

Fast and Scalable Joint Estimators for Learning Sparse Gaussian Graphical Models from Heterogeneous Data with Additional Knowledge

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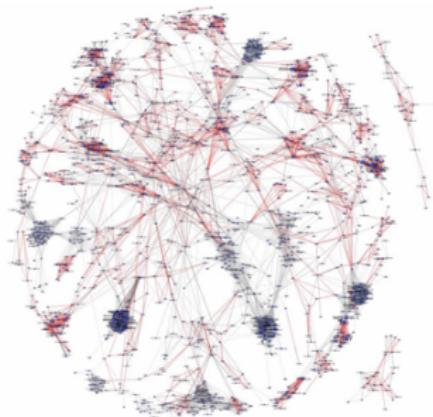
¹University of Virginia

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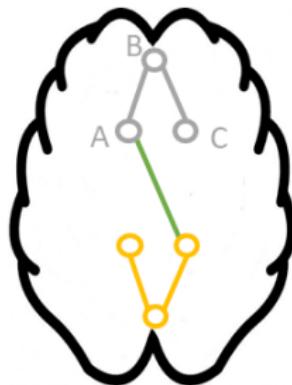
August 24, 2018

Background

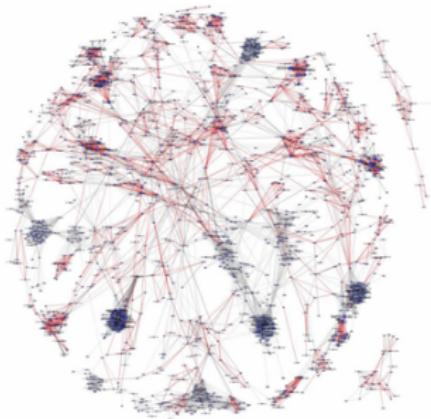
Background: Entity Graph



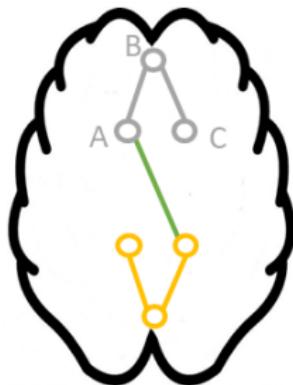
- Many applications need to know interactions among entities:
 - Gene Interactions
 - Brain connectivity



Background: Entity Graph



- Many applications need to know interactions among entities:
 - Gene Interactions
 - Brain connectivity
- Why to study the entity graph
 - Understanding
 - Diagnosis, e.g., marker
 - Treatment, e.g., drug development.



Background: What Type of Edges? Correlation to Conditional dependency

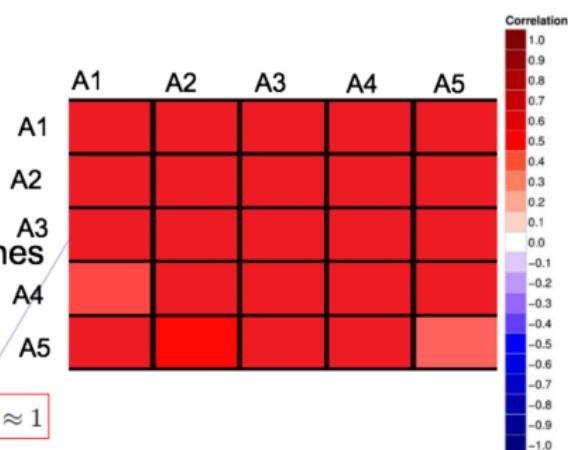
A1: Children swim

A2: Weather is hot

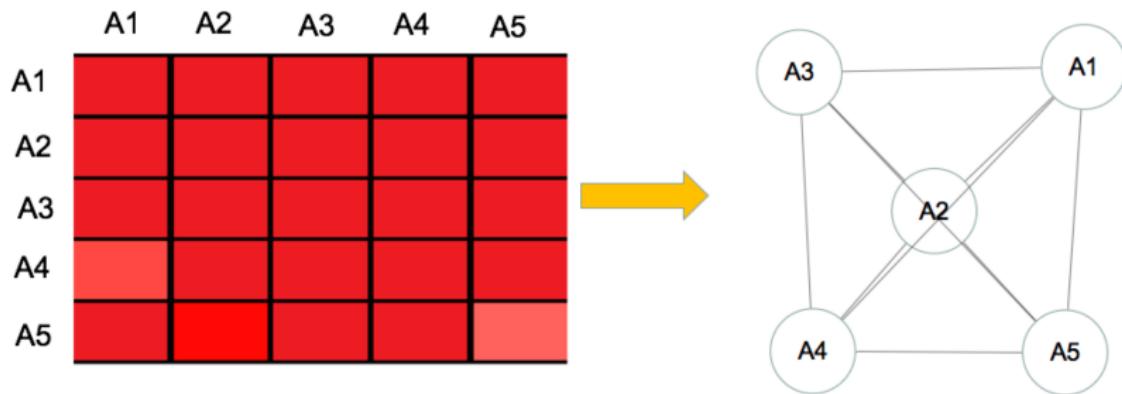
A3: High sale of ice cream

A4: Wear less amount of clothes

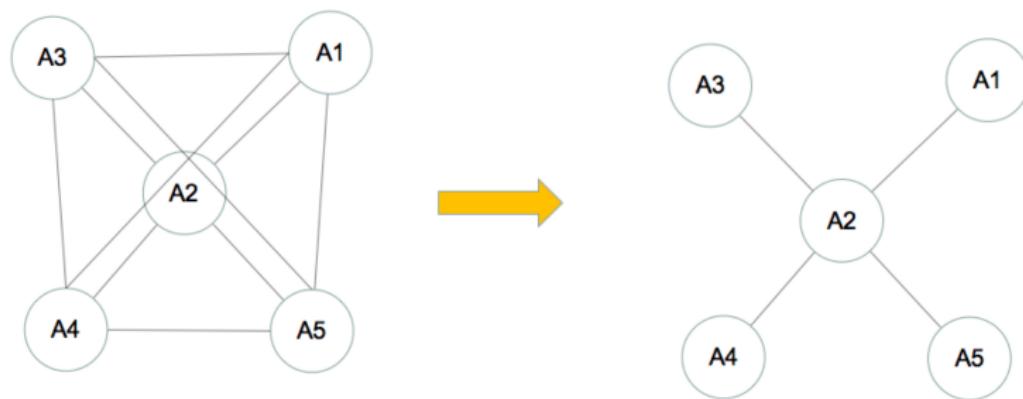
A5: High Electricity
Consumption



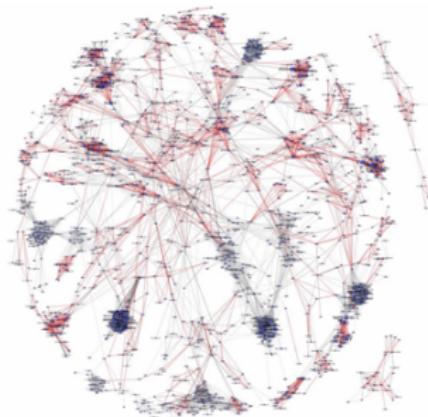
Background: What Type of Edges? Correlation to Conditional dependency



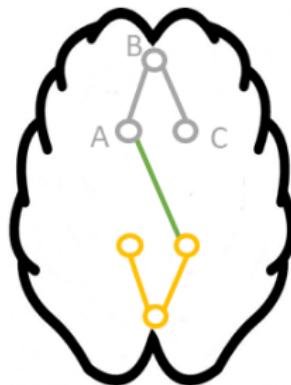
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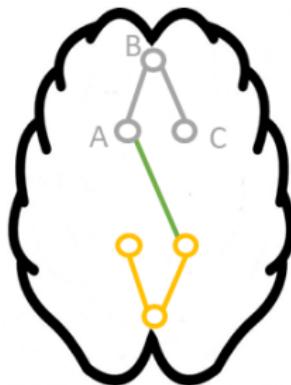
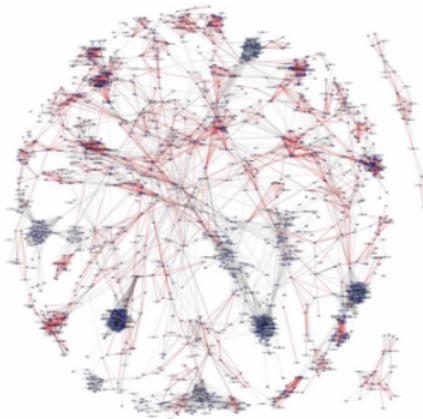
Background: How to Infer Entity Graph?



- To measure conditional dependency interactions physically.
- Largely unknown and hard to measure physically.

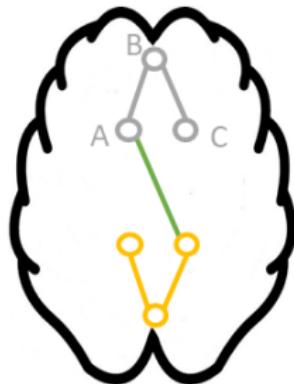
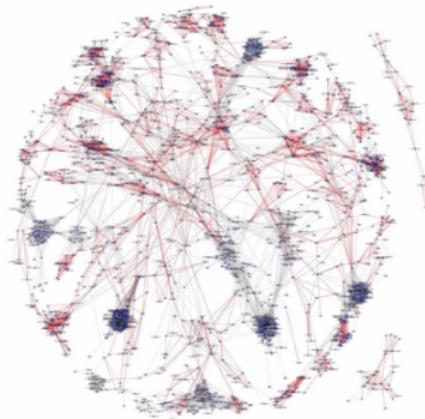


Background: How to Infer Entity Graph?



- To measure conditional dependency interactions physically.
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- #Physical check for all possible conditional dependency edges = 2^p (binary experiments)
 - For example, $p = 160$ important regions in human brain
 - For example, $p = 30000$ genes in human cell

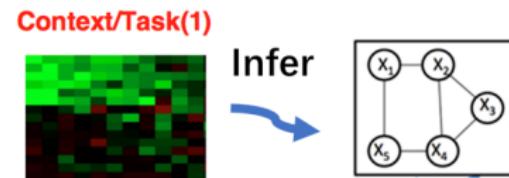
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- To measure conditional dependency interactions physically.
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-
- #Physical check for all possible conditional dependency edges = 2^p (binary experiments)
 - For example, $p = 160$ important regions in human brain
 - For example, $p = 30000$ genes in human cell
-
- Much more than Trillions (2^{40}) of biological experiments

Background: Entity graphs from Observed Samples (Entity as Feature)

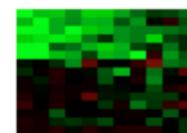
- Trillions of biological experiments \Rightarrow
Data-driven approach
- Experiments (not physically check)
 \Rightarrow Data \Rightarrow Entity Graph



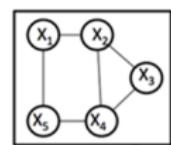
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- n experiments $\rightarrow n$ data samples
 - Each sample is a snapshot of all the entities.
 - Each sample has measurements of p features/entities.

Context/Task(1)

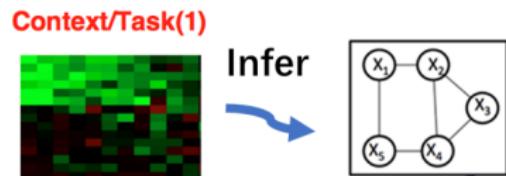


Infer



Background: Entity graphs from Observed Samples (Entity as Feature)

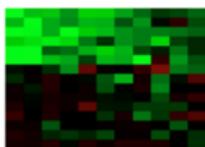
- Trillions of biological experiments \Rightarrow
Data-driven approach
- Experiments (not physically check)
 \Rightarrow Data \Rightarrow Entity Graph
- n experiments $\rightarrow n$ data samples
 - Each sample is a snapshot of all the entities.
 - Each sample has measurements of p features/entities.
- n data samples is enough \rightarrow a well estimated entity graph of p when $n \gg p$ (**low-dimensional**).
- $p > n$ (**high-dimensional**) needs novel approaches



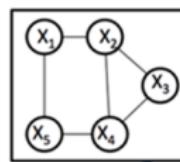
Background: Entity graphs from Heterogeneous Data (Entity as Feature)

- Most applications have heterogeneous samples.
- For example:
 - Totally n_{tot} data samples
 - From K different but related contexts, each has n_i data samples

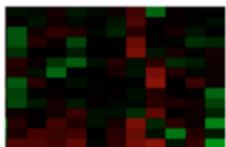
Context/Task(1)



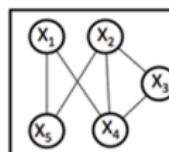
Infer



Context/Task(2)

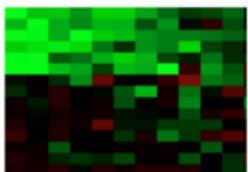


Machine learning approach

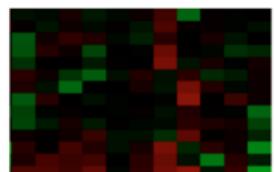


Background: Entity graphs from Heterogeneous Data

Context/Task(1)



Context/Task(2)



Case I:

