

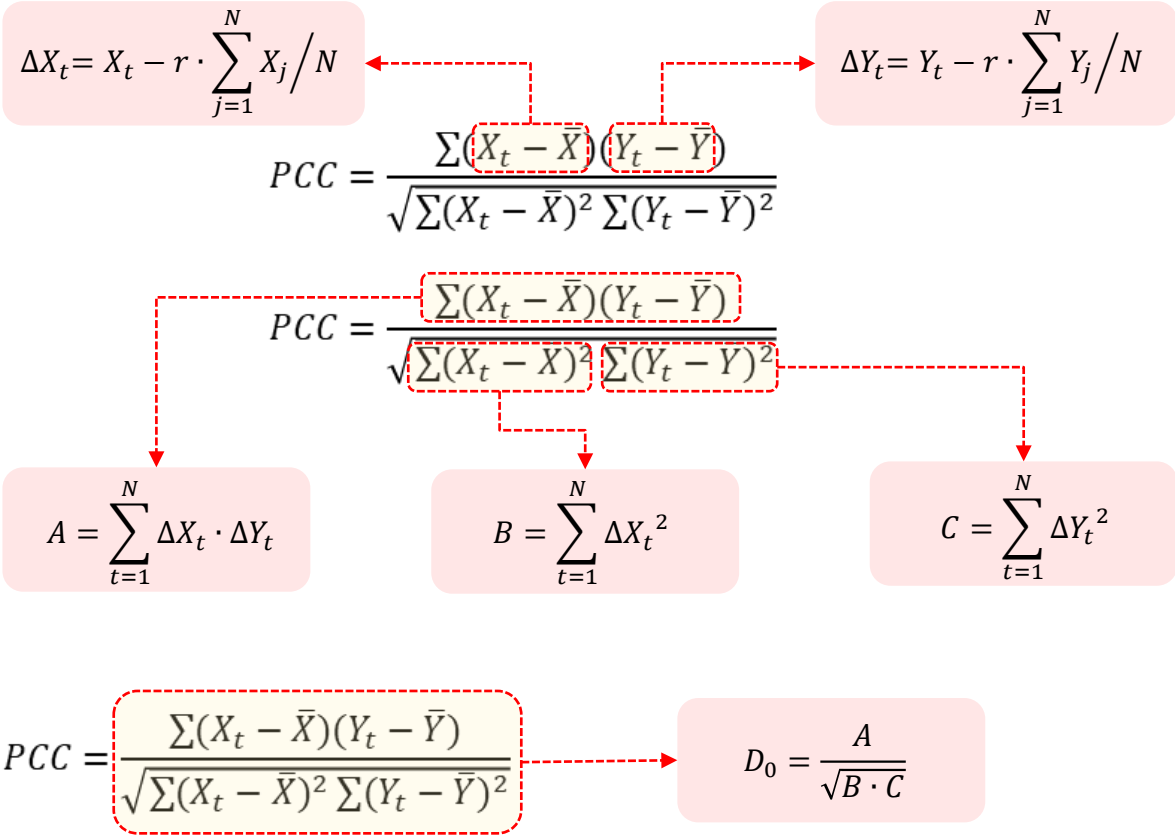
Supp. Figures

Supplementary Figure 1

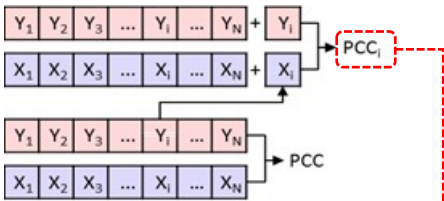
1. The formula of Pearson Correlation Coefficient (PCC):

$$PCC = \frac{\sum(X_t - \bar{X})(Y_t - \bar{Y})}{\sqrt{\sum(X_t - \bar{X})^2 \sum(Y_t - \bar{Y})^2}}$$

2. When r = 1 :



3. When adding Xi and Yi to X list and Y list, respectively :

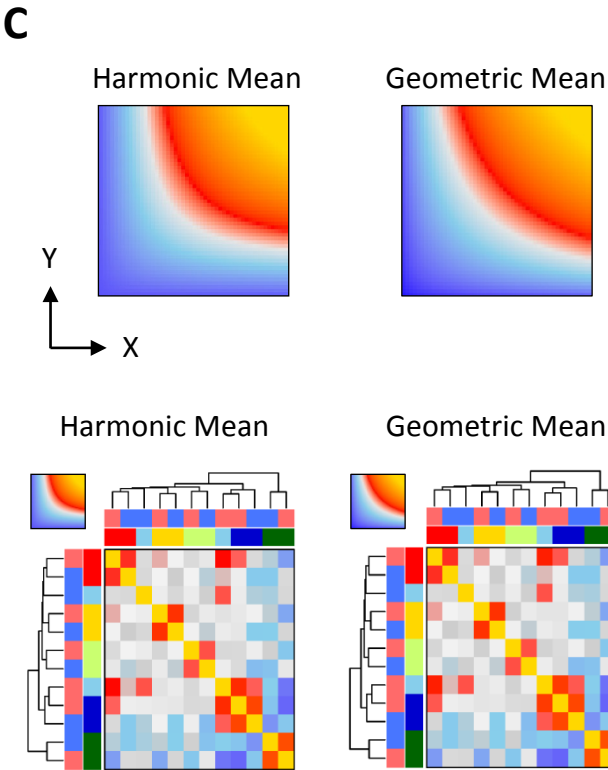
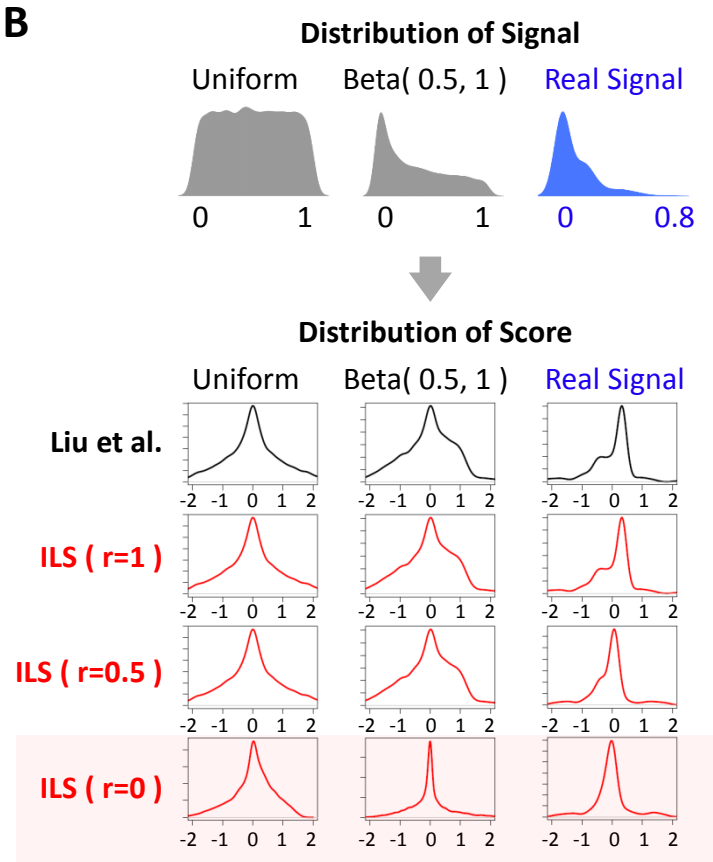
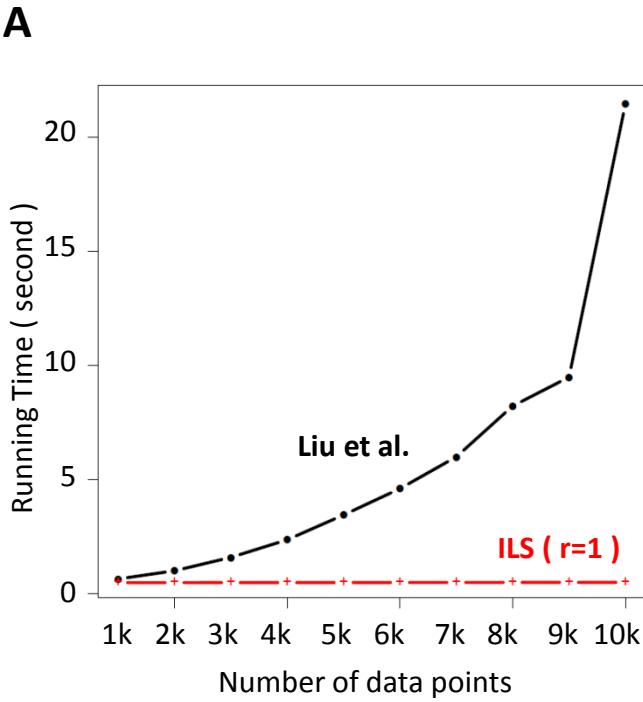


$$PCC_{new} = \frac{[\sum(X_t - \bar{X}_{new})(Y_t - \bar{Y}_{new})] + (X_i - \bar{X}_{new})(Y_i - \bar{Y}_{new})}{\sqrt{[\sum(X_t - \bar{X}_{new})^2 + (X_i - \bar{X}_{new})^2][\sum(Y_t - \bar{Y}_{new})^2 + (Y_i - \bar{Y}_{new})^2]}}$$

