

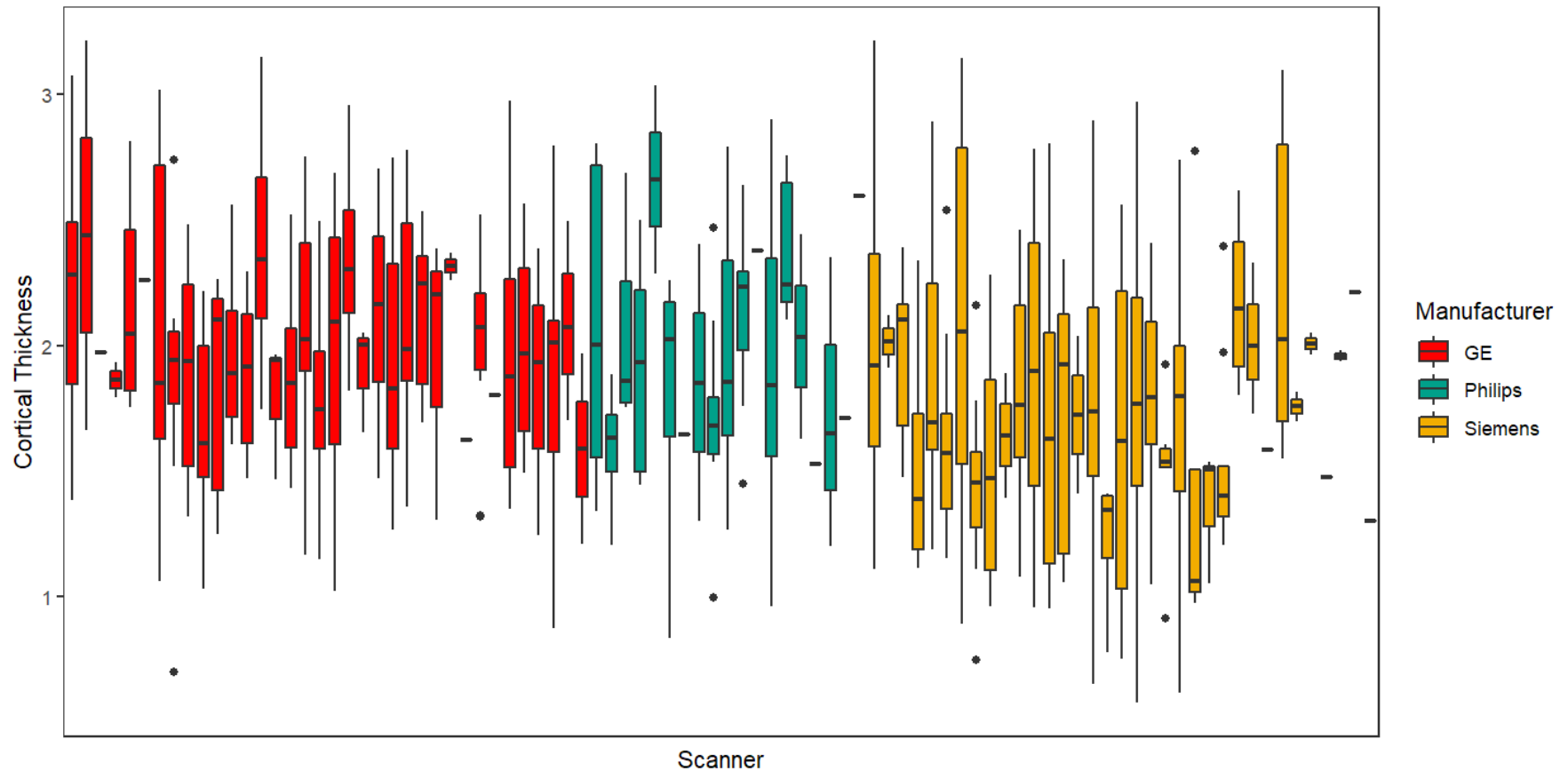


Removal of Scanner Effects in Covariance

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Heterogeneity across Scanners



Current Harmonization Method

Step 1: Assume site effect exists in mean and variance of observations

$$y_{ijv} = \alpha_v + \mathbf{x}'_{ij}\beta_v + \gamma_{iv} + \delta_{iv}e_{ijv}$$

Step 2: Obtain empirical Bayes point estimates by imposing a common prior across features for both mean and variance site effect

