

Supplementary Methods

Modeling expression ranks for noise-tolerant differential expression analysis of scRNA-seq data

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Derivation of the Fisher Information Matrix

Note that, for our DGBD model with likelihood function given by Eq (2) of the main paper, we have

$$\begin{aligned}\frac{\partial^2 \log L}{\partial a^2} &= \left(\sum_{r=1}^{r=N} y_r \right) \frac{\partial^2 \log(A)}{\partial a^2} \\ \frac{\partial^2 \log L}{\partial b^2} &= \left(\sum_{r=1}^{r=N} y_r \right) \frac{\partial^2 \log(A)}{\partial b^2} \\ \frac{\partial^2 \log L}{\partial a \partial b} &= \left(\sum_{r=1}^{r=N} y_r \right) \frac{\partial^2 \log(A)}{\partial a \partial b}\end{aligned}\tag{1}$$

So in order to evaluate the above mentioned double derivatives, the first order derivative $\frac{\partial \log A}{\partial a}$ and $\frac{\partial \log A}{\partial b}$ are determined as follows:

$$\begin{aligned}\log A &= -\log \left(\sum_{r=1}^{r=N} \frac{(N+1-r)^b}{r^a} \right) \\ \frac{\partial \log A}{\partial a} &= \frac{1}{\left(\sum_{r=1}^{r=N} \frac{(N+1-r)^b}{r^a} \right)} \times \sum_{r=1}^{r=N} \frac{(N+1-r)^b \log r}{r^a} \\ \frac{\partial \log A}{\partial b} &= \frac{-1}{\left(\sum_{r=1}^{r=N} \frac{(N+1-r)^b}{r^a} \right)} \times \sum_{r=1}^{r=N} \frac{(N+1-r)^b \log(N+1-r)}{r^a}\end{aligned}\tag{2}$$

Re-writing Equation 2 in a more succinct form in the Equation 3 below, we get

$$\frac{\partial \log A}{\partial a} = \frac{u_{1,0}}{u_{0,0}} \quad \text{and} \quad \frac{\partial \log A}{\partial b} = -\frac{u_{0,1}}{u_{0,0}}\tag{3}$$

where $u_{i,j}$ s are as defined in the main paper. Evaluating the partial derivatives of $u_{1,0}$, $u_{0,0}$ and $u_{0,1}$ with

respect to a and b , in the Equation 4:

$$\begin{aligned}
\frac{\partial u_{1,0}}{\partial a} &= -\sum_{r=1}^{r=N} \frac{(N+1-r)^b (\log r)^2}{r^a} \\
\frac{\partial u_{1,0}}{\partial b} &= \sum_{r=1}^{r=N} \frac{(N+1-r)^b [\log r] [\log (N+1-r)]}{r^a} \\
\frac{\partial u_{0,0}}{\partial a} &= -\sum_{r=1}^{r=N} \frac{(N+1-r)^b \log r}{r^a} \\
\frac{\partial u_{0,0}}{\partial b} &= \sum_{r=1}^{r=N} \frac{(N+1-r)^b \log (N+1-r)}{r^a} \\
\frac{\partial u_{0,1}}{\partial a} &= \sum_{r=1}^{r=N} \frac{(N+1-r)^b [\log r] [\log (N+1-r)]}{r^a} \\
\frac{\partial u_{0,1}}{\partial b} &= -\sum_{r=1}^{r=N} \frac{(N+1-r)^b [\log (N+1-r)]^2}{r^a}
\end{aligned} \tag{4}$$

In a compact form, these can be written more generally, for any $i, j = 0, 1$, as

$$\frac{\partial u_{i,j}}{\partial a} = -u_{i+1,j}, \quad \frac{\partial u_{i,j}}{\partial b} = u_{i,j+1}. \tag{5}$$

Substituting the above expressions in the formula for Fisher information matrix in Eq (3) of the main paper, we get its simplified form for computation within our ROSeq.