4. Because $\bar{X}_{new} \sim \bar{X}$ and $\bar{Y}_{new} \sim \bar{Y}$:

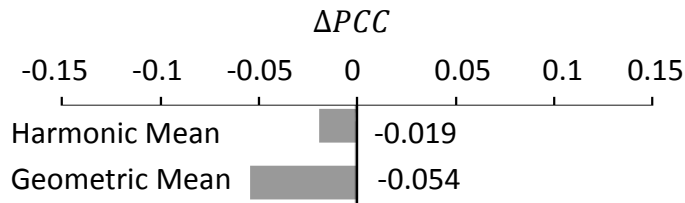
$$PCC_{new} \sim D_i = \frac{A + \Delta X_i \cdot \Delta Y_i}{\sqrt{(B + \Delta X_i^2) \cdot (C + \Delta Y_i^2)}}$$

Supplementary Figure 2



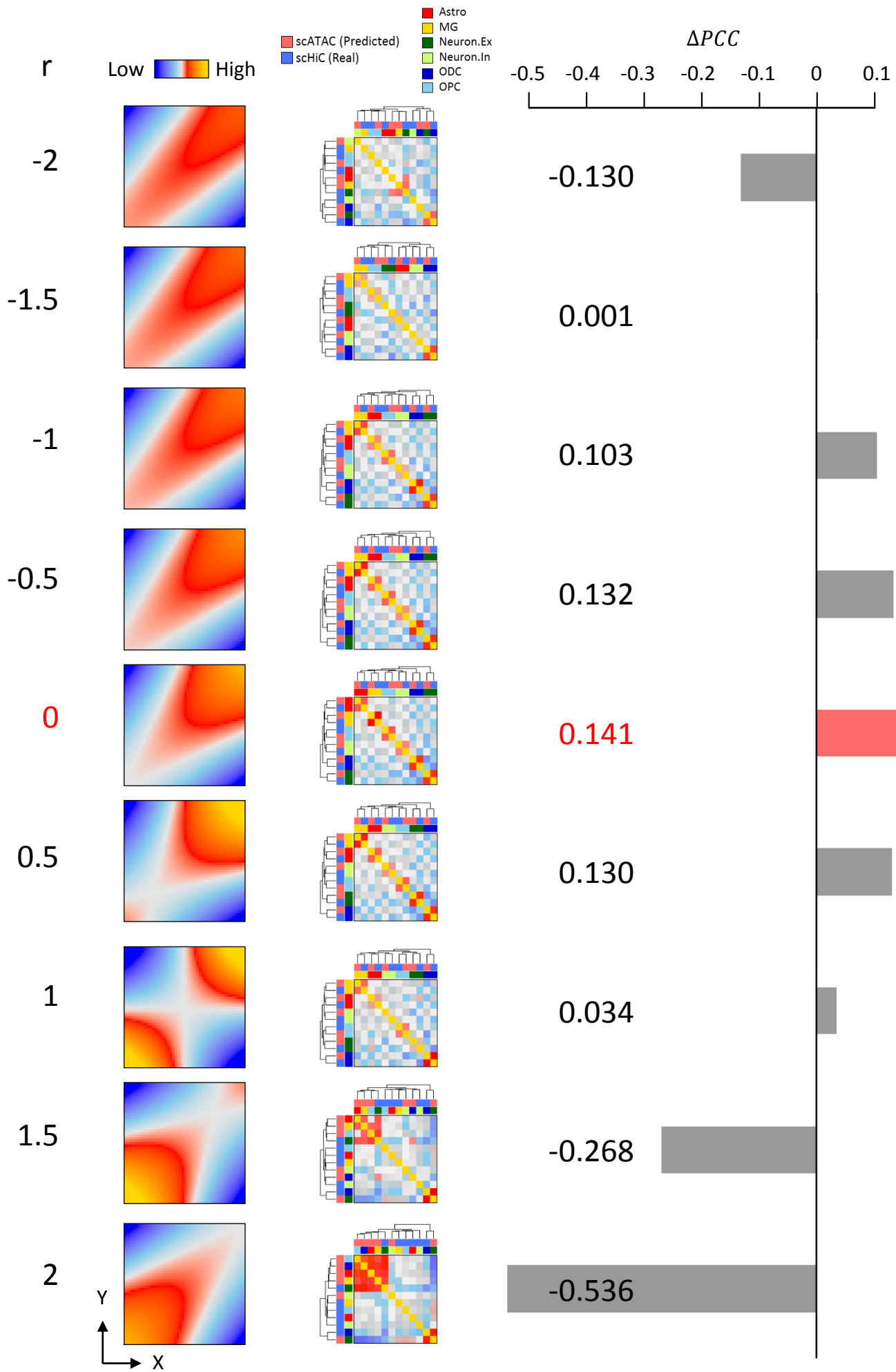
$$\text{Harmonic Mean} = \frac{2 \cdot X \cdot Y}{X + Y + 1}$$