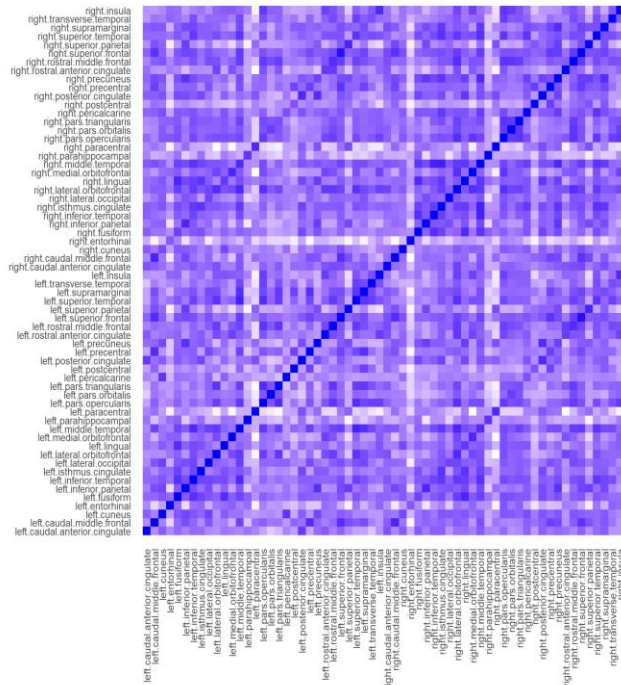
Step 3: Remove site effect while retaining covariate effect

$$y_{ijv}^{ComBat} = \frac{y_{ijv} - \hat{\alpha}_v - \mathbf{x}'_{ij}\hat{\beta}_v - \gamma_{iv}^*}{\delta_{iv}^*} + \hat{\alpha}_v + \mathbf{x}'_{ij}\hat{\beta}_v$$

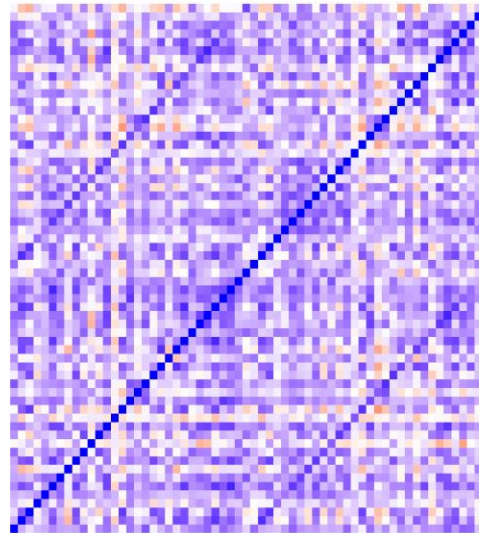


Covariance Differences across Sites

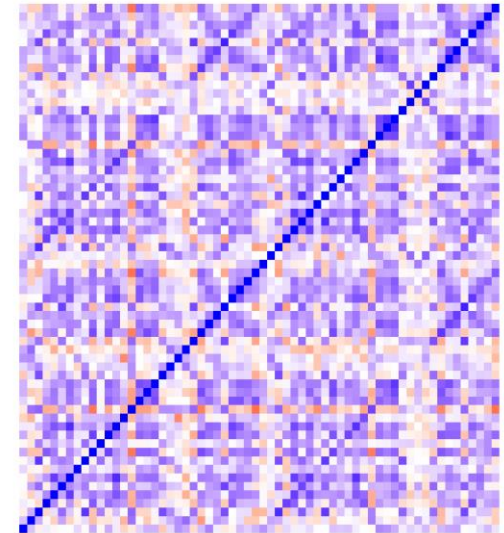
Site A (Siemens)



Site B (GE)



Site B (GE)



Proposed Harmonization: CovBat

Step 1: Apply ComBat to harmonize mean and variance across sites, then residualize on intercept and covariates

Step 2: Perform principal component analysis on full data to obtain

$$\Sigma_i = \sum_{k=1}^q \lambda_{ik} \phi_k \phi_k' \quad e_{ij}^{ComBat} = \sum_{k=1}^q \xi_{ijk} \phi_k$$

