

Supplemental Figures

Anna Hendrika Cornelia Vlot¹, Setareh Maghsudi², and Uwe
Ohler^{1, 3, 4, *}

¹The Berlin Institute for Medical Systems Biology, Max Delbrück
Center for Molecular Medicine, 10115 Berlin, Germany

²Department of Computer Science, University of Tübingen, 72076
Tübingen

³Department of Biology, Humboldt Universität zu Berlin, 10117
Berlin, Germany

⁴Department of Computer Science, Humboldt Universität zu Berlin,
10117 Berlin, Germany

* corresponding author, Uwe.Ohler@mdc-berlin.de

Results

SEMITONES identifies marker genes



Figure S1: Markers of immature classical monocytes (top row) and intermediate and classical monocytes (bottom row).



Figure S2: Cluster-based biological cell type annotations as taken from [1].

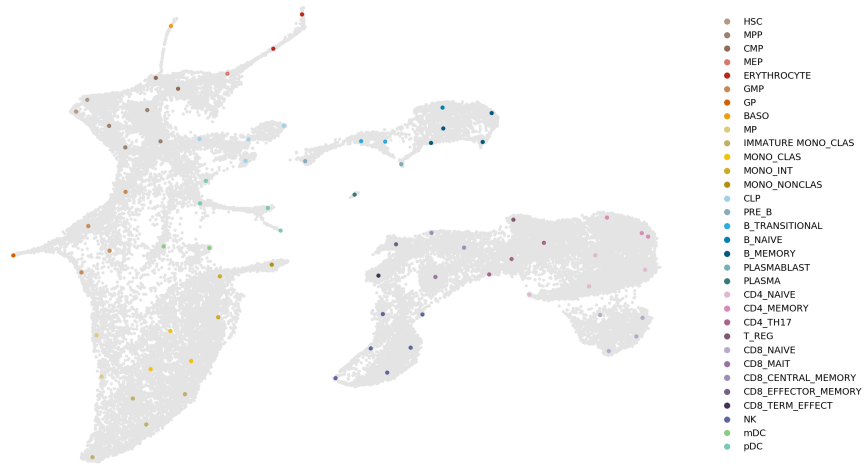


Figure S3: The annotation of manually selected reference cells from scRNA-seq data.



Figure S4: Data-driven cell selection using different embeddings (rows) and distance metrics (columns).



Figure S5: The influence of the radius of influence when using the RBF-kernel as a similarity metric. A higher value of γ is proportional to a larger radius of influence in the RBF-kernel.

SEMITONES identifies transcriptional regulators

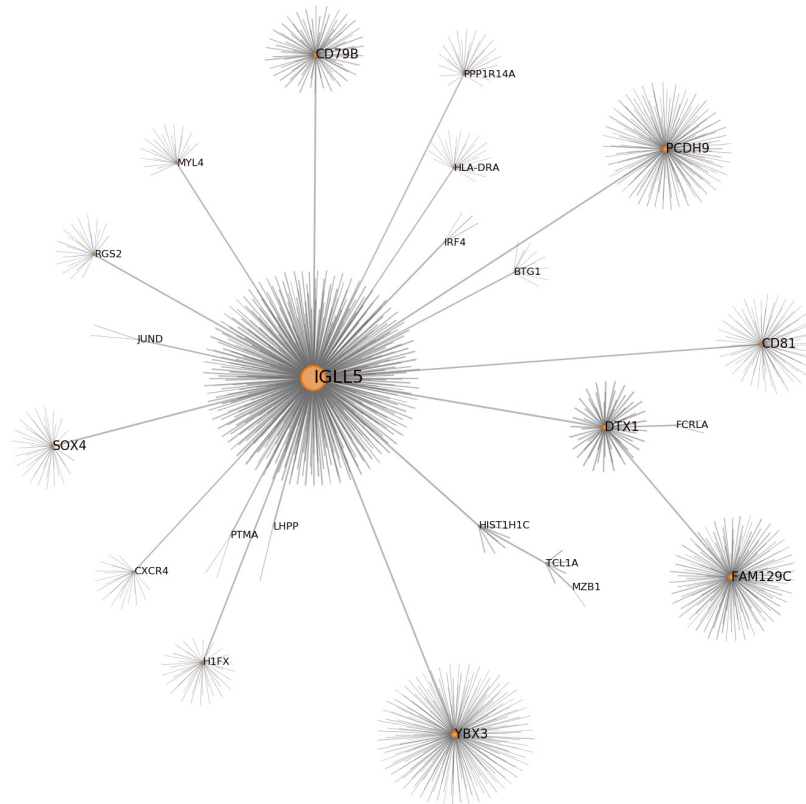


Figure S6: The interaction co-enrichment graph for the transitional B cell neighbourhood.