$$\text{Geometric Mean} = \sqrt{X \cdot Y}$$



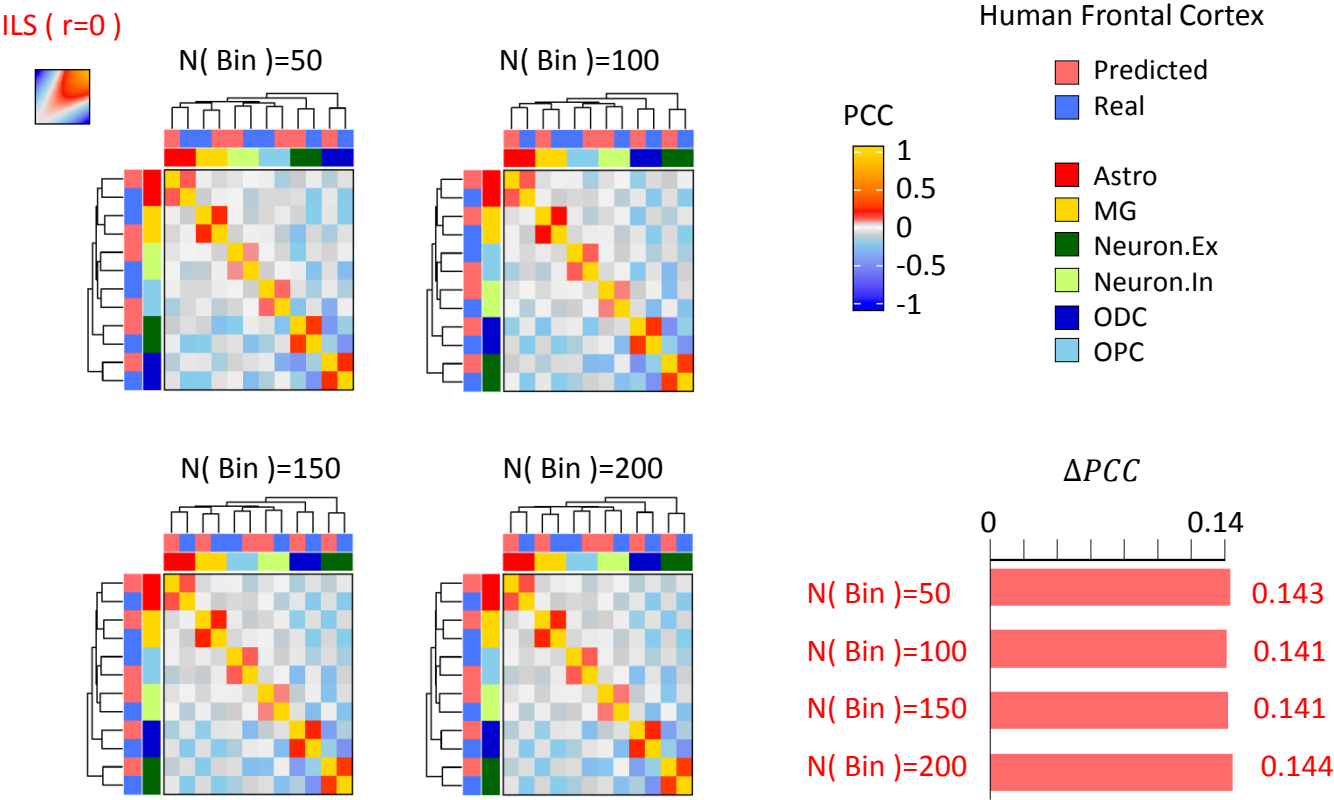
Supplementary Figure 3

Human Frontal Cortex



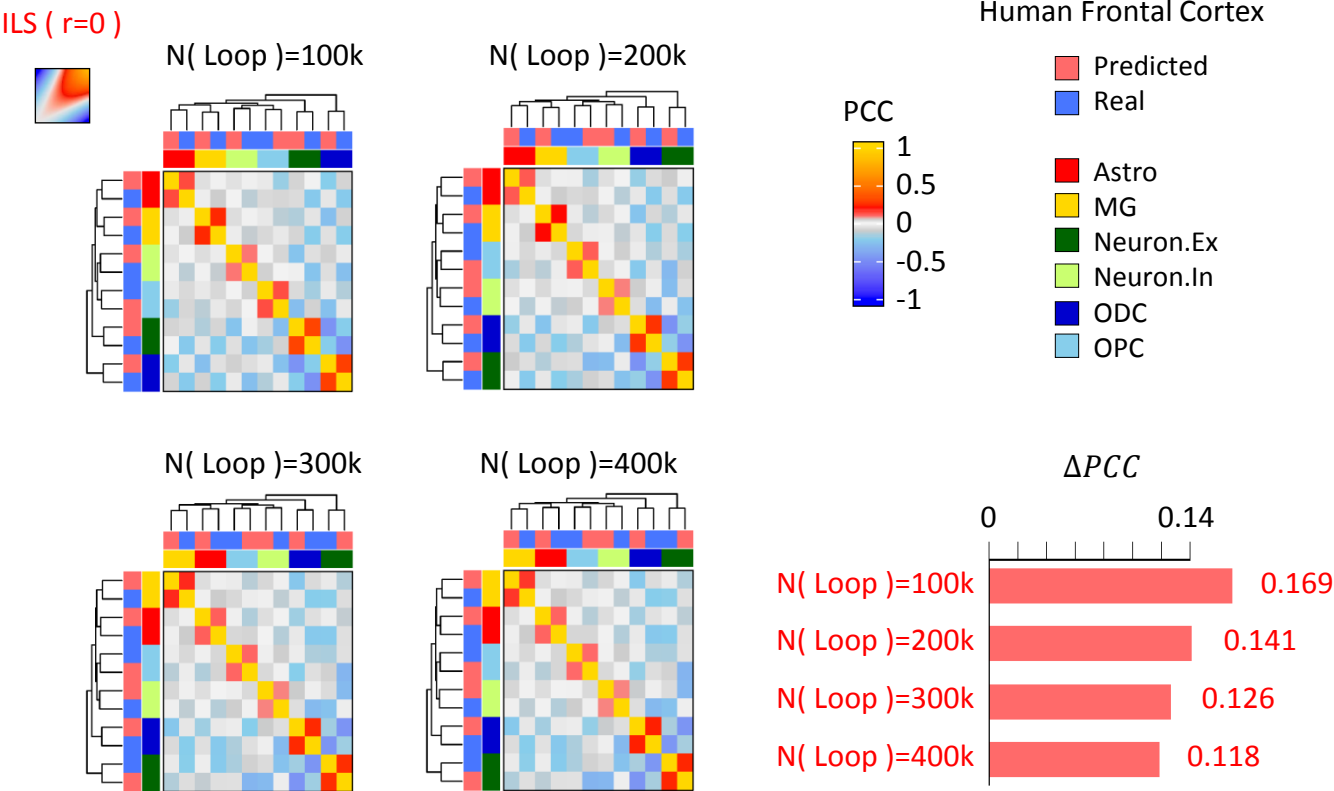
A

Using different number of bins, N(Loop)=200k

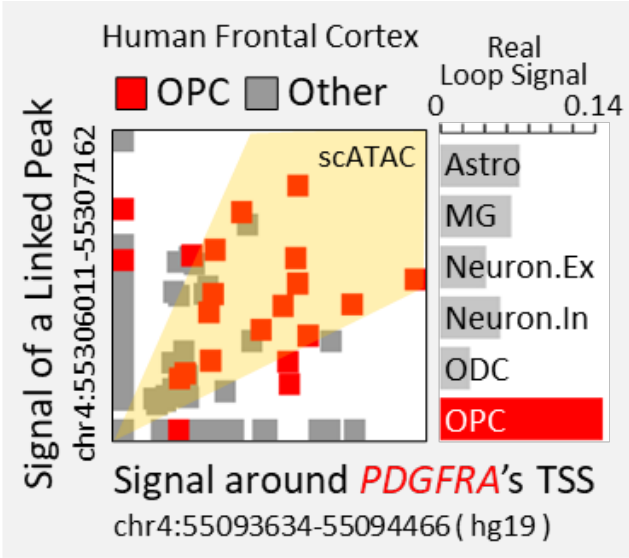
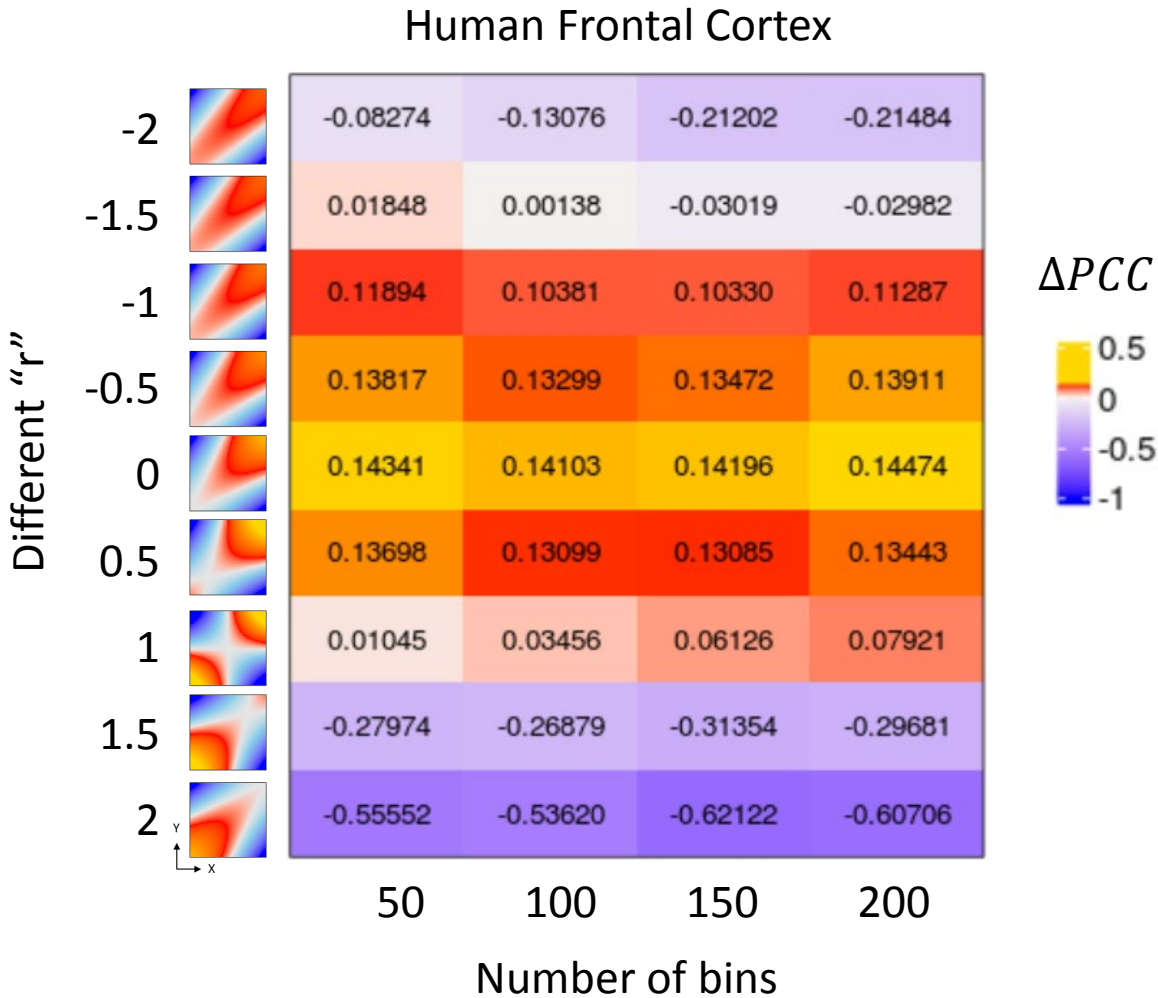


B

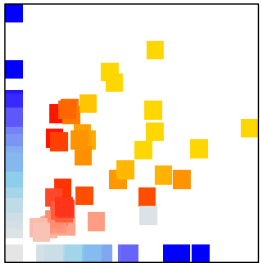
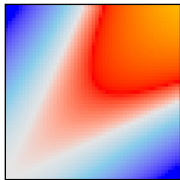
Using different number of top loops of Cicero, N(Bin)=100