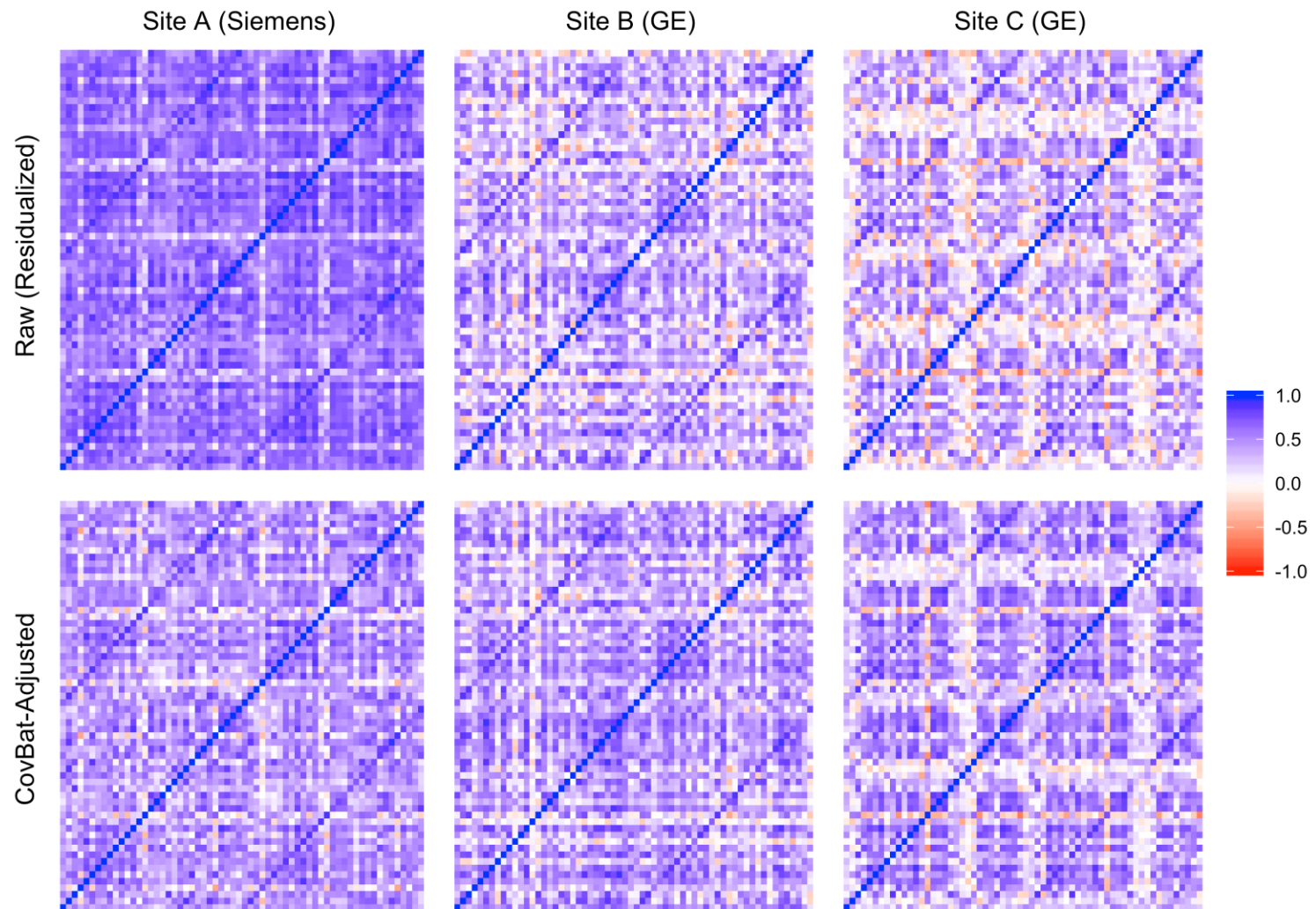
Step 3: Harmonize mean and variance of first K PC scores and reintroduce intercept and covariates

$$e_{ij}^{CovBat} = \sum_{k=1}^K \xi_{ijk}^{CovBat} \phi_k + \sum_{l=K+1}^q \xi_{ijl} \phi_l$$