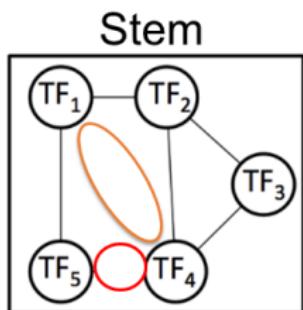
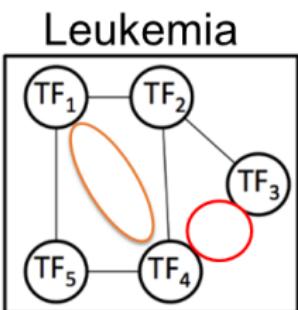
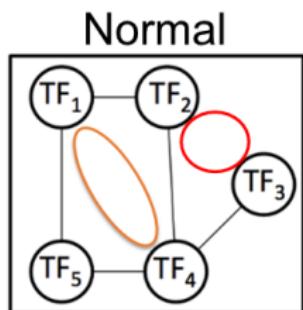


Case II:



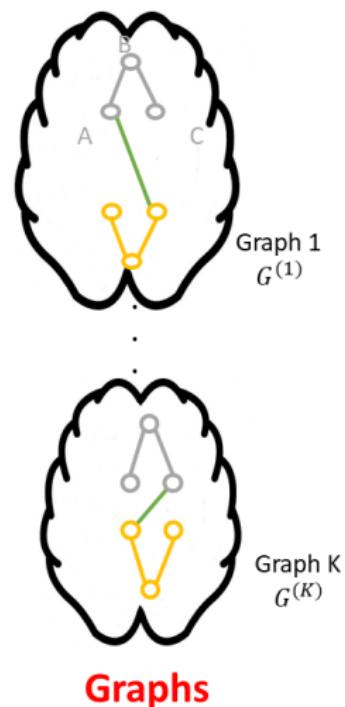
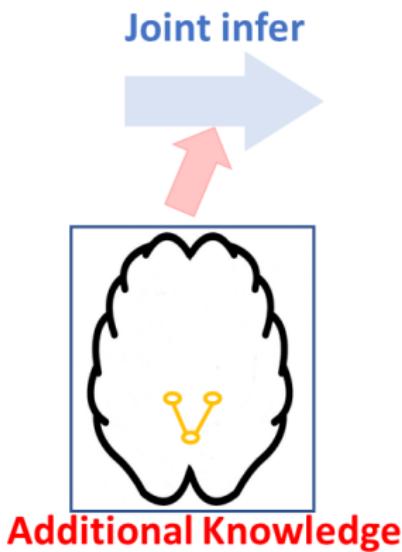
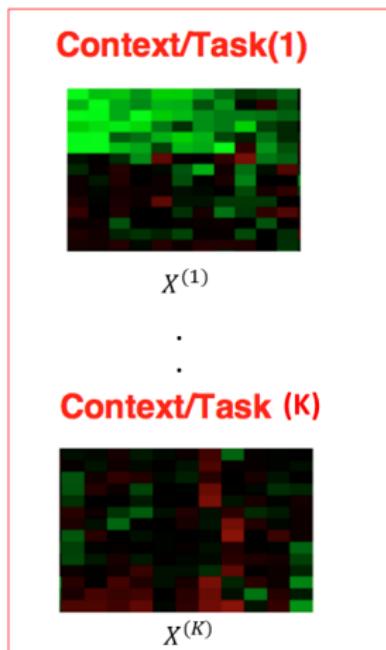
Task I: Learning multiple related graphs

- Learning multiple related graphs
- E.g., TF-TF interactions
 - Three graphs are similar

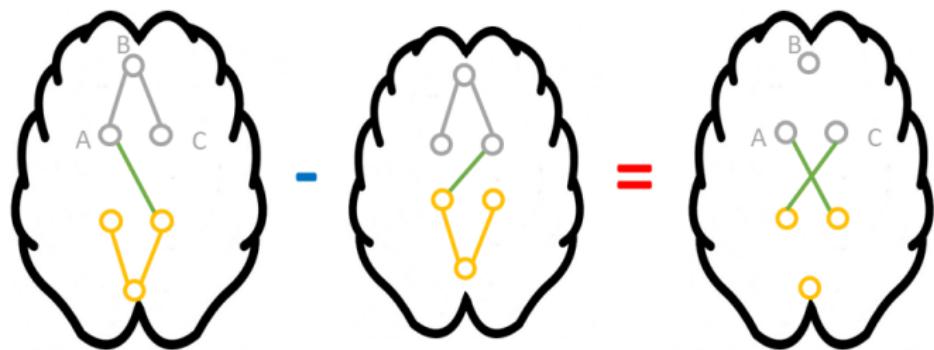


Task II: Integrating additional knowledge

- Integrating known knowledge in Learning multiple related graphs
 - E.g., known knowledge in Brain Connection



Task III: Learning sparse changes between two graphs



- A very interesting task:
 - Find differences in the brains of people with diseases, e.g. Autism, Alzheimer's
 - Use for understanding
 - Use for diagnosis

Notations

$X^{(i)}$ i -th Data matrix.

$\Sigma^{(i)}$ i -th Covariance matrix.

$\Omega^{(i)}$ i -th Inverse of covariance matrix (precision matrix).

p The total number of feature variables.

n_{tot} The total number of samples.

X^{tot} the concatenation of all Data matrices.

Σ^{tot} the concatenation of all Covariance matrices.

Ω^{tot} the concatenation of all Inverse of covariance matrices (precision matrices).

$W_I^{tot} (W_I^{(1)}, W_I^{(2)}, \dots, W_I^{(K)})$

$W_S^{tot} (W_S, W_S, \dots, W_S)$

Motivation

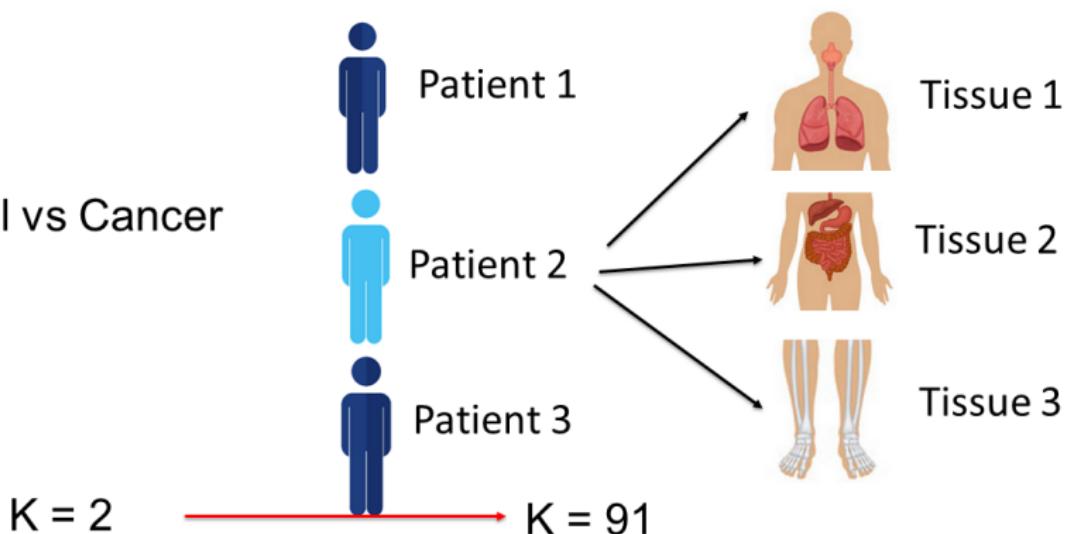
Motivation: More Num of features (p) to consider

- Yeast gene: 6K
↓
Human gene: 30K
- Words interaction, millions of words ($p > 1,000,000$)



Motivation: More num of tasks (K) to consider

Normal vs Cancer



ENCODE Project Consortium et al. An integrated encyclopedia of dna elements in the human genome. *Nature*, 489(7414):57–74, 2012.

Motivation: Limitation I – Slow Computation

| The best baseline of | Task I | Task II | Task III |
|--------------------------|------------------|--------------------|-----------------|
| Computational complexity | $O(Kp^3)$ / iter | $O(K^4 p^5)$ | $O(p^3)$ / iter |
| Bottle neck | SVD | Linear programming | SVD |

- If $K = 91$ and $p = 30K$



| The best baseline of | Task I | Task II | Task III |
|----------------------|-----------------|------------------|---------------|
| Time | 3.5 days / iter | 6 trillion years | 1 hour / iter |

- Can we have a $O(p^2)$ method?

Motivation: Limitation II – No consideration of parallelization



Computer Clusters

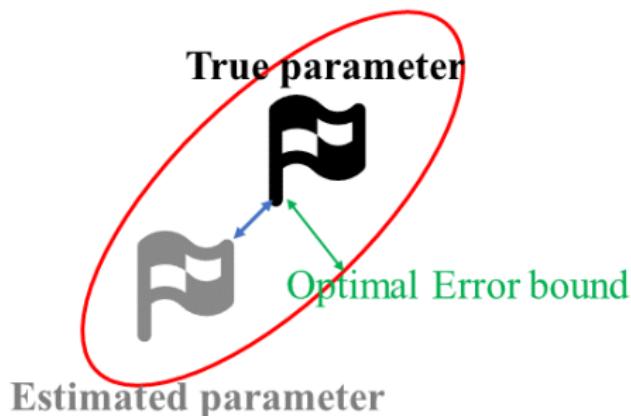


GPU

- Reduce $O(p^2)$ to $O(1)$.

Motivation: Limitation III: Lack of error bound analysis

- $\|\hat{\theta} - \theta^*\|$
- Missing analysis under a high-dimensional setting ($p \geq n$)
- No sacrifices of the accuracy from speeding-up and scaling-up the algorithm



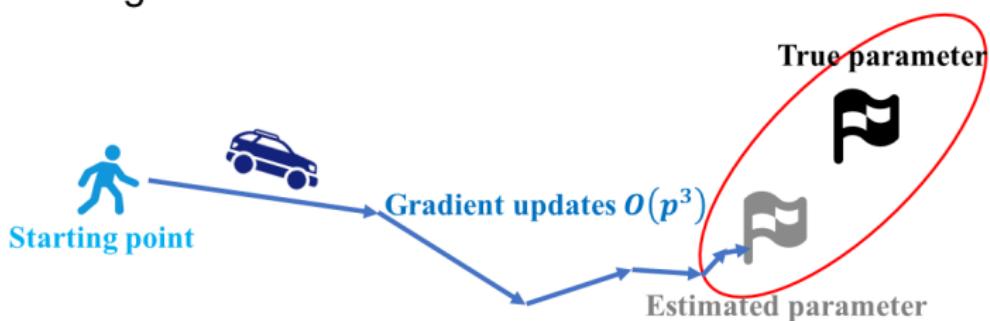
Our Aim: Fast and Scalable estimators for three types of joint graphs estimation

- Fast and scalable estimators for the three tasks
- Parallelizable algorithms
- Integrating additional knowledge
- Sharp convergence rate

Solution for Limitations - Elementary Estimator

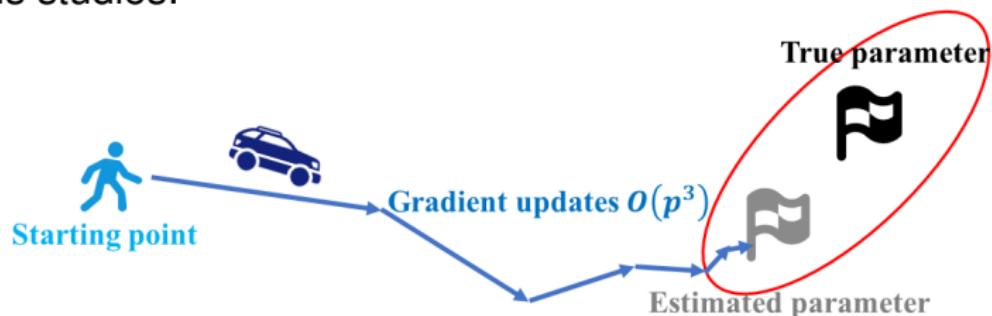
Background: summary of the previous optimization strategy

- e.g., ADMM algorithm

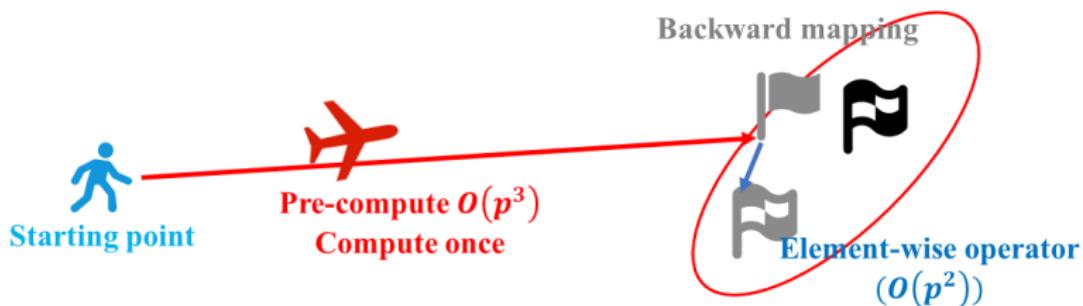


Elementary Estimator (EE) for joint sGGMs tasks

- Previous studies:

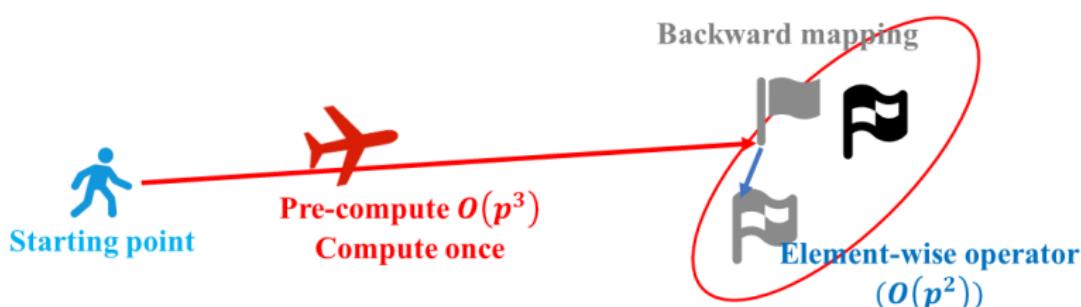


- Elementary Estimator:



Elementary Estimator (EE): Step I – Backward mapping

- Backward mapping $\mathcal{B}^*(\phi)$ of the parameter (Solution of Vanilla Maximum Likelihood Estimator (MLE))
- Vanilla MLE: $\underset{\theta}{\operatorname{argmax}} \mathcal{L}(\theta)$
 - Already close to true parameter
 - But without assumptions e.g., sparse
 - For instance, linear regression solution $(X^T X)^{-1} X^T Y$



Elementary Estimator: Step II – Optimization formulation

Elementary Estimator (EE)

$$\operatorname{argmin}_{\theta} \mathcal{R}(\theta) \quad (3.1)$$

Subject to: $\mathcal{R}^*(\theta - \mathcal{B}^*(\hat{\phi})) \leq \lambda_n$

- Let $\mathcal{R}(\cdot) = \|\cdot\|_1$



$$\operatorname{argmin}_{\theta} \|\theta\|_1 \quad (3.2)$$

Subject to: $\|\theta - \mathcal{B}^*(\hat{\phi})\|_{\infty} \leq \lambda_n$

- Easy to prove the sharp convergence rate when \mathcal{R} and \mathcal{B}^* satisfy certain conditions.