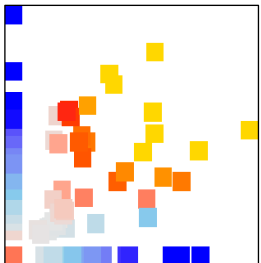
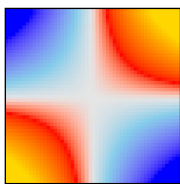
Number of bins & different “r”



r = 0



r = 1



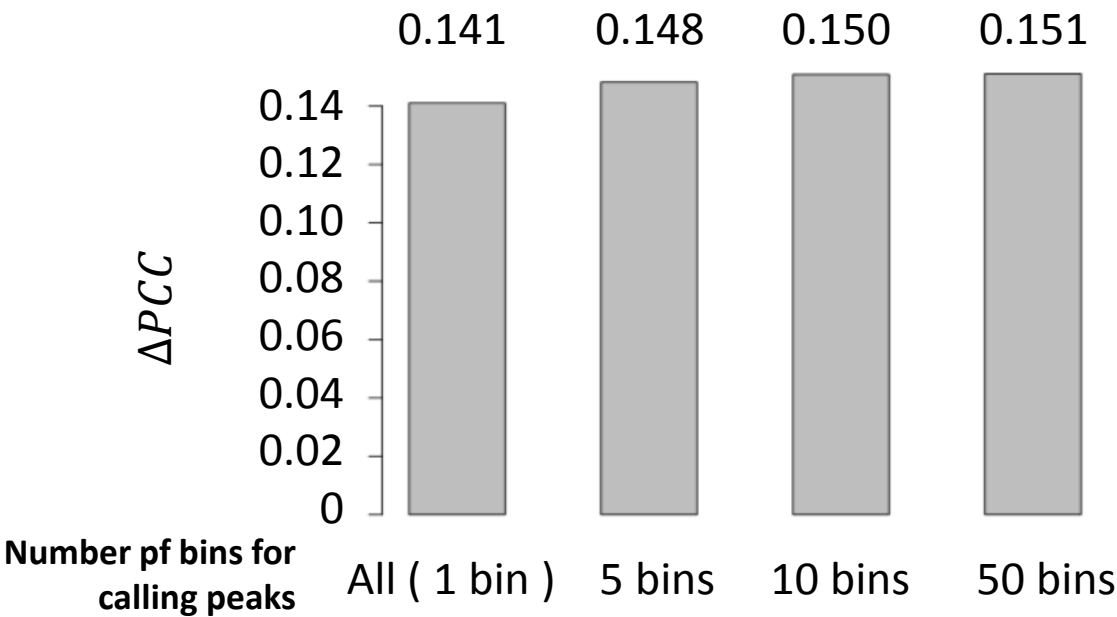
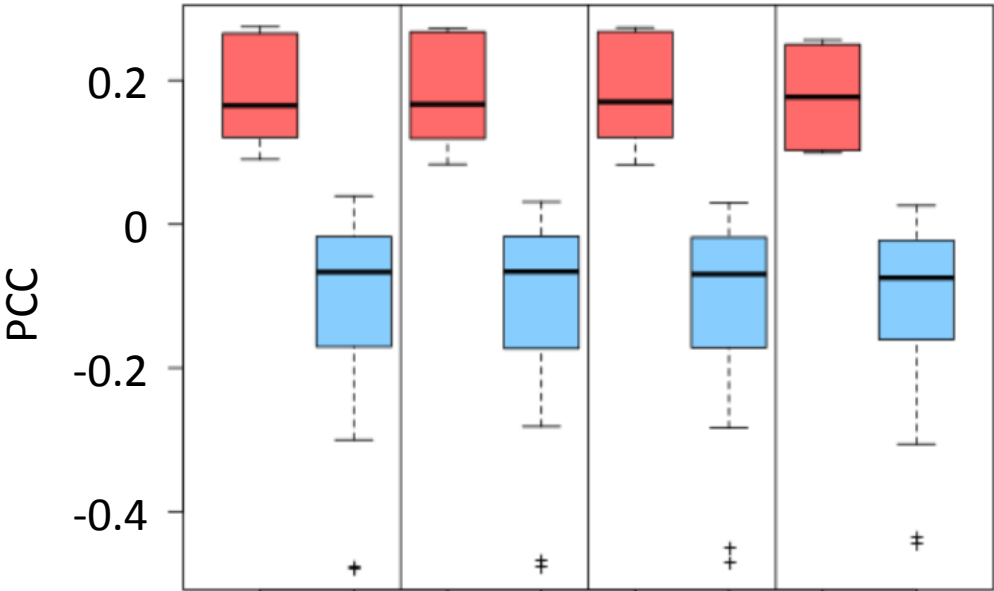
Number of bins & peak calling

Peaks are called independently on each bin of cells and then combined:

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CallPeaks(..., group.by="bin", ...) in Signac
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Human Frontal Cortex

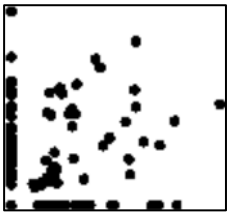
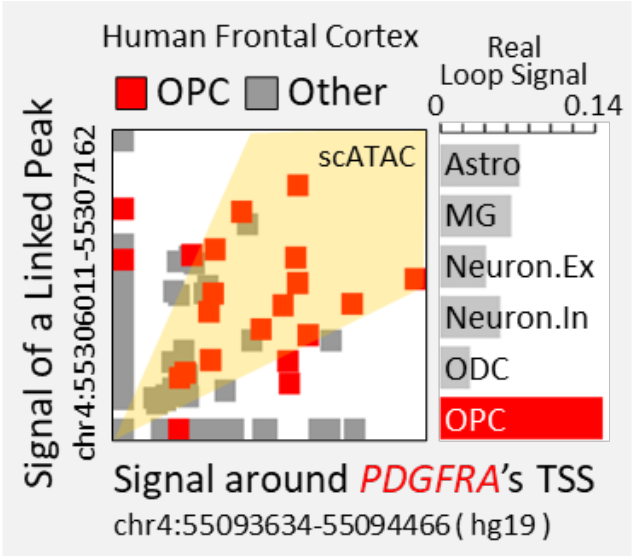
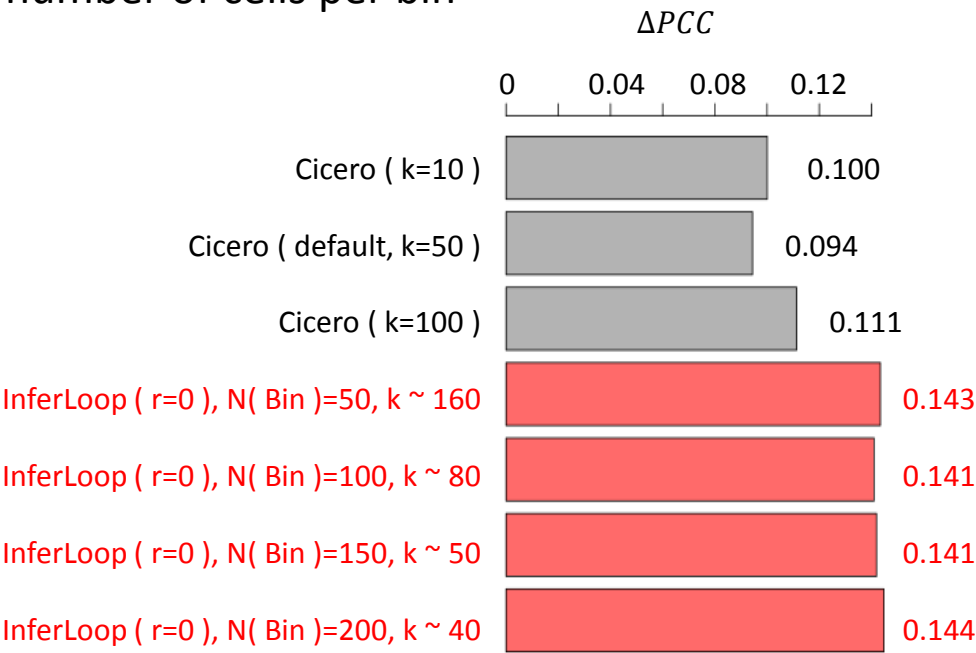
Matched Un-matched