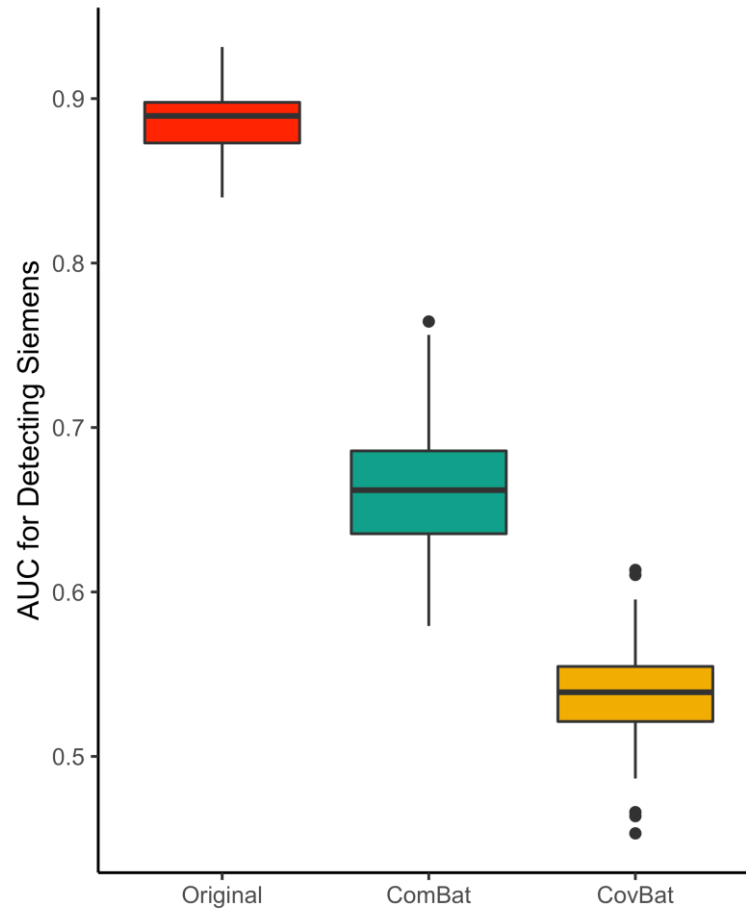
$$y_{ijv}^{CovBat} = e_{ijv}^{CovBat} + \hat{\alpha}_v + \mathbf{x}_{ij}' \hat{\beta}_v$$



