} |\hat{\mathcal{S}}_{j}| &\leq \sum_{(j',g) \in \hat{\mathcal{S}}_{j}} \frac{16}{9n^{2}|g|^{2}\gamma_{n}^{2}} \sum_{k \in g} \left| ((\widehat{\mathbf{E}}_{-j}^{k})^{T} \widehat{\mathbf{E}}^{k} (\widehat{\mathbf{T}}_{j}^{k} - \mathbf{T}_{0,j}^{k}))^{j'} \right|^{2} \\ &\leq \frac{16}{9\gamma_{n}^{2}} \sum_{k=1}^{K} \frac{1}{n} \left\| (\widehat{\mathbf{E}}_{-j}^{k})^{T} \widehat{\mathbf{E}}^{k} (\widehat{\mathbf{T}}_{j}^{k} - \mathbf{T}_{0,j}^{k}) \right\|^{2} \\ &= \frac{16}{9\gamma_{n}^{2}} \sum_{k=1}^{K} (\mathbf{d}_{j}^{k})^{T} \widehat{\mathbf{S}}^{k} \mathbf{d}_{j}^{k} \\ &\leq \frac{8}{3\gamma_{n}} \sum_{j \neq j', g \in \mathcal{G}_{y}^{jj'}} \|\mathbf{d}_{jj'}^{[g]}\| \leq \frac{128s_{j}}{\psi} \end{aligned}$$

using (B.3) and (B.4).

*Proof of Lemma A.2.* This is same as Lemma 2 in Appendix B of Lin et al. (2016) and its proof can be found there.  $\Box$ 

*Proof of Lemma A.3.* This is a part of Lemma 3 of Appendix B in Lin et al. (2016), and is proved therein.  $\Box$ 

Proof of Lemma A.4. To show (A.19) we have

$$\frac{1}{\sqrt{n}\widehat{s}_i}\widehat{\Omega}_y^{1/2}\mathbf{E}^T\mathbf{R}_i = \frac{1}{\sqrt{n}\widehat{s}_i}(\widehat{\Omega}_y^{1/2} - \Omega_y^{1/2})\mathbf{E}^T\mathbf{R}_i + \frac{1}{\sqrt{n}\widehat{s}_i}\Omega_y^{1/2}\mathbf{E}^T\mathbf{R}_i$$

The second summand is distributed as  $\mathcal{N}_q(\mathbf{0}, \mathbf{I})$ . For the first summand,

$$\begin{split} \frac{1}{\sqrt{n}} \left\| (\widehat{\Omega}_{y}^{1/2} - \Omega_{y}^{1/2}) \mathbf{E}^{T} \mathbf{R}_{i} \right\|_{\infty} &\leq \frac{1}{\sqrt{n}} \left\| \widehat{\Omega}_{y}^{1/2} - \Omega_{y}^{1/2} \right\|_{\infty} \left\| \mathbf{E}^{T} \mathbf{R}_{i} \right\|_{1} \\ &\leq \sqrt{n} v_{\Omega} \frac{1}{n} \left[ \left\| \mathbf{E}^{T} (\mathbf{X}_{i} - \mathbf{X}_{-i} \boldsymbol{\zeta}_{i}) \right\|_{1} + \left\| \mathbf{E}^{T} \mathbf{X}_{-i} (\widehat{\boldsymbol{\zeta}}_{i} - \boldsymbol{\zeta}_{0,i}) \right\|_{1} \right] \\ &\leq \sqrt{n} v_{\Omega} \frac{1}{n} \left[ \left\| \mathbf{E}^{T} \mathbf{X}_{i} \right\|_{\infty} + \left\| \mathbf{E}^{T} \mathbf{X}_{-i} \right\|_{\infty} \left\{ \left\| \boldsymbol{\zeta}_{i} \right\|_{1} + \left\| \widehat{\boldsymbol{\zeta}}_{i} - \boldsymbol{\zeta}_{i} \right\|_{1} \right\} \right] \\ &\leq \sqrt{n} v_{\Omega} \left[ \frac{1}{n} \left\| \mathbf{E}^{T} \mathbf{X}_{i} \right\|_{\infty} + \frac{1 + v_{\zeta}}{n} \left\| \mathbf{E}^{T} \mathbf{X}_{-i} \right\|_{\infty} \right] \\ &\leq \sqrt{n} v_{\Omega} (2 + v_{\zeta}) \cdot \frac{1}{n} \left\| \mathbf{E}^{T} \mathbf{X} \right\|_{\infty} \end{split}$$

because  $\Omega_x$  is diagonally dominant implies  $\|\boldsymbol{\zeta}_i\|_1 = \sum_{i' \neq i} |\omega_{x,ii'}|/\omega_{x,ii} \leq 1$ , and using assumption (C1). Applying Lemma A.3, the following holds:

$$\frac{1}{\sqrt{n}} \left\| (\widehat{\Omega}_y^{1/2} - \Omega_y^{1/2}) \mathbf{E}^T \mathbf{R}_i \right\|_{\infty} \le v_{\Omega} (2 + v_{\zeta}) c_2 [\Lambda_{\max}(\Sigma_x) \Lambda_{\max}(\Sigma_e)]^{1/2} \sqrt{\log(pq)}$$
 (B.5)

with probability  $\geq 1 - 6c_1 \exp[-(c_2^2 - 1)\log(pq)]$  for some  $c_1 > 0, c_2 > 1$ .

On the other hand

$$s_i^2 := \frac{1}{n} \|\mathbf{X}_i - \mathbf{X}_{-i} \boldsymbol{\zeta}_i\|^2 \le \hat{s}_i^2 + \frac{1}{n} \|\mathbf{X}_{-i} (\hat{\boldsymbol{\zeta}}_i - \boldsymbol{\zeta}_{0,i})\|^2 \le \hat{s}_i^2 + \|\hat{\boldsymbol{\zeta}}_i - \boldsymbol{\zeta}_{0,i}\|_1^2 \|\frac{1}{n} \mathbf{X}_{-i}^T \mathbf{X}_{-i}\|_{\infty}$$

which implies  $s_i \leq \hat{s}_i + v_{\zeta} \sqrt{V_x}$ . By applying Lemma 8 of Ravikumar et al. (2011),

$$\left\| \frac{1}{n} \mathbf{X}_{-i}^T \mathbf{X}_{-i} \right\|_{\infty} \le \left\| \frac{1}{n} \mathbf{X}^T \mathbf{X} \right\|_{\infty} \le V_x \tag{B.6}$$

with probability  $\geq 1 - 1/p^{\tau_1-2}, \tau_1 > 2$ , and

$$n \ge 512(1 + 4\Lambda_{\max}(\Sigma_x))^4 \max_i (\sigma_{x,ii})^4 \log(4p^{\tau_1})$$
 (B.7)

On the other hand, by Chebyshev inequality, for any  $\epsilon > 0$ 

$$P(|s_i - \sigma_{x,i,-i}| \ge \epsilon) \le \frac{\mathbb{V}s_i}{\epsilon^2} = \frac{\kappa_i}{n\epsilon^2}$$

Taking  $\epsilon = n^{-1/4}$ , we have  $s_i \ge \sigma_{x,i,-i} - n^{-1/4}$  with probability  $\ge 1 - \kappa_i n^{-1/2}$ . Then, for n satisfying (B.6) and  $\sigma_{x,i,-i} - n^{-1/4} > v_\zeta \sqrt{V_x}$ , we get the bound with the above probability:

$$\frac{1}{\widehat{s}_i} \le \frac{1}{\sigma_{x,i,-i} - n^{-1/4} - v_\zeta \sqrt{V_x}} \tag{B.8}$$

Combining (B.5) and (B.8) gives the upper bound for the right hand side of (A.19) with the requisite probability and sample size conditions.

To prove (A.21) we have

$$\frac{1}{n} \|\mathbf{R}_{i}^{T} \mathbf{X}_{-i}\|_{\infty} \leq \frac{1}{n} \|(\mathbf{X}_{i} - \mathbf{X}_{-i} \boldsymbol{\zeta}_{0,i})^{T} \mathbf{X}_{-i}\|_{\infty} + \frac{1}{n} \|\mathbf{X}_{-i}^{T} \mathbf{X}_{-i} (\widehat{\boldsymbol{\zeta}}_{i} - \boldsymbol{\zeta}_{0,i})\|_{\infty}$$
(B.9)

Applying Lemma A.3, for  $n \geq \log(p-1)$  we have

$$\frac{1}{n} \| (\mathbf{X}_i - \mathbf{X}_{-i} \boldsymbol{\zeta}_i)^T \mathbf{X}_{-i} \|_{\infty} \le c_7 [\sigma_{x,i,-i} \Lambda_{\max}(\Sigma_{x,-i})]^{1/2} \sqrt{\frac{\log(p-1)}{n}}$$
(B.10)

with probability  $\geq 1 - 6c_6 \exp[-(c_7^2 - 1)\log(p - 1)]$  for some  $c_6 > 0, c_7 > 1$ . By (B.6), the second term on the right side of (B.9) is bounded above by  $v_\zeta V_x$  with probability  $\geq 1 - 1/p^{\tau_1 - 2}$  and n satisfying (B.7). The bound of (A.21) now follows by conditions (C2), (C3) and (B.8).

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