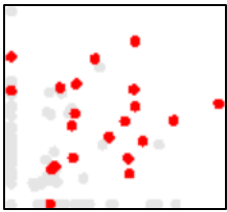
Using different k to run Cicero

Human Frontal Cortex: ~8000 cells

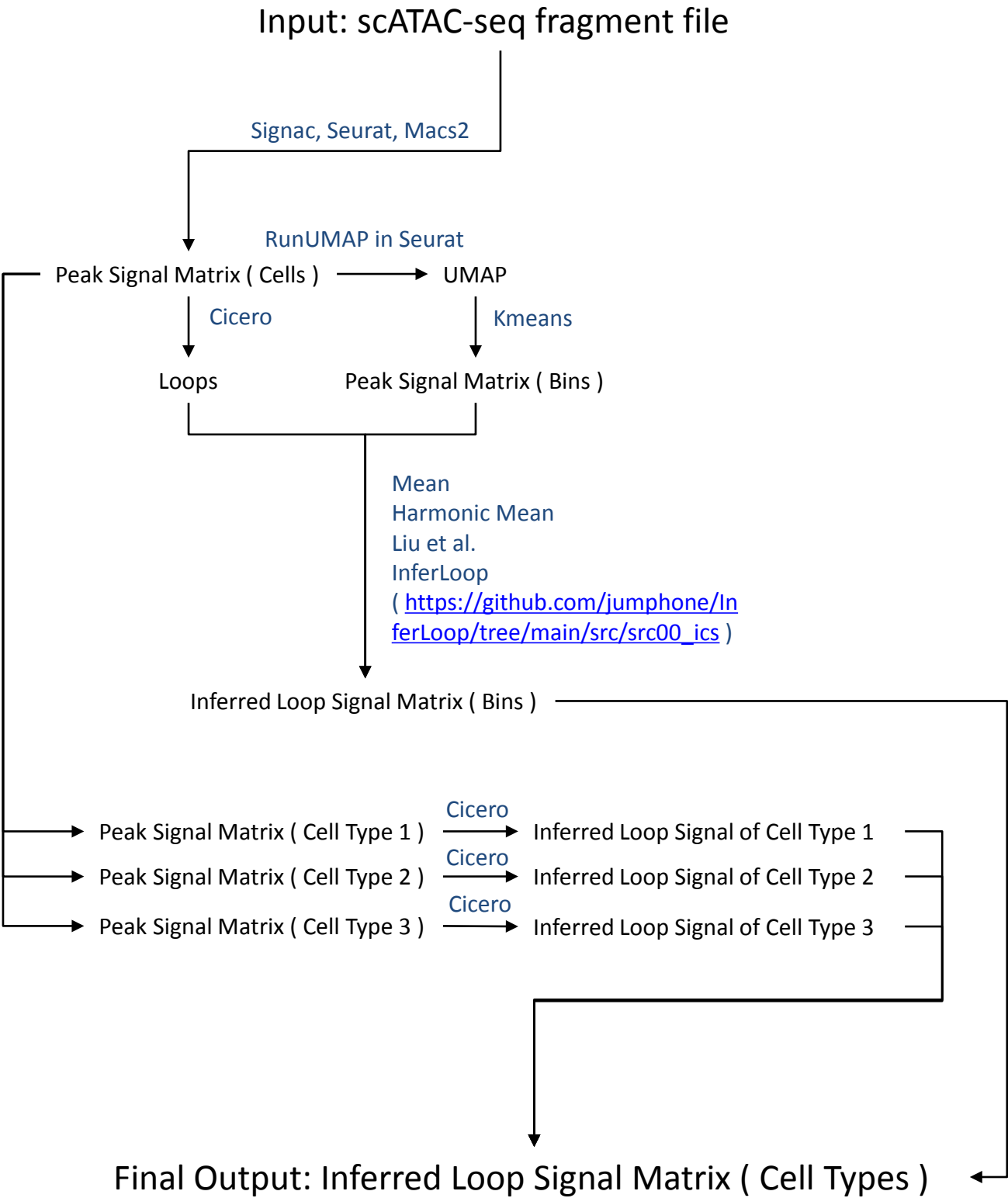
k: number of cells per bin



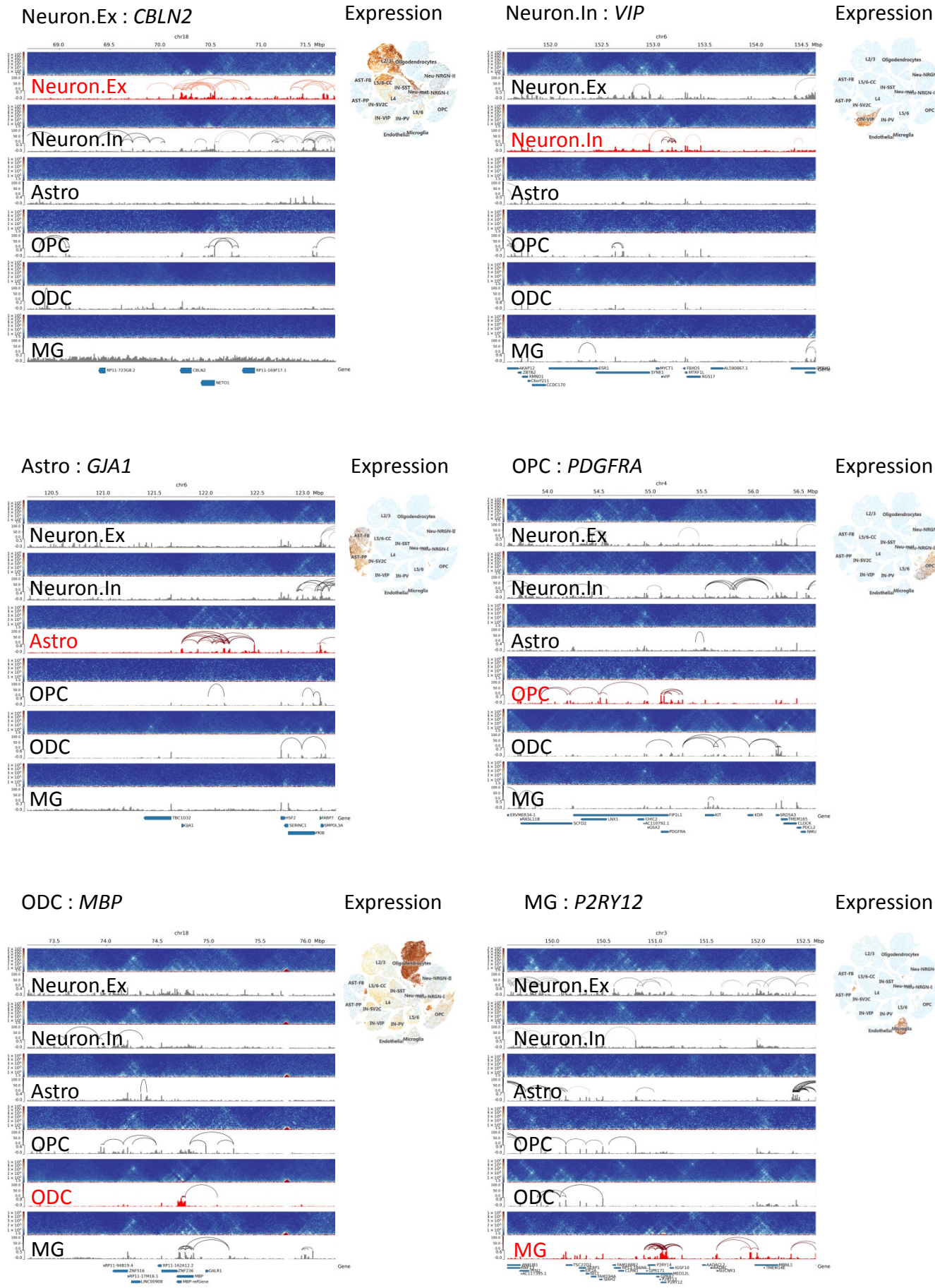