EE-Benefit: Fast and scalable solution

- A soft-thresholding operator (closed form)
- Closed form & $O(p^2)$
- Easy to parallelize in GPU

$$\hat{\theta} = \mathcal{S}_{\lambda_n}(\mathcal{B}^*(\hat{\phi}))$$

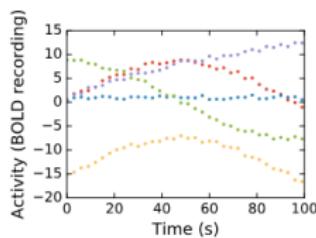
$$[S_\lambda(A)]_{ij} = \text{sign}(A_{ij}) \max(|A_{ij}| - \lambda, 0) \quad (3.3)$$

- Element-wise

$$\Sigma = \text{Cov}(X) = \begin{bmatrix} \sigma_{11} & \sigma_{12} & \cdots & \sigma_{1n} \\ \sigma_{21} & \sigma_{22} & \cdots & \sigma_{2n} \\ \vdots & \vdots & \ddots & \vdots \\ \sigma_{n1} & \sigma_{n2} & \cdots & \sigma_{nn} \end{bmatrix} \quad \Sigma = \text{Cov}(X) = \begin{bmatrix} \sigma_{11} & \sigma_{12} & \cdots & \sigma_{1n} \\ \sigma_{21} & \sigma_{22} & \cdots & \sigma_{2n} \\ \vdots & \vdots & \ddots & \vdots \\ \sigma_{n1} & \sigma_{n2} & \cdots & \sigma_{nn} \end{bmatrix} \quad \Sigma = \text{Cov}(X) = \begin{bmatrix} \sigma_{11} & \sigma_{12} & \cdots & \sigma_{1n} \\ \sigma_{21} & \sigma_{22} & \cdots & \sigma_{2n} \\ \vdots & \vdots & \ddots & \vdots \\ \sigma_{n1} & \sigma_{n2} & \cdots & \sigma_{nn} \end{bmatrix}$$

Apply same operator
Independent calculation

Background: sparse Gaussian Graphical Model (sGGM) to derive Conditional Independence Graph from data



Data

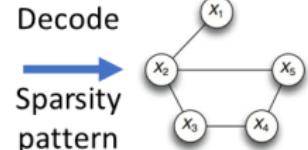
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|-------|-------|-------|-------|-------|
| 1.05 | -0.23 | 0.05 | -0.02 | 0.05 |
| -0.23 | 1.45 | -0.25 | 0.10 | -0.25 |
| 0.05 | -0.25 | 1.10 | -0.24 | 0.10 |
| -0.02 | 0.10 | -0.24 | 1.10 | -0.24 |
| 0.05 | -0.25 | 0.10 | -0.24 | 1.10 |

Covariances matrix Σ

sGGM

| | | | | |
|-----|-----|-----|-----|-----|
| 1 | 0.2 | 0 | 0 | 0 |
| 0.2 | 1 | 0.2 | 0 | 0.2 |
| 0 | 0.2 | 1 | 0.2 | 0 |
| 0 | 0 | 0.2 | 1 | 0.2 |
| 0 | 0.2 | 0 | 0.2 | 1 |

Sparse inversion Ω
(Precision Matrix)



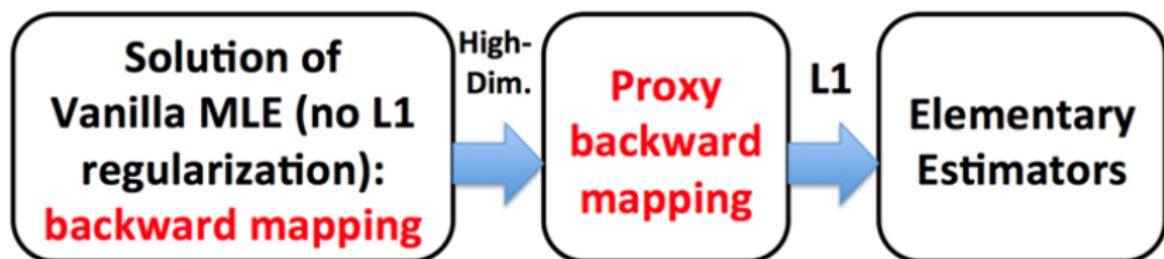
Connectome

EE-GM: Elementary Estimator for sGGM

- Vanilla MLE: $\underset{\Omega}{\operatorname{argmin}} - \log(\det(\Omega)) + \langle \Omega, \Sigma \rangle$
- Backward mapping of Ω is Σ^{-1}
- Not invertible when $p \geq n$

EE-GM: Elementary Estimator for sGGM

- Vanilla MLE: $\underset{\Omega}{\operatorname{argmin}} - \log(\det(\Omega)) + \langle \Omega, \Sigma \rangle$
- Backward mapping of Ω is Σ^{-1}
- Not invertible when $p \geq n$
- Need approximated backward mapping
 - proxy backward mapping $\hat{\theta}_n \approx \mathcal{B}^*(\hat{\phi})$
 - In sGGM, $\hat{\theta}_n = [T_v(\hat{\Sigma})]^{-1}$



EE-GM: Elementary Estimator for sGGM

$$\operatorname{argmin}_{\theta} \|\theta\|_1 \quad (3.4)$$

Subject to: $\|\theta - \mathcal{B}^*(\hat{\phi})\|_{\infty} \leq \lambda_n$

$$\hat{\theta}_n = [T_V(\hat{\Sigma})]^{-1} \quad \Downarrow$$

EE-sGGM

$$\operatorname{argmin}_{\Omega} \|\Omega\|_{1,\text{off}} \quad (3.5)$$

subject to: $\|\Omega - [T_V(\hat{\Sigma})]^{-1}\|_{\infty,\text{off}} \leq \lambda_n$

- | EE | $\mathcal{R}(\cdot)$ | θ | $\hat{\theta}_n$ | \mathcal{R}^* |
|---------|----------------------|----------|----------------------------|----------------------|
| EE-sGGM | $\ \cdot\ _1$ | Ω | $[T_V(\hat{\Sigma})]^{-1}$ | $\ \cdot\ _{\infty}$ |

EE-Benefit: Easy to prove error bound

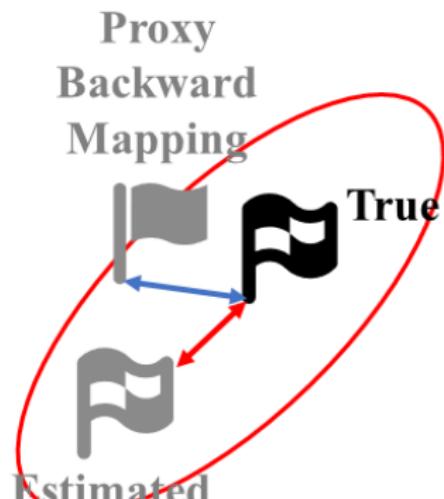
- Error bound:

$$\begin{aligned} \|\hat{\theta} - \theta^*\|_\infty &\leq 2\lambda_n \\ \|\hat{\theta} - \theta^*\|_F &\leq 4\sqrt{s}\lambda_n \\ \|\hat{\theta} - \theta^*\|_1 &\leq 8s\lambda_n \end{aligned} \quad (3.6)$$

- Condition:

$$\lambda_n \geq \|\hat{\theta}_n - \theta^*\|_\infty \quad (3.7)$$

- Constant: s is the num of non-zero entries.



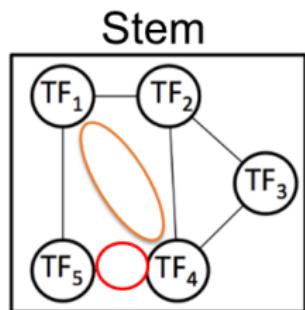
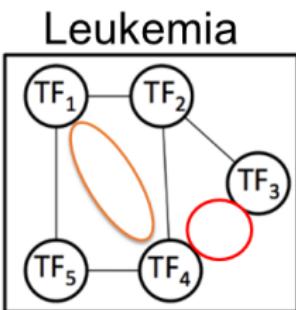
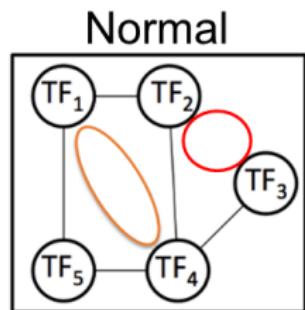
Method I: FASJEM

Outline

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 - Method
 - Results
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 - Background
 - Method
 - Results
- 6 Method III: DIFFEE
 - Method
 - Results
- 7 Discussion
 - Questions from Proposal
 - Future works

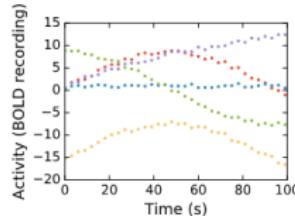
Task I: Learning multiple related graphs

- Learning multiple related graphs
- E.g., TF-TF interactions
 - Three graphs are similar



Background: Multi-task sGGMs

- A pipeline to infer Multiple Related Graphs from heterogeneous datasets $\mathbf{X}^{(1)}, \dots, \mathbf{X}^{(K)}$ ¹.

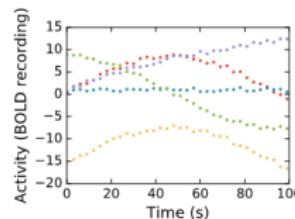
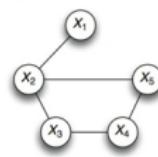


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| -0.02 | 0.10 | -0.24 | 1.10 | -0.24 |
| 0.05 | -0.25 | 0.10 | -0.24 | 1.10 |

Multi-task
GGM

| | | | | |
|-----|-----|-----|-----|-----|
| 1 | 0.2 | 0 | 0 | 0 |
| 0.2 | 1 | 0.2 | 0 | 0.2 |
| 0 | 0.2 | 1 | 0.2 | 0 |
| 0 | 0 | 0.2 | 1 | 0.2 |
| 0 | 0.2 | 0 | 0.2 | 1 |

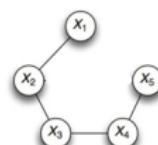
Decode
Sparsity pattern



| | | | | |
|-------|-------|-------|-------|-------|
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| 0.05 | -0.25 | 1.10 | -0.24 | 0.10 |
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| | | | | |
|-----|-----|-----|-----|-----|
| 1 | 0.2 | 0 | 0 | 0 |
| 0.2 | 1 | 0.2 | 0 | 0 |
| 0 | 0.2 | 1 | 0.2 | 0 |
| 0 | 0 | 0.2 | 1 | 0.2 |
| 0 | 0 | 0 | 0.2 | 1 |

Decode
Sparsity pattern



¹ X^{tot} : the concatenation of $(X^{(1)}, X^{(2)}, \dots, X^{(K)})$.

Σ^{tot} : the concatenation of $(\Sigma^{(1)}, \Sigma^{(2)}, \dots, \Sigma^{(K)})$.

Ω^{tot} : the concatenation of $(\Omega^{(1)}, \Omega^{(2)}, \dots, \Omega^{(K)})$.

Background: Joint Graphical Lasso

Graphical Lasso

$$\underset{\Omega}{\operatorname{argmin}} -\log \det(\Omega) + \langle \Omega, \Sigma \rangle + \lambda_n \|\Omega\|_1 \quad (4.1)$$

- Add $\mathcal{R}'(\cdot)$



Joint Graphical Lasso

$$\begin{aligned} & \underset{\Omega^{(i)} > 0}{\operatorname{argmin}} \sum_i (-L(\Omega^{(i)}) + \lambda_1 \sum_i \|\Omega^{(i)}\|_1 \\ & \quad + \lambda_2 \mathcal{R}'(\Omega^{(1)}, \Omega^{(2)}, \dots, \Omega^{(K)})) \end{aligned} \quad (4.2)$$

- $\Omega_{tot} = (\Omega^{(1)}, \Omega^{(2)}, \dots, \Omega^{(K)})$.