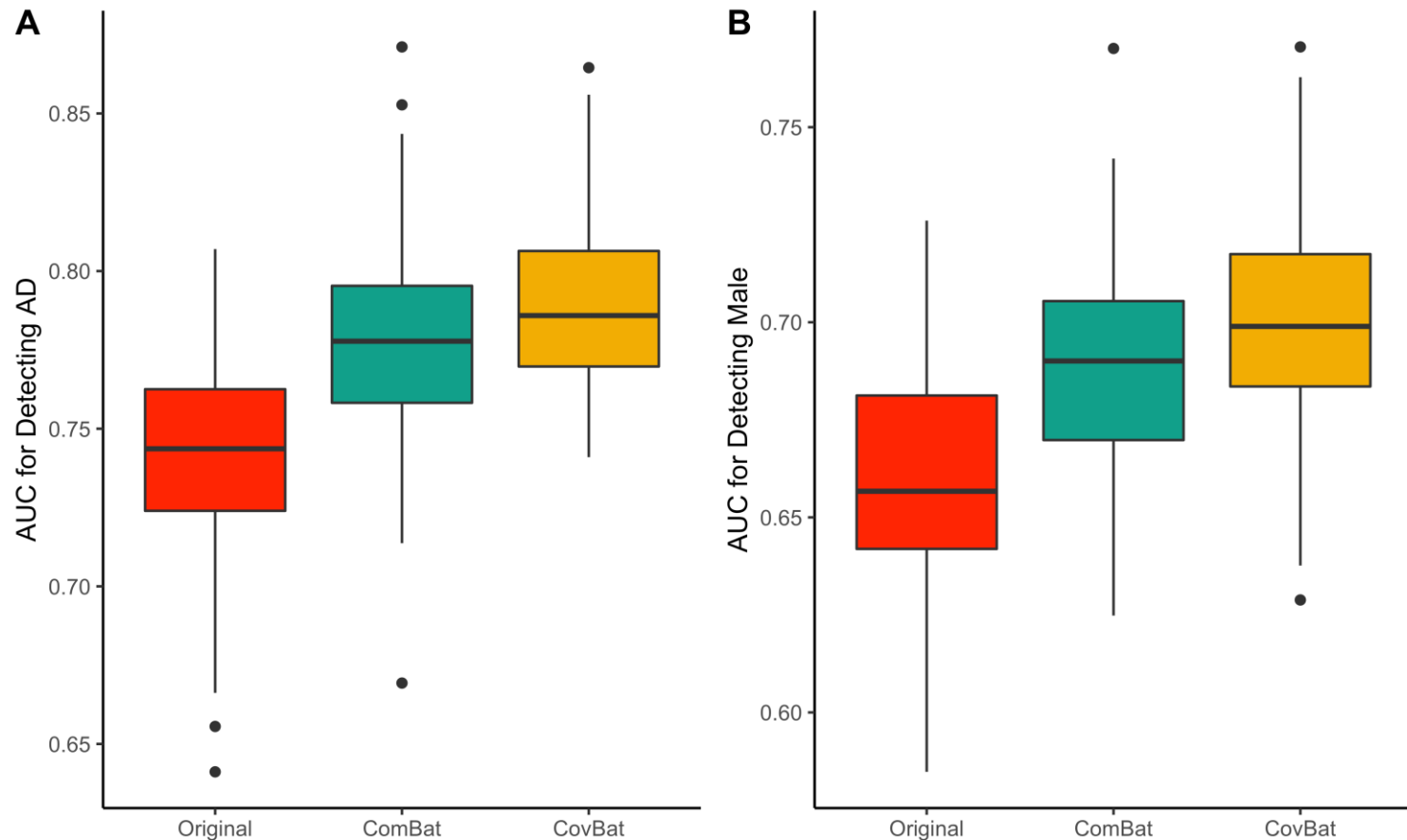
CovBat Harmonizes Covariance



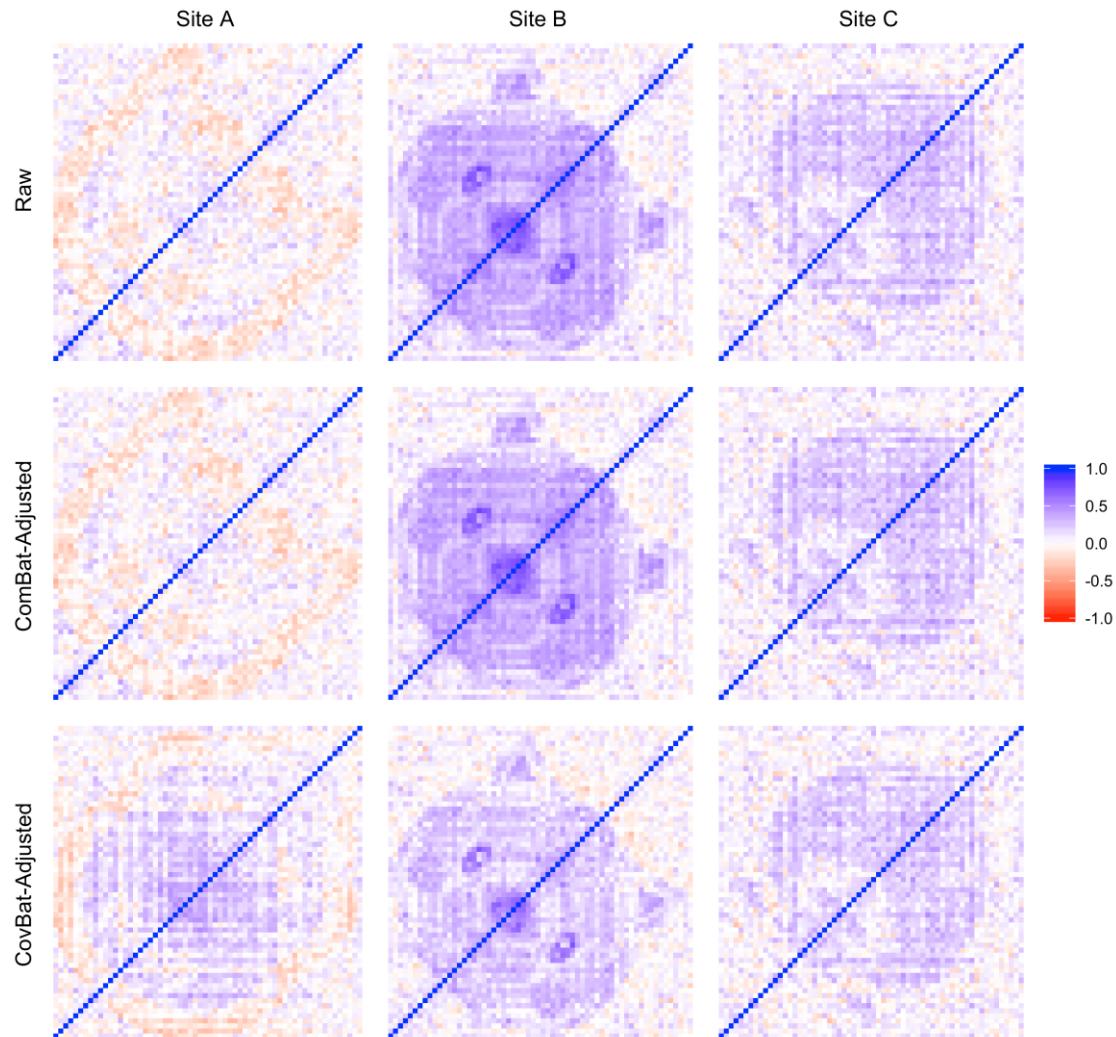
Obscures Detection of Scanner



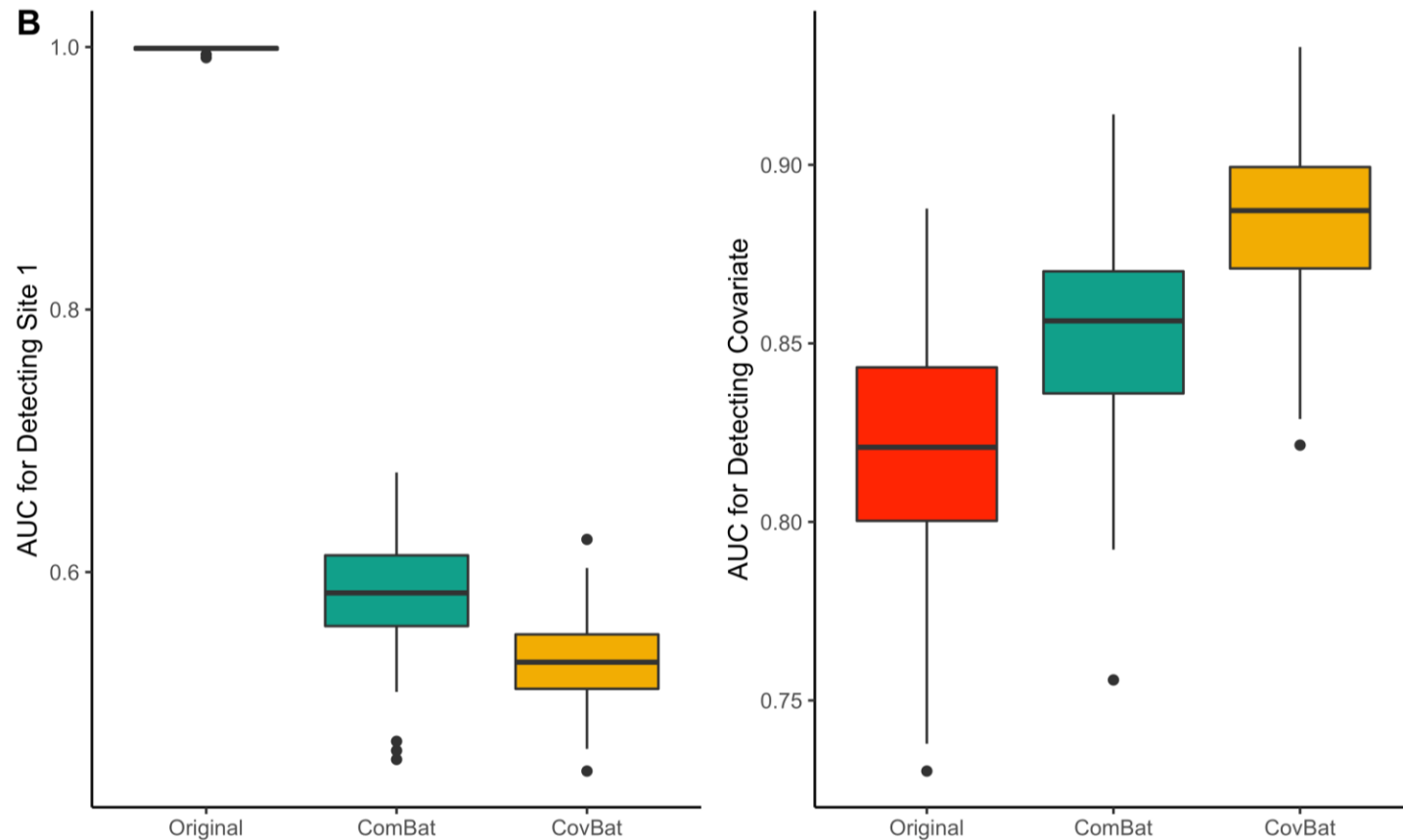
Improves Detection of Covariates



Simulation Results



Simulation Results



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References

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