PCC (all bins)
= 0.27



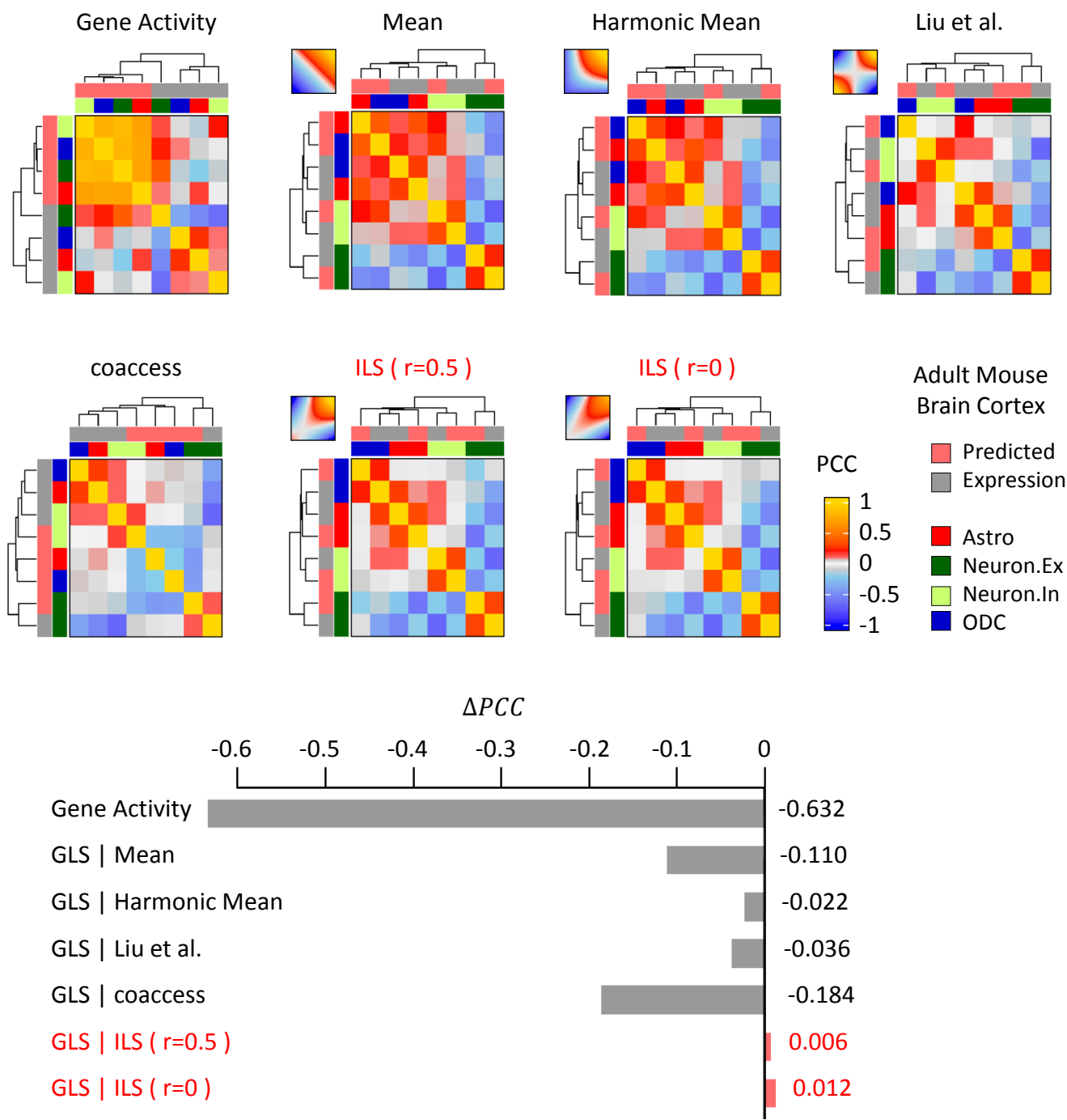
PCC (bins of OPC)
= 0.03



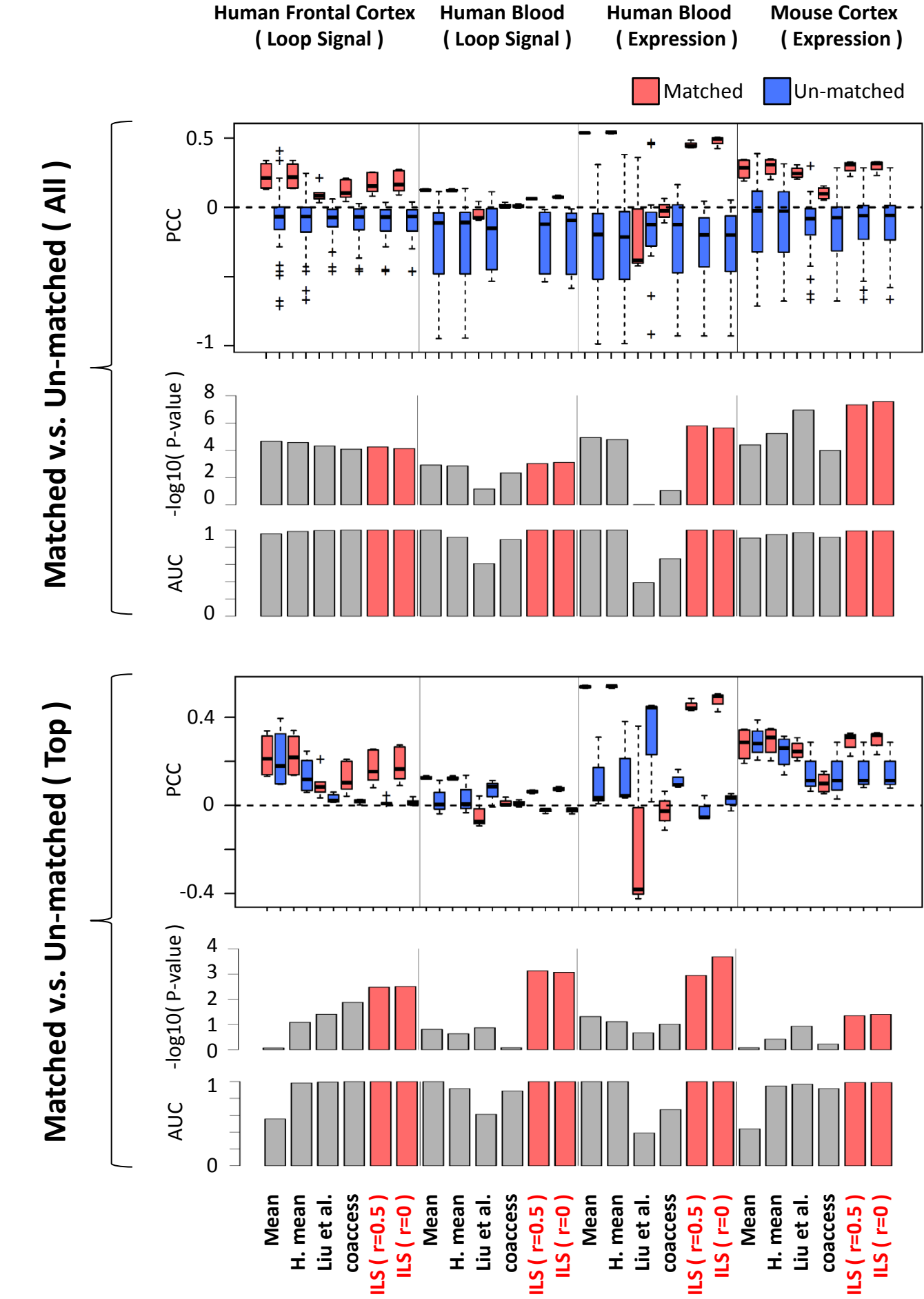
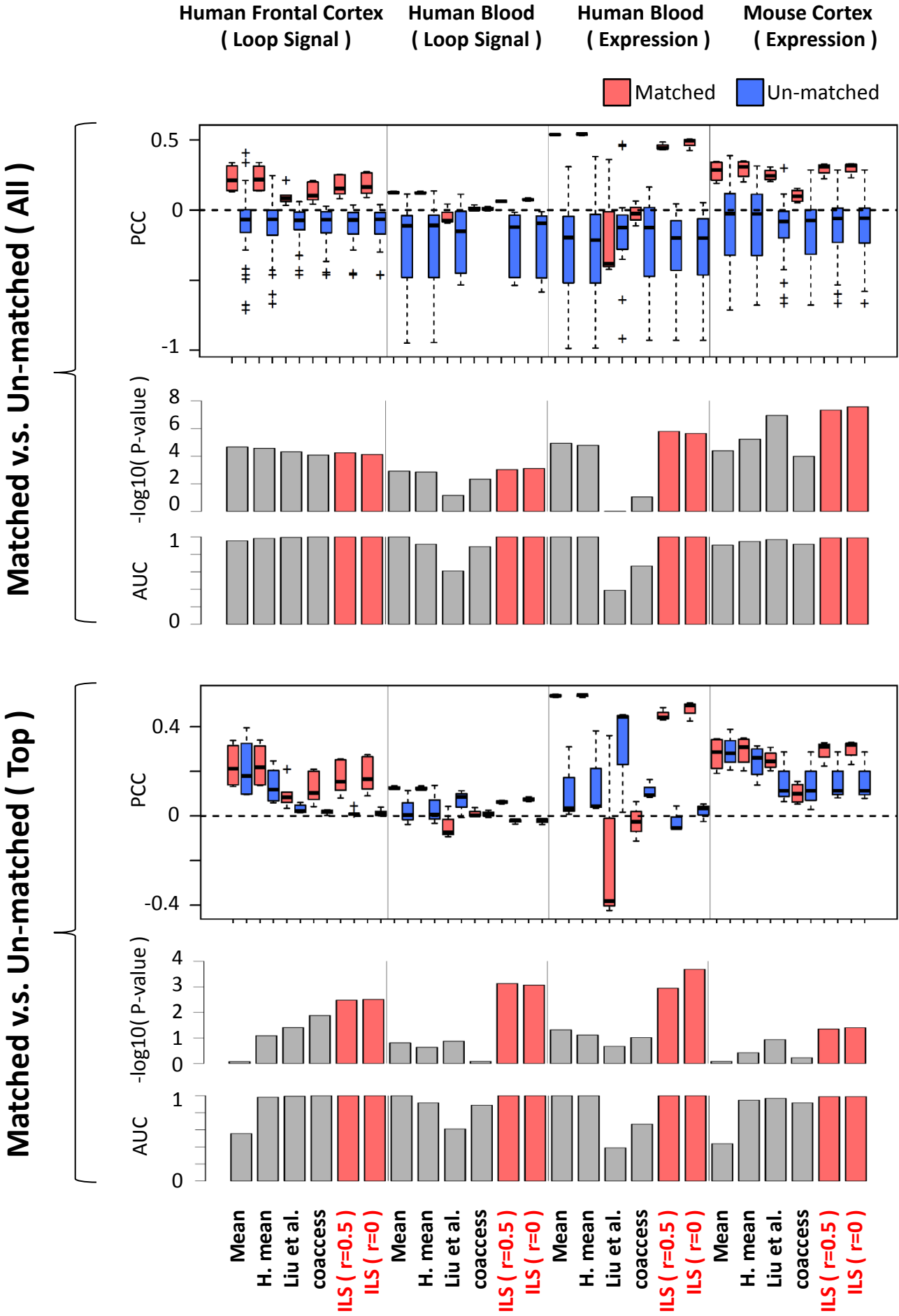
Supplementary Figure 9