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Enforcing relatedness of multiple graphs through Regularization: FASJEM-norm

EE-sGGM

$$\underset{\Omega}{\operatorname{argmin}} \|\Omega\|_{1,\text{off}} \quad (4.3)$$

subject to: $\|\Omega - [T_V(\widehat{\Sigma})]^{-1}\|_{\infty,\text{off}} \leq \lambda_n$

- Add $\mathcal{R}'(\cdot)$



FASJEM-norm

$$\mathcal{R}(\Omega_{tot}) = \|\Omega_{tot}\|_1 + \mathcal{R}'(\Omega_{tot}) \quad (4.4)$$

Elementary Estimator (EE)

$$\operatorname{argmin}_{\theta} \mathcal{R}(\theta) \quad (4.5)$$

Subject to: $\mathcal{R}^*(\theta - \mathcal{B}^*(\hat{\phi})) \leq \lambda_n$

| EE | $\mathcal{R}(\cdot)$ | θ | $\hat{\theta}_n$ | $\mathcal{R}^*(\cdot)$ |
|---------|------------------------------|----------------|--------------------------------|--|
| EE-sGGM | $\ \cdot\ _1$ | Ω | $[T_v(\hat{\Sigma})]^{-1}$ | $\ \cdot\ _\infty$ |
| FASJEM | $\ \cdot\ _1 + \mathcal{R}'$ | Ω^{tot} | $inv[T_v(\hat{\Sigma}^{tot})]$ | $\max(\ \cdot\ _\infty, \mathcal{R}'^*)$ |

FASJEM

$$\begin{aligned} & \operatorname{argmin}_{\Omega_{tot}} \|\Omega_{tot}\|_1 + \mathcal{R}'(\Omega_{tot}) \\ & s.t. \|\Omega_{tot} - inv(T_v(\hat{\Sigma}_{tot}))\|_\infty \leq \lambda_n \end{aligned} \quad (4.6)$$

$$\mathcal{R}'^*(\Omega_{tot} - inv(T_v(\hat{\Sigma}_{tot}))) \leq \lambda_n$$

FASJEM: Variations

- FASJEM-G:

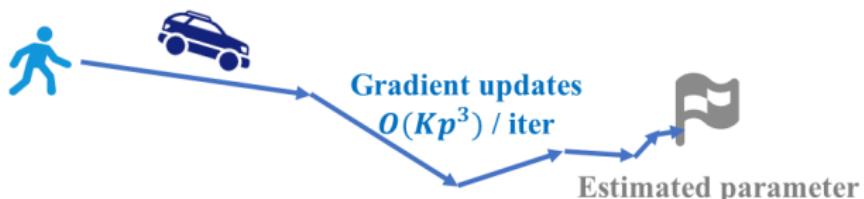
$$\begin{aligned}\mathcal{R}'(\cdot) &= \|\cdot\|_{\mathcal{G},2} \\ \|\Omega_{tot}\|_{\mathcal{G},2} &= \sum_{j=1}^p \sum_{k=1}^p \|(\Omega_{j,k}^{(1)}, \Omega_{j,k}^{(2)}, \dots, \Omega_{j,k}^{(i)}, \dots, \Omega_{j,k}^{(K)})\|_2\end{aligned}\tag{4.7}$$

- FASJEM-I:

$$\begin{aligned}\mathcal{R}'(\cdot) &= \|\cdot\|_{\mathcal{G},\infty} \\ \|\Omega_{tot}\|_{\mathcal{G},\infty} &= \sum_{j=1}^p \sum_{k=1}^p \|(\Omega_{j,k}^{(1)}, \Omega_{j,k}^{(2)}, \dots, \Omega_{j,k}^{(i)}, \dots, \Omega_{j,k}^{(K)})\|_\infty\end{aligned}\tag{4.8}$$

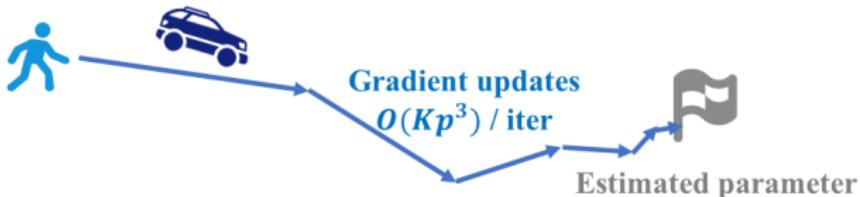
FASJEM: Optimization Solution

- JGL solution:

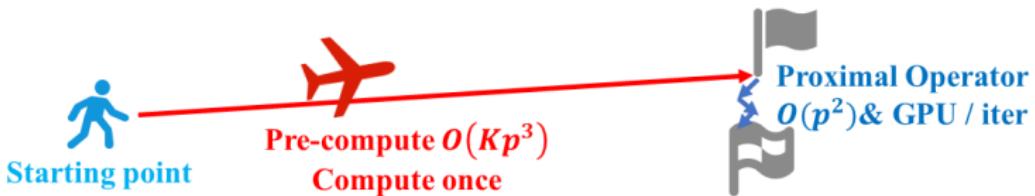


FASJEM: Optimization Solution

- JGL solution:

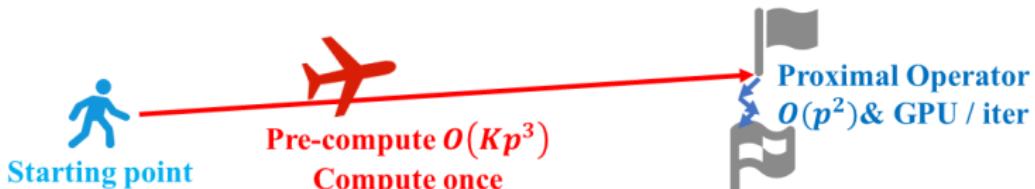


- FASJEM solution:



FASJEM: Optimization Solution – Proximal algorithm

- FASJEM solution:



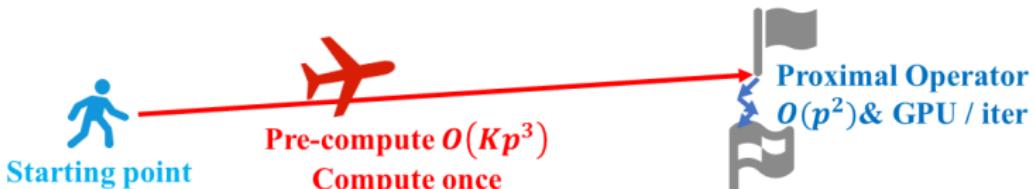
- In each iteration, a proximal operator
- Element-wise operator, $O(p^2)$

$$\text{prox}_{\gamma \|\cdot\|_1}(x) = \begin{cases} x_{j,k}^{(i)} - \gamma, & x_{j,k}^{(i)} > \gamma \\ 0, & |x_{j,k}^{(i)}| \leq \gamma \\ x_{j,k}^{(i)} + \gamma, & x_{j,k}^{(i)} < -\gamma \end{cases} \quad (4.9)$$

$$\begin{aligned} \text{prox}_{\gamma \|\cdot\|_1}(x) &= \max((x_{j,k}^{(i)} - \gamma), 0) \quad (4.10) \\ &\quad + \min(0, (x_{j,k}^{(i)} + \gamma)) \end{aligned}$$

FASJEM: Optimization Solution – Proximal algorithm

- FASJEM solution:



- In each iteration, a proximal operator
- Element-wise operator, $O(p^2)$
- GPU-parallelizable $O(1)$
 - e.g., proximity of ℓ_1

$$\text{prox}_{\gamma \|\cdot\|_1}(x) = \begin{cases} x_{j,k}^{(i)} - \gamma, & x_{j,k}^{(i)} > \gamma \\ 0, & |x_{j,k}^{(i)}| \leq \gamma \\ x_{j,k}^{(i)} + \gamma, & x_{j,k}^{(i)} < -\gamma \end{cases} \quad \Rightarrow \quad \begin{aligned} \text{prox}_{\gamma \|\cdot\|_1}(x) &= \max((x_{j,k}^{(i)} - \gamma), 0) \quad (4.10) \\ &+ \min(0, (x_{j,k}^{(i)} + \gamma)) \end{aligned}$$

(4.9)

FASJEM: Computational Complexity

| The best baseline of | Task I | Task II | Task III |
|--------------------------|------------------|--------------------|-----------------|
| Computational complexity | $O(Kp^3)$ / iter | $O(K^4 p^5)$ | $O(p^3)$ / iter |
| Bottle neck | SVD | Linear programming | SVD |
| Our approach | FASJEM | | |
| Computational complexity | $O(Kp^2)$ / iter | | |
| Parallelization | $O(K)$ / iter | | |

Summary

| | EE | $\mathcal{R}(\cdot)$ | θ | $\widehat{\theta}_n$ | $\mathcal{R}^*(\cdot)$ |
|----------|---------|------------------------------|----------------|------------------------------------|--|
| | EE-sGGM | $\ \cdot\ _1$ | Ω | $[T_v(\widehat{\Sigma})]^{-1}$ | $\ \cdot\ _\infty$ |
| Task I | FASJEM | $\ \cdot\ _1 + \mathcal{R}'$ | Ω^{tot} | $inv[T_v(\widehat{\Sigma}^{tot})]$ | $\max(\ \cdot\ _\infty, \mathcal{R}'^*)$ |
| Task II | | | | | |
| Task III | | | | | |

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Results: Theoretical Analysis

- $p' = \max(Kp, n_{tot})$
- **Error Bound:** $\|\widehat{\Omega}_{tot} - \Omega_{tot}^*\|_F \leq 32 \frac{4\kappa_1 a}{\kappa_2} \sqrt{\frac{s \log p'}{n_{tot}}}$

| Multi-task: | K Single-task: |
|--|------------------------------------|
| $O\left(\frac{\log(Kp)}{n_{tot}}\right)$ | $O\left(\frac{\log p}{n_i}\right)$ |

- By assuming $n_i = \frac{n_{tot}}{K}$:

Results: Theoretical Analysis

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- By assuming $n_i = \frac{n_{tot}}{K}$:
- We can conclude that $\frac{\log(Kp)}{n_{tot}} < K \frac{\log p}{n_{tot}}$

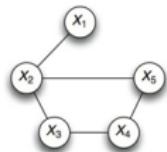
Results: Theoretical Analysis

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- **Error Bound:** $\|\widehat{\Omega}_{tot} - \Omega_{tot}^*\|_F \leq 32 \frac{4\kappa_1 a}{\kappa_2} \sqrt{\frac{s \log p'}{n_{tot}}}$

| Multi-task: | K Single-task: |
|--|------------------------------------|
| $O\left(\frac{\log(Kp)}{n_{tot}}\right)$ | $O\left(\frac{\log p}{n_i}\right)$ |

- By assuming $n_i = \frac{n_{tot}}{K}$:
- We can conclude that $\frac{\log(Kp)}{n_{tot}} < K \frac{\log p}{n_{tot}}$
- This indicates that the multi-task estimator is better!!!

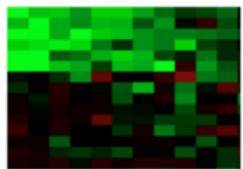
Results: Synthetic Data generation process



| | | | | |
|-----|-----|-----|-----|-----|
| 1 | 0.2 | 0 | 0 | 0 |
| 0.2 | 1 | 0.2 | 0 | 0.2 |
| 0 | 0.2 | 1 | 0.2 | 0 |
| 0 | 0 | 0.2 | 1 | 0.2 |
| 0 | 0.2 | 0 | 0.2 | 1 |

| | | | | |
|-------|-------|-------|-------|-------|
| 1.05 | -0.23 | 0.05 | -0.02 | 0.05 |
| -0.23 | 1.45 | -0.25 | 0.10 | -0.25 |
| 0.05 | -0.25 | 1.10 | -0.24 | 0.10 |
| -0.02 | 0.10 | -0.24 | 1.10 | -0.24 |
| 0.05 | -0.25 | 0.10 | -0.24 | 1.10 |

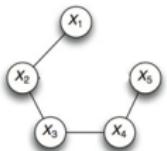
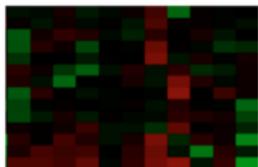
Context/Task(1)



Inverse

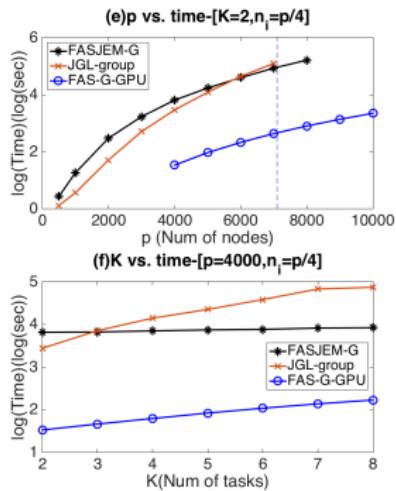
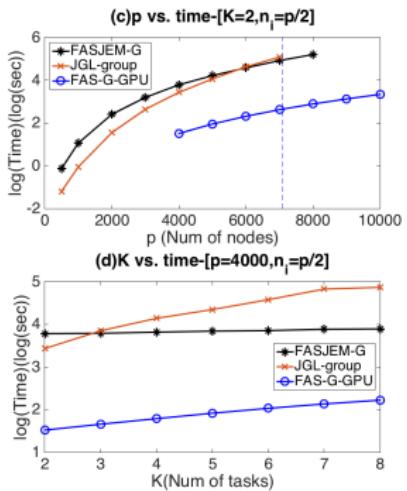
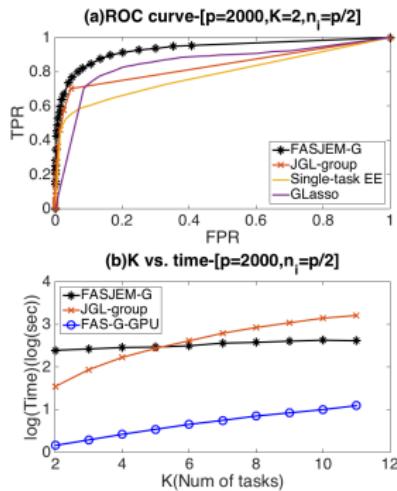
| | | | | |
|-------|-------|-------|-------|-------|
| 1.05 | -0.23 | 0.05 | -0.02 | 0.05 |
| -0.23 | 1.45 | -0.25 | 0.10 | -0.10 |
| 0.05 | -0.25 | 1.10 | -0.24 | 0.10 |
| -0.02 | 0.10 | -0.24 | 1.10 | -0.24 |
| 0.05 | -0.10 | 0.10 | -0.24 | 1.10 |

