Consider $W_{ij} = Z_i^{\mathrm{T}} \beta + X_i + U_{ij}$. where i = 1, ..., n and $j = 1, ..., m_j$. Assume:

- $Z_i \sim N(\mu_z, \sigma_z^2 \mathbb{I})$
- $U_{ij} \sim N\{0, \sigma_U^2(Z_i)\}$ where $\sigma^2(\cdot)$ is an unknown smooth function
- $Cor(U_{ij}, U_{ik}) = 0$ for all $j \neq k$ (I think this assumption is reasonable)
- $X_i \sim N(0, \sigma_x^2)$

We have

$$Var(W_{ij}) = \beta^{\mathrm{T}} \sigma_z \beta + \sigma_x^2 + \sigma_u^2(Z_i) + \beta^{\mathrm{T}} \Sigma_{ZU} \beta + \beta^{\mathrm{T}} \Sigma_{ZX} \beta.$$
 (1)

If we can estimate all the other components, we can solve for $\hat{\sigma}_x^2$. If X and Z are uncorrelated, we have one less term to worry about since $\Sigma_{ZX} = 0$. This seems like a reasonable assumption to me. Is it possible to assume $\Sigma_{ZU} = 0$? I don't think Z and U can be independent if the variance of U is a function of Z.

Consider the replication differences, $W_{ij} - W_{ik} = U_{ij} - U_{ik}$, and note $Var(W_{ij} - W_{ik}) = Var(U_{ij}) + Var(U_{ik}) - 2Cor(U_{ij}, U_{ij}) = Var(U_{ij}) + Var(U_{ik}) = 2\sigma^2(Z_i)$. Then $1/\sqrt{2}(W_{ij} - W_{ik}) \sim N\{0, \sigma(Z_i)\}$ Now consider all permutations of the replication differences, the set $\{W_{ij} - W_{ik} | k \neq j\}$. This gives us $m_j(m_j - 1)$ observations for each Z_i to model the variance function, $\sigma_U^2(Z_i)$. Denote these differences as $D_{i\ell}$, for $\ell = 1, \ldots, m_j(m_j - 1)$ and $i = 1, \ldots, n$.

? wrote a paper about variance estimation, and I'm stealing their general idea. I doubt the asymptotics hold. The paper is technical and I haven't gone through it in detail. I think we can use his idea though. You and Marie Davidian wrote a similar, though less general, paper: ?

Regress $D_{i\ell} = m(Z_i) + \epsilon_i$ where $m(\cdot)$ is a specified or unspecified mean function which should hopefully be estimated very close to 0 and ϵ_i are iid $N\{0, \sigma^2(Z_i)\}$. Take the residuals r_i and regression them against Z_i to get $\widehat{\sigma}^2(\cdot)$. That is $r_i = \sigma^2(Z_i) + \epsilon_i$. I think this will work with $Var(U_{ij}) = \sigma^2(X_i, Z_i)$ as well. We can fit $\widehat{\sigma}^2(\cdot)$ with multivariate spline model.

<u>The good</u>: I am actually able to get a decent estimate of σ_x^2 . I am able to get reasonable estimates of $\hat{\sigma}^2(\cdot)$, though there are problems at the tails of the data. We can estimate

everything else in (1) with sample variance or covariance. All our estimates are \sqrt{n} consistent except $\hat{\sigma}^2(\cdot)$, though we can get \sqrt{n} consistency if we use a parametric model

<u>The bad</u>: The feels very ad-hoc. Maybe we can see this analogous to calculating the reliability ratio for regression coefficients. I think we'll have a hard time developing any theory for this method, except demonstrating consistency. I have no idea how to include uncertainty in the estimate of σ_x^2

<u>Next?</u>: Maybe you have a different idea for how to solve this. Maybe a likelihood based method?

1 Illustrate Point

What if we maximize the sum of squares and find the worst fitting line?

$$max(\boldsymbol{\beta}) ||Y - \boldsymbol{\beta}X||$$