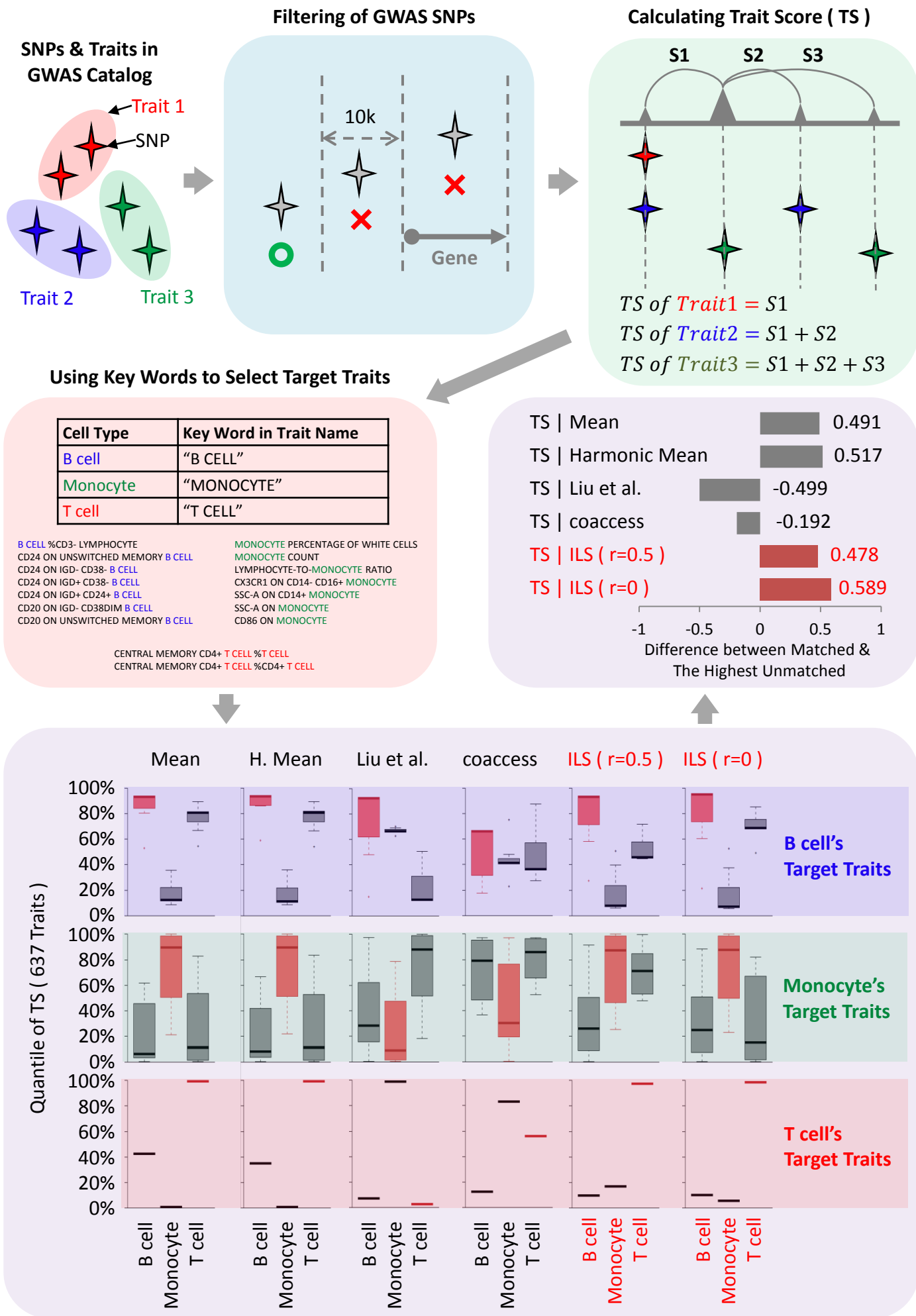
Supplementary Figure 10



Supplementary Figure 11

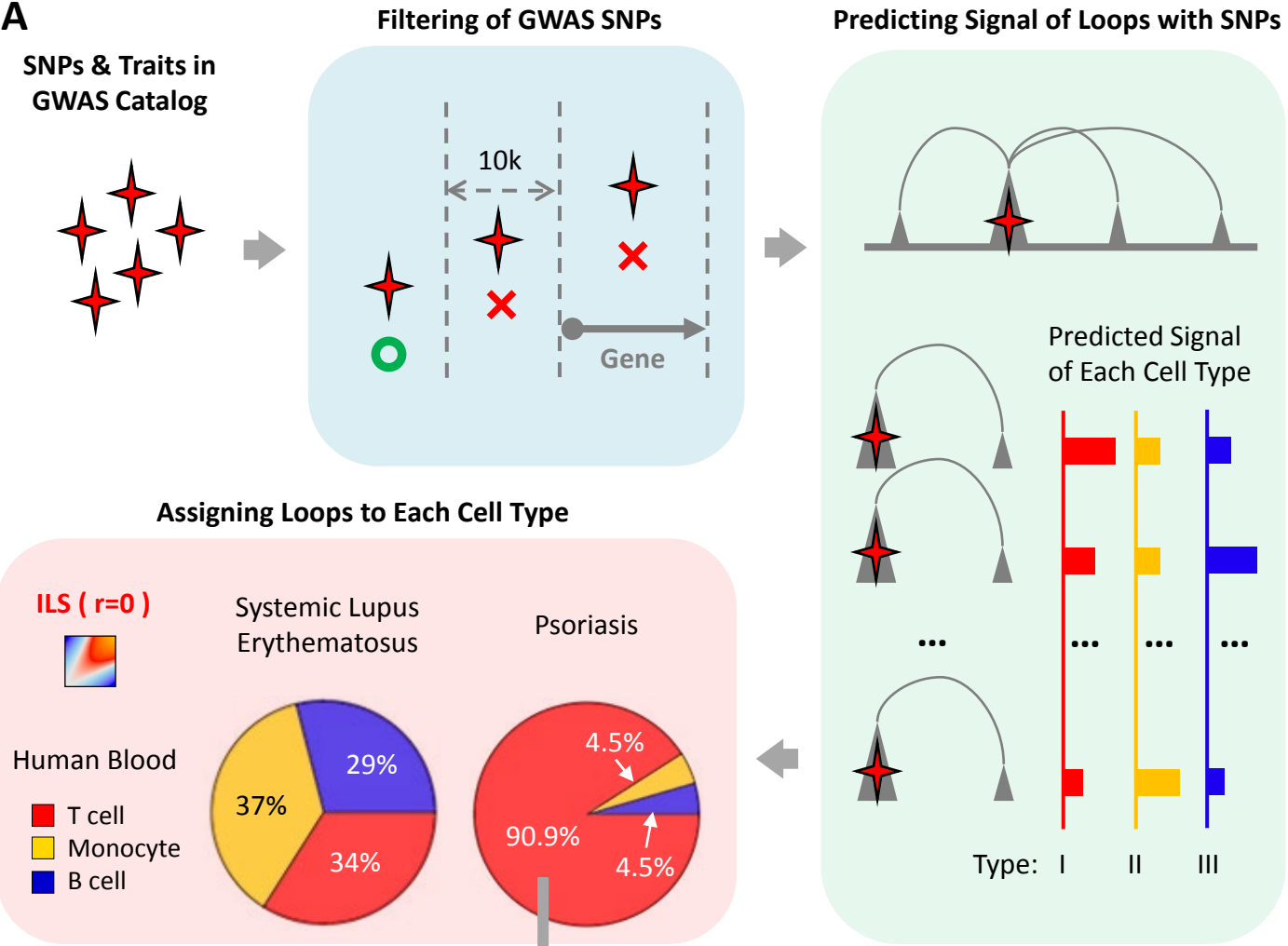


Supplementary Figure 12

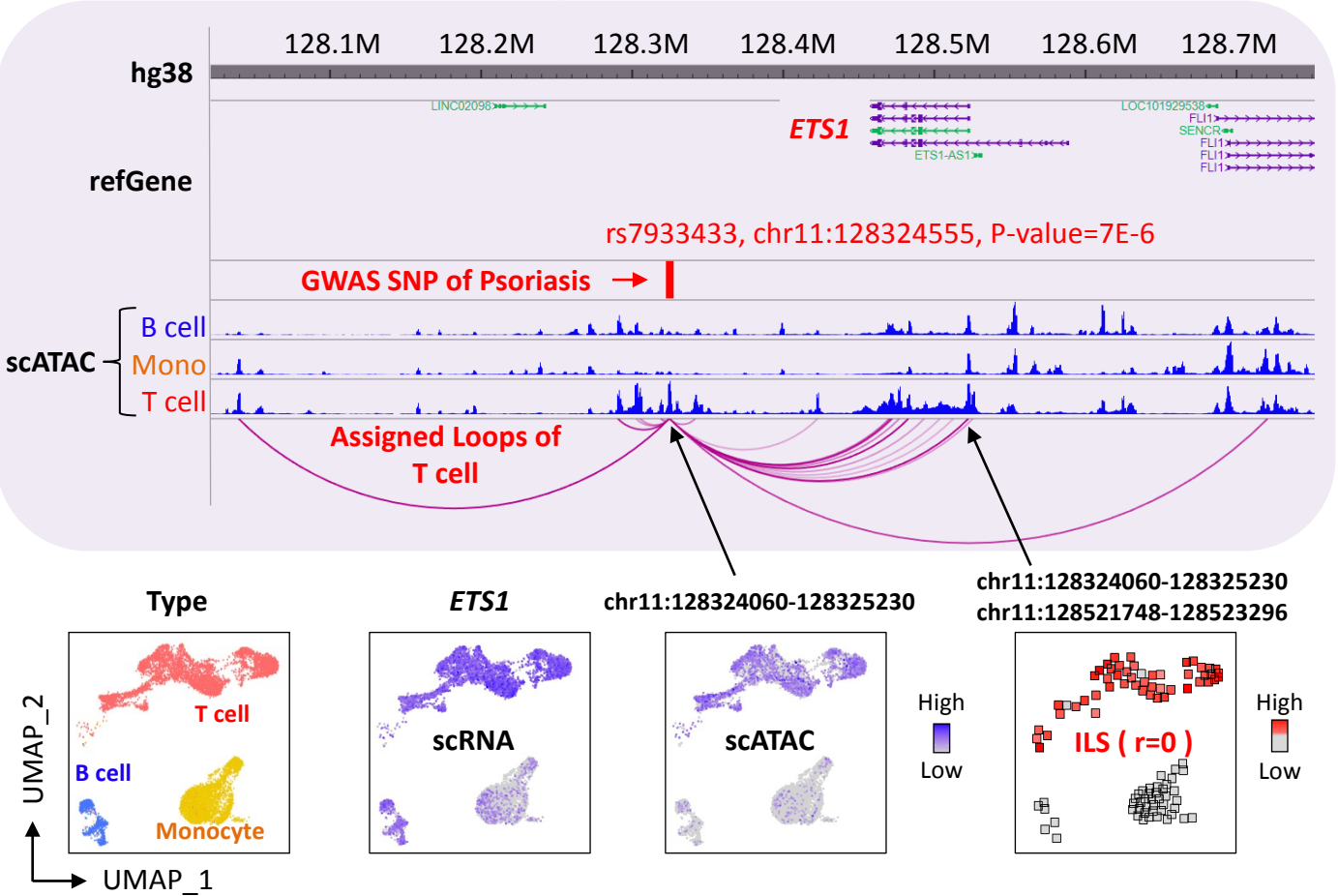


Supplementary Figure 13

A



B



Our study (hg38)

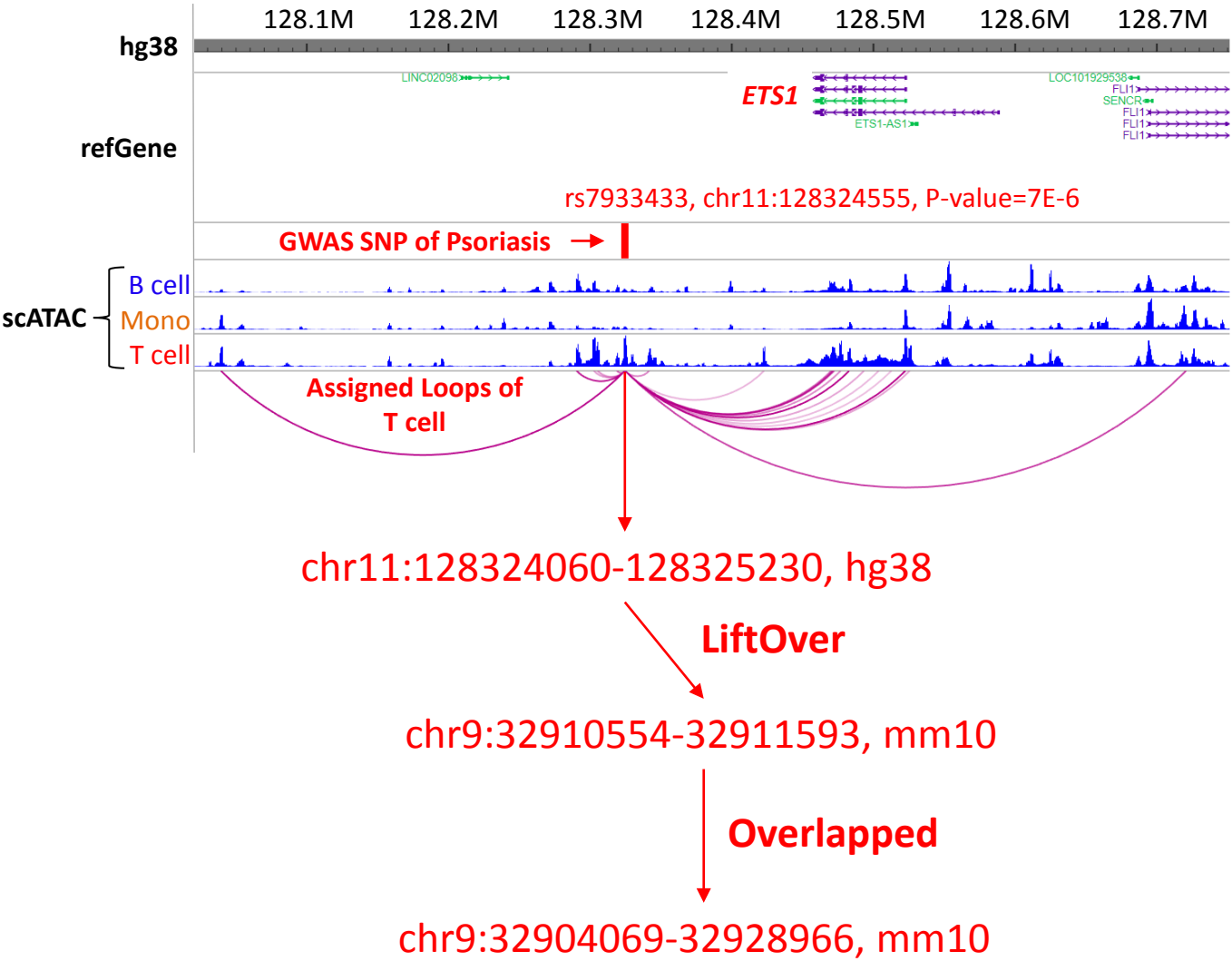