Context/Task(2)



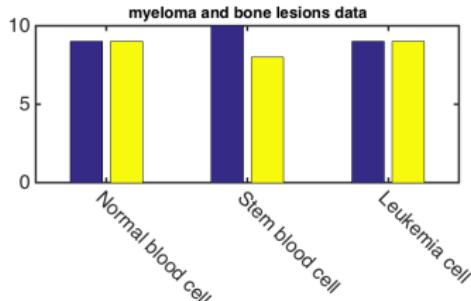
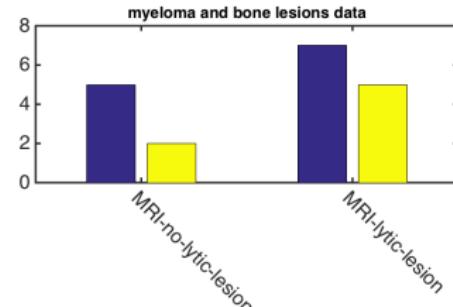
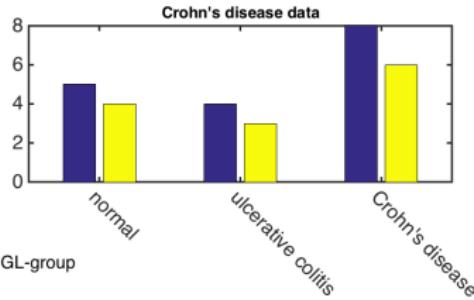
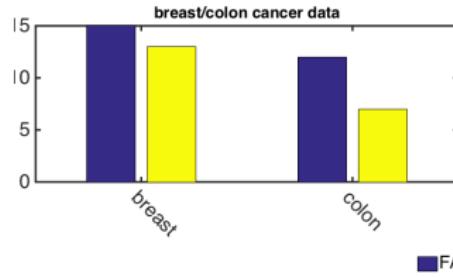
| | | | | |
|-----|-----|-----|-----|-----|
| 1 | 0.2 | 0 | 0 | 0 |
| 0.2 | 1 | 0.2 | 0 | 0 |
| 0 | 0.2 | 1 | 0.2 | 0 |
| 0 | 0 | 0.2 | 1 | 0.2 |
| 0 | 0 | 0 | 0.2 | 1 |

Results: Synthetic Data Results



Results: Real-world Data Results – Number of Matched Edges versus the Existing Domain Databases

- Validation by counting the overlapped interactions according to the existing bio-databases (MInact)



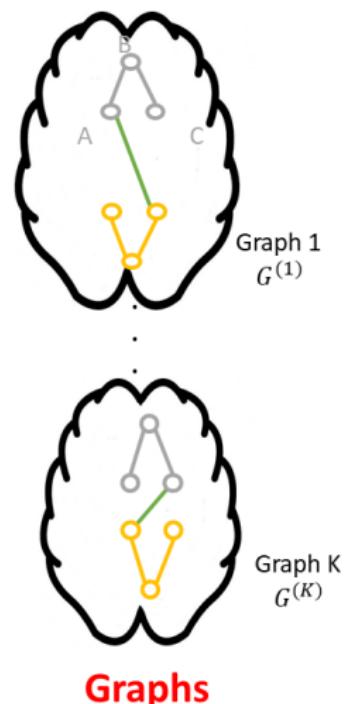
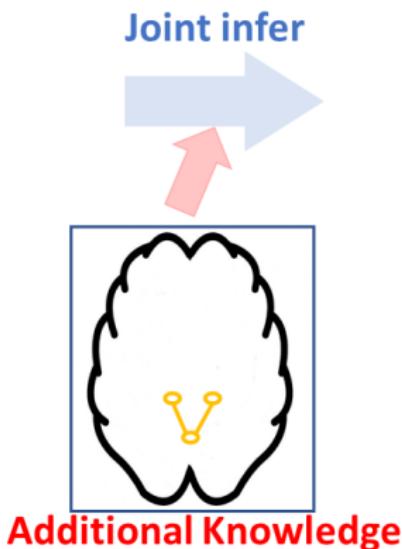
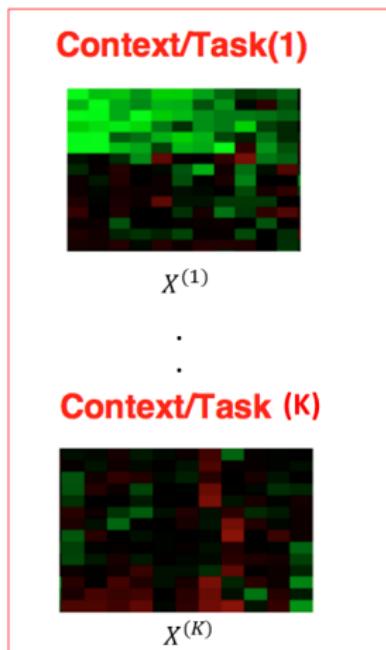
Method II: JEEK

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Task II: Integrating additional knowledge

- Integrating known knowledge in Learning multiple related graphs
 - E.g., known knowledge in Brain Connection



Solution: Using Knowledge as Weight in Regularization (KW-norm)

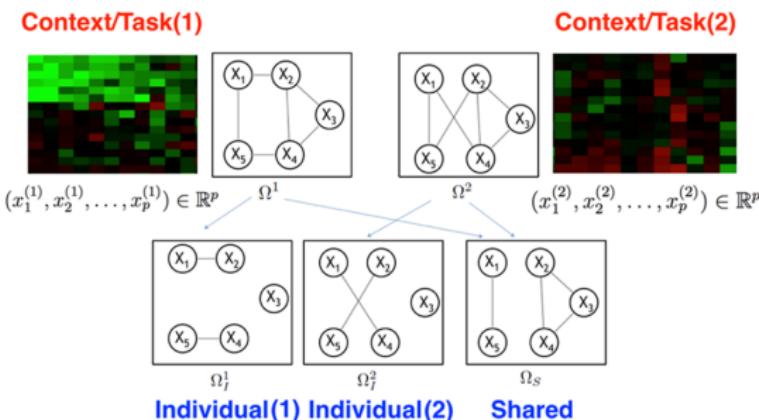
- Integrating additional knowledge through a novel regularization function $\mathcal{R}(\cdot)$

KW-norm

$$\mathcal{R}(\{\Omega^{(i)}\}) = \sum_{i=1}^K \|W_I^{(i)} \circ \Omega_I^{(i)}\|_1 + \sum_{i=1}^K \|W_S \circ \Omega_S\|_1 \quad (5.1)$$

- $\Omega^{(i)} = \Omega_I^{(i)} + \Omega_S$
- $\{W_I^{(i)}\}$: weights describing knowledge of each individual graph.
- W_S : weights describing knowledge of the shared graph.

Background: Shared and Task-Specific Subgraph Representation



- Know both
 - House keeping interactions
 - Context-specific networks

Solution: Using Knowledge as Weight in Regularization (KW-norm)

- Use *tot* notation

KW-norm

$$\mathcal{R}(\Omega^{tot}) = ||W_I^{tot} \circ \Omega_I^{tot}||_1 + ||W_S^{tot} \circ \Omega_S^{tot}||_1 \quad (5.2)$$

- W_I^{tot} : weights describing knowledge of each individual graph.
- W_S^{tot} : weights describing knowledge of the shared graph.

Solution: Using Knowledge as Weight in Regularization (KW-norm)

- Use *tot* notation

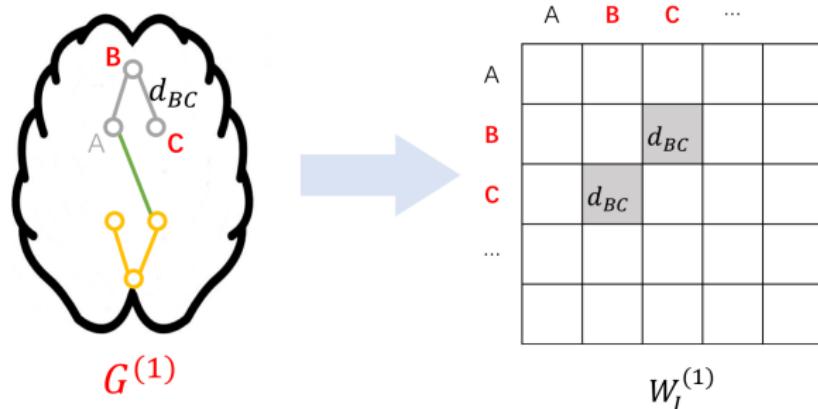
KW-norm

$$\mathcal{R}(\Omega^{tot}) = ||W_I^{tot} \circ \Omega_I^{tot}||_1 + ||W_S^{tot} \circ \Omega_S^{tot}||_1 \quad (5.2)$$

- W_I^{tot} : weights describing knowledge of each individual graph.
- W_S^{tot} : weights describing knowledge of the shared graph.
- No need to design knowledge-specific optimization
- KW-norm is **flexible**.

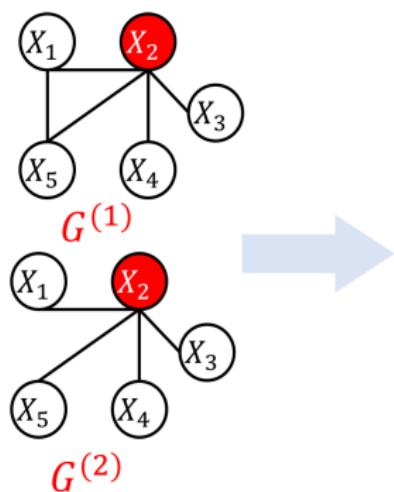
Example I: KW-norm representing the edge-level knowledge

- e.g., Spatial distance among brain regions;



Example II: KW-norm describing the node-level knowledge

- e.g., X_2 is a known hub node;



| | 1 | 2 | 3 | 4 | 5 |
|---|------------|------------|------------|------------|------------|
| 1 | | $1/\gamma$ | 1 | 1 | 1 |
| 2 | $1/\gamma$ | | $1/\gamma$ | $1/\gamma$ | $1/\gamma$ |
| 3 | 1 | $1/\gamma$ | | 1 | 1 |
| 4 | 1 | $1/\gamma$ | 1 | | 1 |
| 5 | 1 | $1/\gamma$ | 1 | 1 | |

W_s

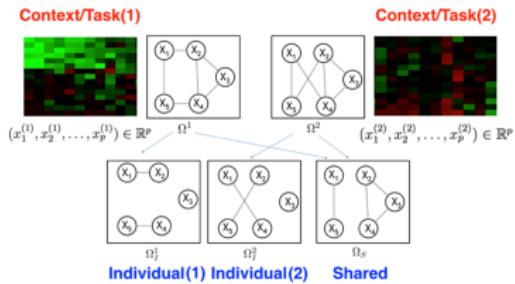
Background: SIMULE

- Decompose $\Omega^{(i)} = \Omega_I^{(i)} + \Omega_S$

- An ℓ_1 minimization approach

$$\hat{\Omega}_I^{(1)}, \hat{\Omega}_I^{(2)}, \dots, \hat{\Omega}_I^{(K)}, \hat{\Omega}_S = \operatorname{argmin}_{\Omega_I^{(i)}, \Omega_S} \sum_i \|\Omega_I^{(i)}\|_1 + \epsilon K \|\Omega_S\|_1$$

Subject to: $\|\Sigma^{(i)}(\Omega_I^{(i)} + \Omega_S) - I\|_\infty \leq \lambda_n, i = 1, \dots, K$



Background: WSIMULE: A weighted SIMULE estimator

SIMULE

$$\widehat{\Omega}_I^{(1)}, \widehat{\Omega}_I^{(2)}, \dots, \widehat{\Omega}_I^{(K)}, \widehat{\Omega}_S = \operatorname{argmin}_{\Omega_I^{(i)}, \Omega_S} \sum_i \|\Omega_I^{(i)}\|_1 + \epsilon K \|\Omega_S\|_1$$

Subject to: $\|\Sigma^{(i)}(\Omega_I^{(i)} + \Omega_S) - I\|_\infty \leq \lambda_n, i = 1, \dots, K$

- ADD $W_I^{(i)}, W_S$



W-SIMULE

$$\widehat{\Omega}_I^{(1)}, \dots, \widehat{\Omega}_I^{(K)}, \widehat{\Omega}_S = \operatorname{argmin}_i \|\mathcal{W}_I^{(i)} \circ \Omega_I^{(i)}\|_1 + K \|\mathcal{W}_S \circ \Omega_S\|_1 \quad (5.3)$$

Subject to: $\|\Sigma^{(i)}(\Omega_I^{(i)} + \Omega_S) - I\|_\infty \leq \lambda, i = 1, \dots, K.$

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Proposed Method: Combine EE and KW-norm

Elementary Estimator

$$\operatorname{argmin}_{\theta} \mathcal{R}(\theta) \quad (5.4)$$

Subject to: $\mathcal{R}^*(\theta - \mathcal{B}^*(\hat{\phi})) \leq \lambda_n$

+

KW-norm

$$\mathcal{R}(\Omega^{tot}) = ||W_I^{tot} \circ \Omega_I^{tot}||_1 + ||W_S^{tot} \circ \Omega_S^{tot}||_1 \quad (5.5)$$

Proposed Method: Joint Elementary Estimator incorporating additional Knowledge (JEEK)

| EE | $\mathcal{R}(\cdot)$ | θ | $\widehat{\theta}_n$ | $\mathcal{R}^*(\cdot)$ |
|---------|----------------------|----------------|------------------------------------|------------------------|
| EE-sGGM | $\ \cdot\ _1$ | Ω | $[T_v(\widehat{\Sigma})]^{-1}$ | $\ \cdot\ _\infty$ |
| JEEK | kw-norm | Ω^{tot} | $inv[T_v(\widehat{\Sigma}^{tot})]$ | kw-dual |

JEEK

$$\operatorname{argmin}_{\Omega_I^{tot}, \Omega_S^{tot}} \|W_I^{tot} \circ \Omega_I^{tot}\|_1 + \|W_S^{tot} \circ \Omega_S^{tot}\|$$

$$\text{Subject to: } \|W_I^{tot} \circ (\Omega^{tot} - inv(T_v(\widehat{\Sigma}^{tot})))\|_\infty \leq \lambda_n \quad (5.6)$$

$$\|W_S^{tot} \circ (\Omega^{tot} - inv(T_v(\widehat{\Sigma}^{tot})))\|_\infty \leq \lambda_n$$

$$\Omega^{tot} = \Omega_S^{tot} + \Omega_I^{tot}$$

Proposed method: JEEK – Solution

- Fast and Scalable solution² – p^2 small linear programming subproblems with only $K + 1$ variables:

$$\operatorname{argmin}_{a_i, b} \sum_i |w_i a_i| + K |w_s b|$$

$$\text{Subject to: } |a_i + b - c_i| \leq \frac{\lambda_n}{\min(w_i, w_s)}, \quad (5.7)$$
$$i = 1, \dots, K$$

² $a_i := \Omega_{I,j,k}^{(i)}$ (the $\{j, k\}$ -th entry of $\Omega^{(i)}$)

$b := \Omega_{Sj,k}$

$c_i = [T_V(\widehat{\Sigma}^{(i)})]_{j,k}^{-1}$.

$W_{j,k}^{(i)} = w_i$ and $W_{j,k}^S = w_s$.

Why JEEK is better

- Rich and flexible for integrating additional knowledge
 - e.g., spatial, anatomy, hub, pathway, location, known edges;

Why JEEK is better

- Rich and flexible for integrating additional knowledge
 - e.g., spatial, anatomy, hub, pathway, location, known edges;
- Parallelizable optimization with small sub-problems.

Why JEEK is better

- Rich and flexible for integrating additional knowledge
 - e.g., spatial, anatomy, hub, pathway, location, known edges;
- Parallelizable optimization with small sub-problems.
- Theoretical guaranteed

JEEK: Computational Complexity

| The best baseline of computational complexity | Task I | Task II | Task III |
|---|------------------|--------------------|-----------------|
| Computational complexity | $O(Kp^3)$ / iter | $O(K^4p^5)$ | $O(p^3)$ / iter |
| Bottle neck | SVD | Linear programming | SVD |
| Our approach | FASJEM | JEEK | |
| Computational complexity | $O(Kp^2)$ / iter | $O(K^4p^2)$ | |
| Parallelization | $O(K)$ / iter | $O(K^4)$ | |

Summary

| | EE | $\mathcal{R}(\cdot)$ | θ | $\widehat{\theta}_n$ | $\mathcal{R}^*(\cdot)$ |
|----------|---------|------------------------------|----------------|------------------------------------|--|
| | EE-sGGM | $\ \cdot\ _1$ | Ω | $[T_v(\widehat{\Sigma})]^{-1}$ | $\ \cdot\ _\infty$ |
| Task I | FASJEM | $\ \cdot\ _1 + \mathcal{R}'$ | Ω^{tot} | $inv[T_v(\widehat{\Sigma}^{tot})]$ | $\max(\ \cdot\ _\infty, \mathcal{R}'^*)$ |
| Task II | JEEK | kw-norm | Ω^{tot} | $inv[T_v(\widehat{\Sigma}^{tot})]$ | kw-dual |
| Task III | | | | | |

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Theoretical Results

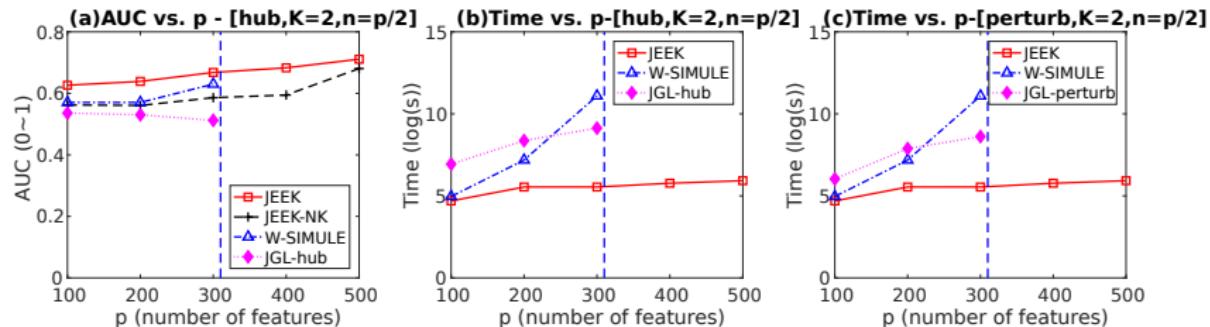
- Sharp convergence rate as the state-of-art

$$\begin{aligned} \|\widehat{\Omega}^{tot} - \Omega^{tot*}\|_F &\leq 4\sqrt{k_i + k_s}\lambda_n \\ \max(\|W_I^{tot} \circ (\widehat{\Omega}^{tot} - \Omega^{tot*})\|_\infty, \|W_S^{tot} \circ (\widehat{\Omega}^{tot} - \Omega^{tot*})\|_\infty) &\leq 2\lambda_n \quad (5.8) \\ \|W_I^{tot} \circ (\widehat{\Omega}_I^{tot} - \Omega_I^{tot*})\|_1 + \|W_S^{tot} \circ (\widehat{\Omega}_S^{tot} - \Omega_S^{tot*})\|_1 &\leq 8(k_i + k_s)\lambda_n \end{aligned}$$

Where a , c , κ_1 and κ_2 are constants

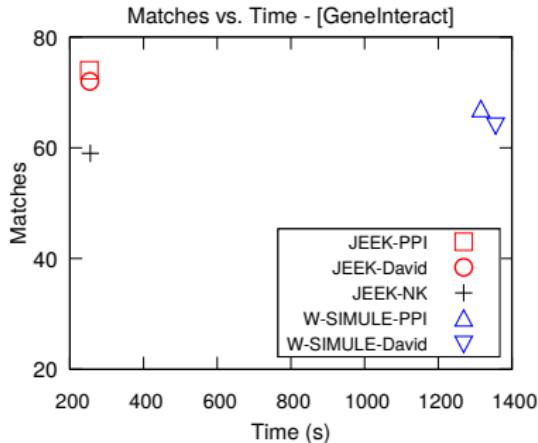
$$\begin{aligned} \|\widehat{\Omega}^{tot} - \Omega^{tot*}\|_F \\ \leq \frac{16\kappa_1 a \max_{j,k}(W_I^{tot}_{j,k}, W_S^{tot}_{j,k})}{\kappa_2} \sqrt{\frac{(k_i + k_s) \log(Kp)}{n_{tot}}} \quad (5.9) \end{aligned}$$

Empirical Results on Multiple Synthetic Datasets

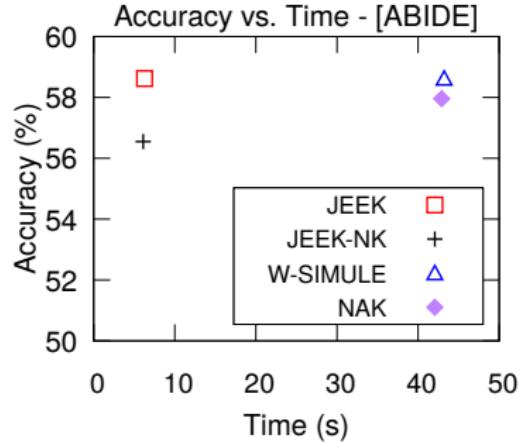


- **JEEK** outperforms the speed of the state-of arts significantly faster ($\sim 5000\times$ improvement);
- **JEEK** obtains better AUC as the state-of-the-art;
- **JEEK** obtains better AUC than JEEK-NK (no additional knowledge).

Empirical Results on Two Real-world Datasets



(a)



(b)

- (a). On real-world gene expression data about leukemia cells vs. normal blood cells. Used multiple types of additional knowledge;
- (b). On real-world Brain fMRI dataset: ABIDE. Using LDA as a downstream classification for evaluating JEEK vs. baselines.

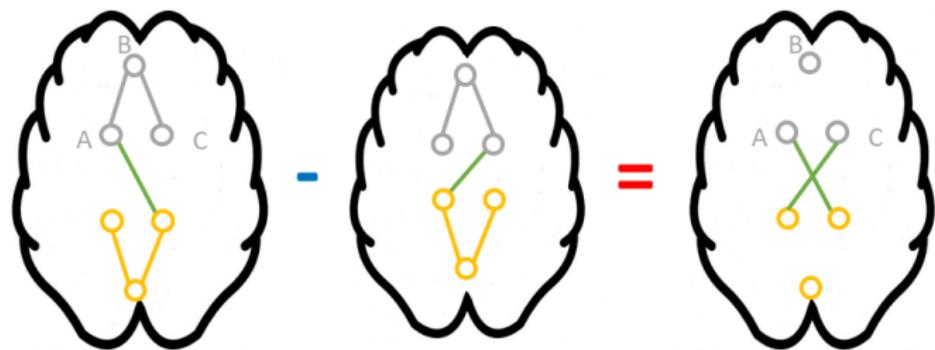
Method III: DIFFEE

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Takes III: Learning sparse changes between two graphs

- Each graph may be dense or sparse, differential net is sparse



Proposed Method III: DIFFEE

- Two cases : d (disease) & c (control)

$$\begin{aligned} & \underset{\theta}{\operatorname{argmin}} \|\theta\|_1 \\ \text{Subject to: } & \quad (6.1) \quad \Delta = \Omega_d - \Omega_c \\ & \|\theta - \mathcal{B}^*(\hat{\phi})\|_{\infty} \leq \lambda_n \end{aligned}$$

$$\begin{aligned} & \underset{\Delta}{\operatorname{argmin}} \|\Delta\|_1 \\ \text{Subject to: } & \quad (6.2) \quad \|\Delta - \mathcal{B}^*(\hat{\Sigma}_d, \hat{\Sigma}_c)\|_{\infty} \leq \lambda_n \end{aligned}$$

Proposed Method III: DIFFEE

Elementary Estimator (EE)

$$\operatorname{argmin}_{\theta} \mathcal{R}(\theta) \quad (6.3)$$

Subject to: $\mathcal{R}^*(\theta - \mathcal{B}^*(\hat{\phi})) \leq \lambda_n$

| EE | $\mathcal{R}(\cdot)$ | θ | $\hat{\theta}_n$ | $\mathcal{R}^*(\cdot)$ |
|---------|----------------------|----------|--|------------------------|
| EE-sGGM | $\ \cdot\ _1$ | Ω | $[T_v(\hat{\Sigma})]^{-1}$ | $\ \cdot\ _\infty$ |
| DIFFEE | $\ \cdot\ _1$ | Δ | $\left([T_v(\hat{\Sigma}_d)]^{-1} - [T_v(\hat{\Sigma}_c)]^{-1} \right)$ | $\ \cdot\ _\infty$ |

DIFFEE

$$\operatorname{argmin}_{\Delta} \|\Delta\|_1 \quad (6.4)$$

Subject to: $\|\Delta - \left([T_v(\hat{\Sigma}_d)]^{-1} - [T_v(\hat{\Sigma}_c)]^{-1} \right)\|_\infty \leq \lambda_n$

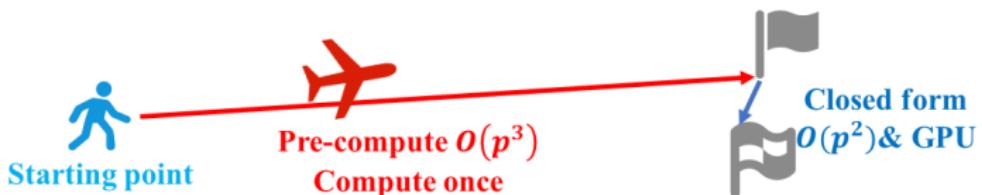
DIFFEE: Optimization Solution

- Close form

$$\widehat{\Delta} = S_{\lambda_n}([T_v(\widehat{\Sigma}_d)]^{-1} - [T_v(\widehat{\Sigma}_c)]^{-1}) \quad (6.5)$$

$$[S_\lambda(A)]_{ij} = \text{sign}(A_{ij}) \max(|A_{ij}| - \lambda, 0) \quad (6.6)$$