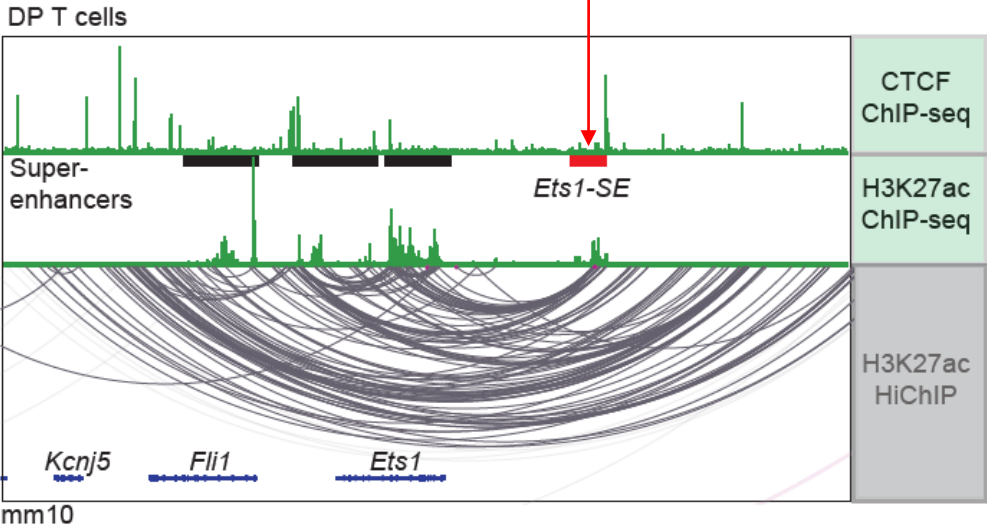


Figure 1d in Chandra's paper (mm10)



A multi-enhancer hub at the *Ets1* locus controls T cell differentiation and allergic inflammation through 3D genome topology, *bioRxiv*, 2022.10.28, <https://doi.org/10.1101/2022.10.28.514213>