- GPU-parallelizable



DIFFEE: Computational Complexity

| The best baseline of | Task I | Task II | Task III |
|--------------------------|------------------|--------------------|-----------------|
| Computational complexity | $O(Kp^3)$ / iter | $O(K^4p^5)$ | $O(p^3)$ / iter |
| Bottle neck | SVD | Linear programming | SVD |
| Our approach | FASJEM | JEEK | DIFFEE |
| Computational complexity | $O(Kp^2)$ / iter | $O(K^4p^2)$ | $O(p^3)$ |
| Parallelization | $O(K)$ / iter | $O(K^4)$ | $O(p^3)$ |

Summary

| | EE | $\mathcal{R}(\cdot)$ | θ | $\widehat{\theta}_n$ | $\mathcal{R}^*(\cdot)$ |
|----------|---------|------------------------------|----------------|---|--|
| | EE-sGGM | $\ \cdot\ _1$ | Ω | $[T_v(\widehat{\Sigma})]^{-1}$ | $\ \cdot\ _\infty$ |
| Task I | FASJEM | $\ \cdot\ _1 + \mathcal{R}'$ | Ω^{tot} | $inv[T_v(\widehat{\Sigma}^{tot})]$ | $\max(\ \cdot\ _\infty, \mathcal{R}'^*)$ |
| Task II | JEEK | kw-norm | Ω^{tot} | $inv[T_v(\widehat{\Sigma}^{tot})]$ | kw-dual |
| Task III | DIFFEE | $\ \cdot\ _1$ | Δ | $[T_v(\widehat{\Sigma}_d)]^{-1}$ $-[T_v(\widehat{\Sigma}_c)]^{-1}$ | $\ \cdot\ _\infty$ |

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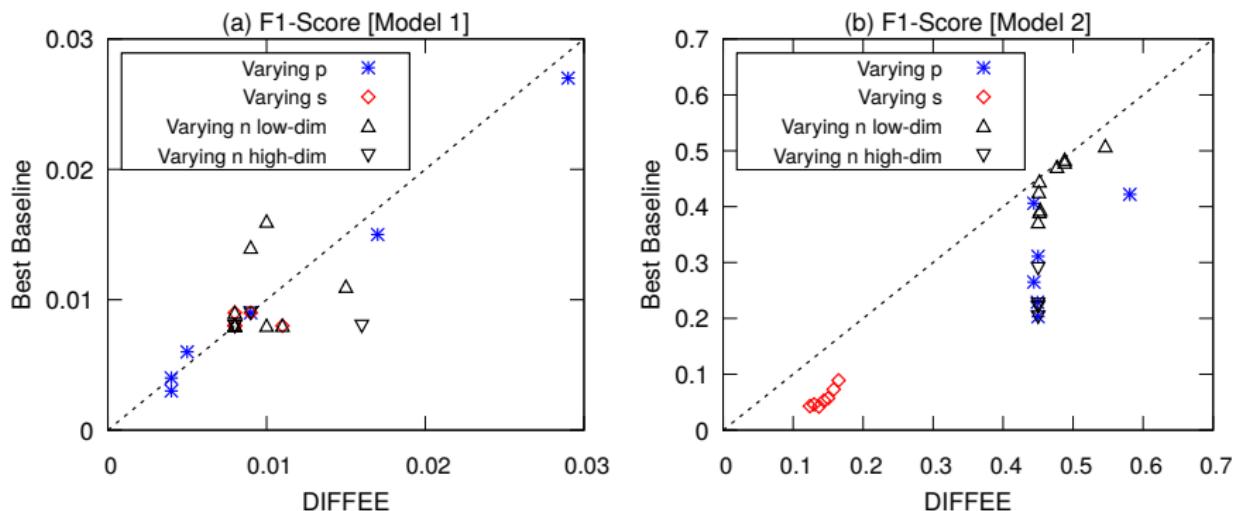
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Results: Theoretical Analysis

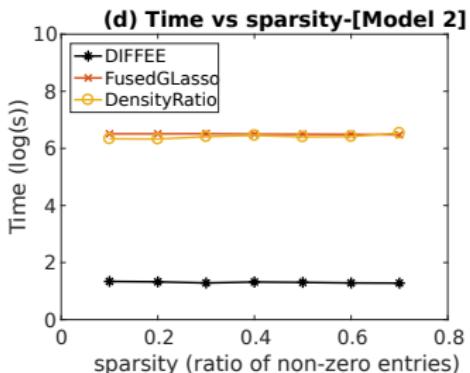
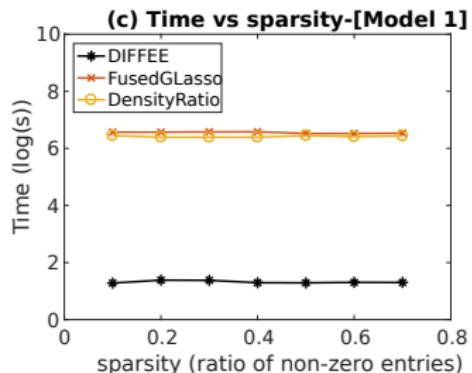
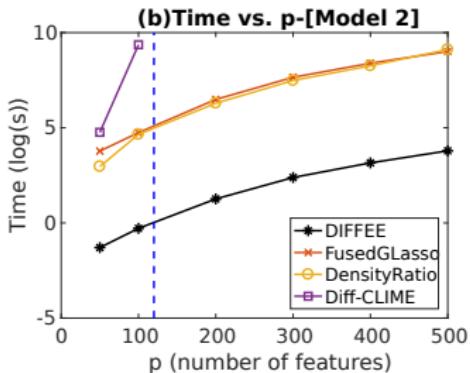
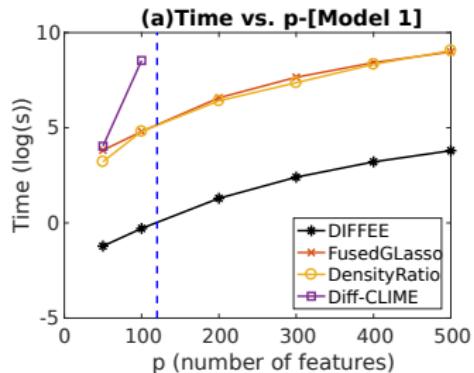
- Sharp convergence rate as the state-of-art

$$\begin{aligned} \|\hat{\Delta} - \Delta^*\|_\infty &\leq \frac{16\kappa_1 a}{\kappa_2} \sqrt{\frac{\log p}{\min(n_c, n_d)}} \\ \|\hat{\Delta} - \Delta^*\|_F &\leq \frac{32\kappa_1 a}{\kappa_2} \sqrt{\frac{k \log p}{\min(n_c, n_d)}} \\ \|\hat{\Delta} - \Delta^*\|_1 &\leq \frac{64\kappa_1 a}{\kappa_2} k \sqrt{\frac{\log p}{\min(n_c, n_d)}} \end{aligned} \tag{6.7}$$

Results: Synthetic Data Results



Results: Synthetic Data Results



Results: Real-world Data Results

- Apply to Brain image data (fMRI)
- Use the estimated different network in LDA
- Compare the accuracy with the state-of-art methods

| Method | DIFFEE | FusedGLasso | Diff-CLIME |
|--------------|---------------|-------------|------------|
| Accuracy (%) | 57.58% | 56.90% | 53.79% |

Discussion

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Support Analysis Results

- DIFFEE as an example

Lemma

$$\|\Delta^* - \mathcal{B}^*(\widehat{\Sigma}_d, \widehat{\Sigma}_c)\|_\infty \leq \lambda_n \quad (7.1)$$

Support Analysis Results

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Corollary

$$\Delta_{i,j}^* = 0 \implies |\mathcal{B}^*(\widehat{\Sigma}_d, \widehat{\Sigma}_c)_{i,j}| \leq \lambda_n \quad (7.2)$$



$$\widehat{\Delta} = \mathcal{S}_{\lambda_n}(\mathcal{B}^*(\widehat{\Sigma}_d, \widehat{\Sigma}_c)) \quad (7.3)$$

Support Analysis Results

- DIFFEE as an example

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$$\|\Delta^* - \mathcal{B}^*(\widehat{\Sigma}_d, \widehat{\Sigma}_c)\|_\infty \leq \lambda_n \quad (7.1)$$



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$$\widehat{\Delta} = \mathcal{S}_{\lambda_n}(\mathcal{B}^*(\widehat{\Sigma}_d, \widehat{\Sigma}_c)) \quad (7.3)$$

Result

$$\Delta_{i,j}^* = 0 \implies \widehat{\Delta}_{i,j} = 0 \quad (7.4)$$

- $\text{supp}(\widehat{\Delta}) \subseteq \text{supp}(\Delta^*)$

Support Analysis Result

- Additional Assumption:

Assumption

$$\min_{s \in supp(\Delta^*)} |\Delta_s^*| \geq 3 \|\Delta^* - \mathcal{B}^*(\widehat{\Sigma}_d, \widehat{\Sigma}_c)\|_\infty \quad (7.5)$$

Support Analysis Result

- Additional Assumption:

Assumption

$$\min_{s \in supp(\Delta^*)} |\Delta_s^*| \geq 3 \|\Delta^* - \mathcal{B}^*(\widehat{\Sigma}_d, \widehat{\Sigma}_c)\|_\infty \quad (7.5)$$

-

$$supp(\Delta^*) \subseteq supp(\widehat{\Delta}) \quad (7.6)$$

Support Analysis Result

- Additional Assumption:

Assumption

$$\min_{s \in supp(\Delta^*)} |\Delta_s^*| \geq 3 \|\Delta^* - \mathcal{B}^*(\widehat{\Sigma}_d, \widehat{\Sigma}_c)\|_\infty \quad (7.5)$$



$$supp(\Delta^*) \subseteq supp(\widehat{\Delta}) \quad (7.6)$$

- Combine the above results

$$supp(\Delta^*) = supp(\widehat{\Delta}) \quad (7.7)$$

Standardized Covariance Matrices

- Real world: Different tasks → different value scale
 - e.g., fMRI vs RNA sequencing
- Problem: hard to choose λ_n in different scales

Standardized Covariance Matrices

- Real world: Different tasks → different value scale
 - e.g., fMRI vs RNA sequencing
- Problem: hard to choose λ_n in different scales
- Solution: Covariance matrices \Rightarrow Correlation matrices

Theorem

The inverse of Correlation matrices have the same support set as the inverse of covariance matrices

- Nonparanormal extensions – Relax the Gaussian Assumption
- Added in all the packages

Iteration number T

- linearly converge method: $T = O(n \log(\frac{1}{TOL}))$
- TOL is the error bound

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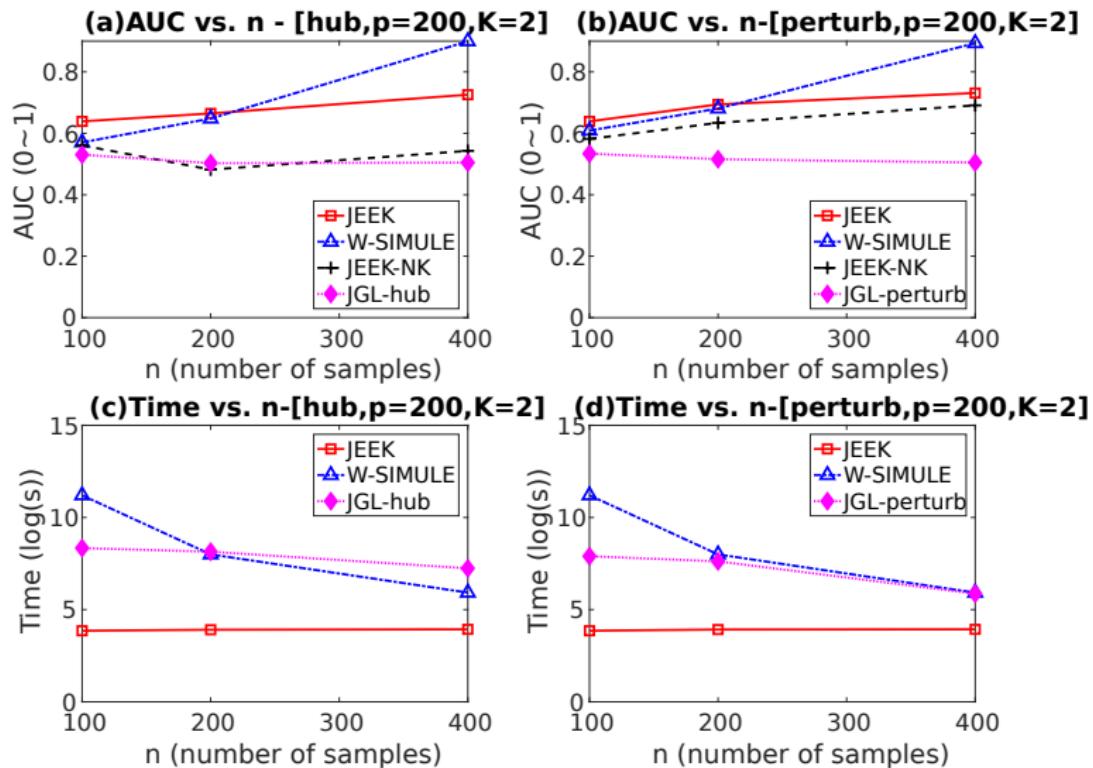
Iteration number T

- linearly converge method: $T = O(n \log(\frac{1}{TOL}))$
- TOL is the error bound
- FASJEM error bound: $O(\frac{\log(Kp)}{n_{tot}})$
- $T = O(\frac{n_{tot} \log(n_{tot})}{\log(\log(Kp))})$

Trade-off

- proxy backward mapping still $O(p^3)$
- In practice, fast in our three tasks
- Thanks to excellent low-level implementation
- Not well performed in low-dimensional case
- $p' = \max(n, p)$

Trade-off



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KW-norm for FASJEM

- Revise the ℓ_1 norm in FASJEM to a KW-norm

KW-norm for FASJEM

$$\begin{aligned}\mathcal{R}(\{\Omega^{(i)}\}) &= \sum_{i=1}^K ||W^{(i)} \circ \Omega^{(i)}||_1 \\ &= ||W^{tot} \circ \Omega^{tot}||_1\end{aligned}\tag{7.8}$$

- $\{W^{(i)}\}$: weights describing knowledge of each graph.

Future work: FASJEM with additional knowledge – FASJEM-K

FASJEM-K

$$\begin{aligned} & \underset{\Omega_{tot}}{\operatorname{argmin}} ||W_{tot} \circ \Omega_{tot}||_1 + \epsilon \mathcal{R}'(\Omega_{tot}) \\ & s.t. ||W_{tot} \circ (\Omega_{tot} - \text{inv}(T_v(\hat{\Sigma}_{tot})))||_\infty \leq \lambda_n \\ & \mathcal{R}'^*(\Omega_{tot} - \text{inv}(T_v(\hat{\Sigma}_{tot}))) \leq \epsilon \lambda_n \end{aligned} \tag{7.9}$$

KW-norm for Differential Network: kEV-norm

- Integrating both edge-level and node-level additional knowledge through a novel regularization function $\mathcal{R}(\cdot)$

kEV-norm

$$\mathcal{R}(\Delta) = ||W_E \circ \Delta_{E \setminus \mathcal{G}_V}||_1 + \epsilon ||\Delta_{\mathcal{G}_V}||_{\mathcal{G}_V,2} \quad (7.10)$$

- \mathcal{G}_V is a node group.
- W_E represents the weights for edges.

Future work: DIFFEE-K

- Combine kEV-norm and Elementary Estimator

DIFFEE-K

$$\underset{\Delta}{\operatorname{argmin}} \|\mathcal{W}_E \circ \Delta_{E \setminus \mathcal{G}_V}\|_1 + \epsilon \|\Delta_{\mathcal{G}_V}\|_{\mathcal{G}_V,2}$$

Subject to: $\|\mathcal{W}_E \circ \left(\Delta - \left([T_V(\widehat{\Sigma}_d)]^{-1} - [T_V(\widehat{\Sigma}_c)]^{-1} \right) \right)\|_{\infty} \leq \lambda_n \quad (7.11)$

$$\epsilon \|\Delta - \left([T_V(\widehat{\Sigma}_d)]^{-1} - [T_V(\widehat{\Sigma}_c)]^{-1} \right)\|_{\mathcal{G}_V,2}^* \leq \lambda_n$$

Publications

- FASJEM
 - A Fast and Scalable Joint Estimator for Learning Multiple Related Sparse Gaussian Graphical Models, **B Wang, J Gao, Y Qi, AISTATS 2017**
- DIFFEE
 - Fast and Scalable Learning of Sparse Changes in High-Dimensional Gaussian Graphical Model Structure, **B Wang, A Sekhon, Y Qi, AISTATS 2018**
- W-SIMULE
 - A constrained ℓ_1 minimization approach for estimating multiple sparse Gaussian or nonparanormal graphical models, **B Wang, R Singh, Y Qi, Machine Learning 106 (9-10), 1381-1417**
 - A Constrained, Weighted-L1 Minimization Approach for Joint Discovery of Heterogeneous Neural Connectivity Graphs, **C Singh, B Wang, Y Qi, Advances in Modeling and Learning Interactions from Complex Data, NIPS 2017 Workshop**

Publications

- JEEK
 - A Fast and Scalable Joint Estimator for Integrating Additional Knowledge in Learning Multiple Related Sparse Gaussian Graphical Models, B Wang, A Sekhon, Y Qi, ICML 2018
- DIFFEE-K
 - A Fast and Scalable Estimator for Using Additional Knowledge in Learning Sparse Structure Change of High-Dimensional Gaussian Graphical Models, B Wang, A Sekhon, Y Qi, submit to NIPS 2018

R Package is Available !!!

- The project website: <http://jointggm.org/>
- R package "simule":
 - `install.packages("simule")`
 - `demo(simule) !`
- R package "fasjem":
 - `install.packages("fasjem")`
 - `demo(fasjem) !`
- R package "diffee":
 - `install.packages("diffee")`
 - `demo(diffee) !`
- R package "jeek":
 - `install.packages("jeek")`
 - `demo(jeek) !`
- A complete package "jointNet" will be ready by this summer.

Acknowledgement

- Advisor: Yanjun Qi
- Co-authors: Rita, Arshdeep, Ji, Chandan
- Lab mates: Zhaoyang, Jack, Weilin
- My Family
- Thanks!

Back-up: Difficulty in combining FASJEM and JEEK

$$\operatorname{argmin}_{\Omega_I^{tot}, \Omega_S^{tot}} ||W_I^{tot} \circ \Omega_I^{tot}||_1 + ||W_S^{tot} \circ \Omega_S^{tot}|| + \epsilon \mathcal{R}'(\Omega^{tot})$$

Subject to: $||W_I^{tot} \circ (\Omega^{tot} - \text{inv}(T_V(\widehat{\Sigma}^{tot})))||_\infty \leq \lambda_n$ (7.12)

$$||W_S^{tot} \circ (\Omega^{tot} - \text{inv}(T_V(\widehat{\Sigma}^{tot})))||_\infty \leq \lambda_n$$

$$\mathcal{R}^{*' }(\Omega^{tot}) \leq \epsilon \lambda_n$$

- Hard to optimize
- Lose fast and scalable property

Back-up: How to choose v in $T_v(\widehat{\Sigma})$

- line search
- v from the set $\{0.001i | i = 1, 2, \dots, 1000\}$
- pick a value that makes $T_v(\widehat{\Sigma})$ and be invertible

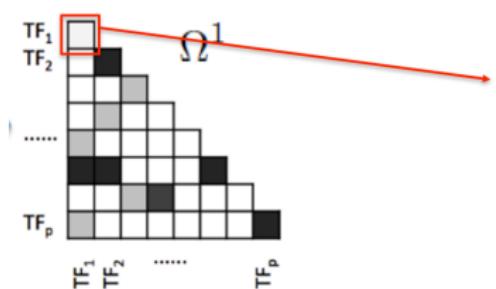
Back-up: Connecting to Bayesian Statistics

$$\begin{aligned} & -\log(\mathbb{P}(\Omega^{(i)} | X^{(i)}, \mu^{(i)}, W_{I,j,k}^{(i)}, W_{Sj,k})) \\ & \propto -\log(\det(\Omega^{(i)-1})) + \langle \Omega^{(i)}, \widehat{\Sigma}^{(i)} \rangle \\ & + \sum_{j,k} (W_{I,j,k}^{(i)} |\Omega_{I,j,k}^{(i)}| + W_S |\Omega_{Sj,k}|) \end{aligned} \tag{7.13}$$

Back-up: Proximal algorithm Basics

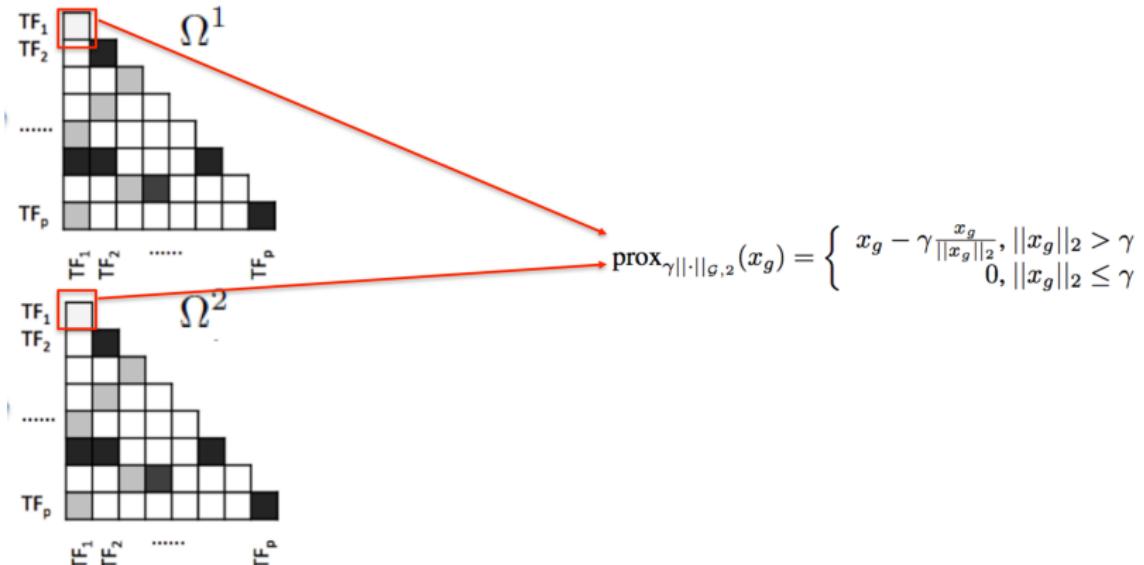
- proximity definition:
- $\text{prox}_h(x) = \underset{u}{\operatorname{argmin}}(h(u) + \frac{1}{2}||u - x||_2^2)$
- $\underset{x}{\operatorname{argmin}} f(x) = g(x) + h(x)$
- proximal gradient descent:
- $x^{(k)} = \text{prox}_{t_k h}(x^{(k-1)} - t_k \triangledown g(x^{(k-1)}))$

Back-up: Proximal algorithm for FASJEM

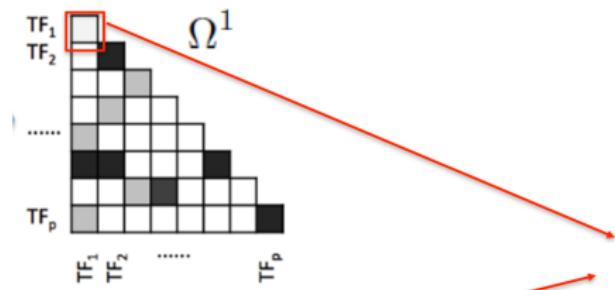


$$(\text{prox}_{\gamma \|\cdot\|_1}(x))_i = \begin{cases} x_i - \gamma, & x_i > \gamma \\ 0, & |x_i| \leq \gamma \\ x_i + \gamma, & x_i < -\gamma \end{cases}$$

Back-up: Proximal algorithm for FASJEM



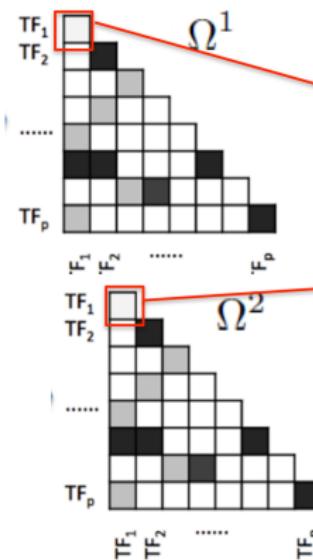
Back-up: Proximal algorithm for FASJEM



$$\text{proj}_{||x-a||_\infty \leq \lambda} = \begin{cases} x_i, & |x_i - a_i| \leq \lambda \\ a_i + \lambda, & x_i > a_i + \lambda \\ a_i - \lambda, & x_i < a_i - \lambda \end{cases}$$

$$\mathcal{B}^{*(1)} := \begin{pmatrix} b_{11}^{(1)} & b_{12}^{(1)} & \cdots & b_{1p}^{(1)} \\ b_{21}^{(1)} & b_{22}^{(1)} & \cdots & b_{2p}^{(1)} \\ \vdots & \vdots & \ddots & \vdots \\ b_{p1}^{(1)} & b_{p2}^{(1)} & \cdots & b_{pp}^{(1)} \end{pmatrix}$$

Back-up: Proximal algorithm for FASJEM



$$\mathcal{B}^{*(1)} := \begin{pmatrix} b_{11}^{(1)} & b_{12}^{(1)} & \dots & b_{1p}^{(1)} \\ b_{21}^{(1)} & b_{22}^{(1)} & \dots & b_{2p}^{(1)} \\ \vdots & \vdots & \ddots & \vdots \\ b_{p1}^{(1)} & b_{p2}^{(1)} & \dots & b_{pp}^{(1)} \end{pmatrix}$$

$$\text{prox}_{\gamma g_2}(x) = \text{proj}_{||x-a||_{G,2}^*, \leq \lambda}$$

$$= \begin{cases} x_g, & ||x_g - a_g||_2 \leq \lambda \\ \lambda \frac{x_g - a_g}{||x_g - a_g||_2} + a_g, & ||x_g - a_g||_2 > \lambda \end{cases}$$

$$\mathcal{B}^{*(2)} := \begin{pmatrix} b_{11}^{(2)} & b_{12}^{(2)} & \dots & b_{1p}^{(2)} \\ b_{21}^{(2)} & b_{22}^{(2)} & \dots & b_{2p}^{(2)} \\ \vdots & \vdots & \ddots & \vdots \\ b_{p1}^{(2)} & b_{p2}^{(2)} & \dots & b_{pp}^{(2)} \end{pmatrix}$$