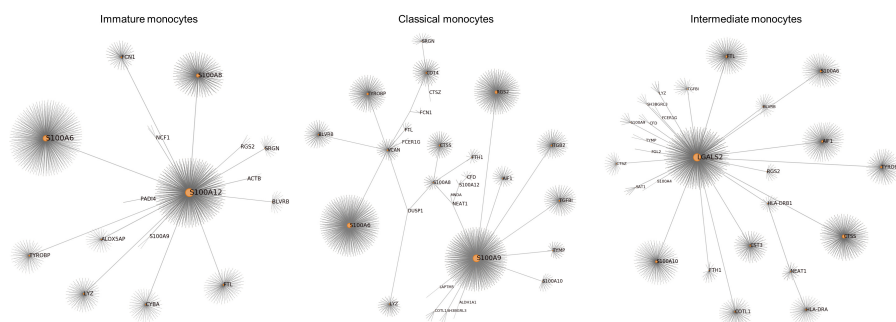


Figure S7: The interaction co-enrichment graphs for distinct, but highly similar, monocyte populations.

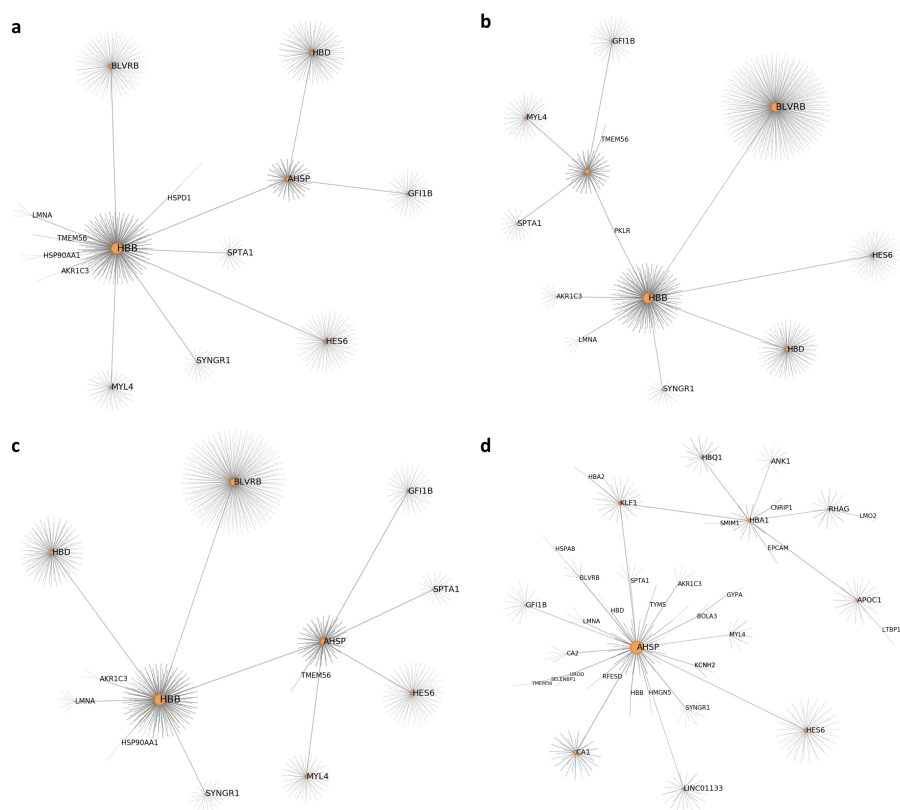


Figure S8: Erythrocyte a) interaction co-enrichment graph, b) maximum-value co-enrichment graph, c) median-value co-enrichment graph, d) minimum-value co-enrichment

SEMITONES for feature selection



Figure S9: UMAPs obtained using different numbers of top enriched (left) or highly variable genes (right).

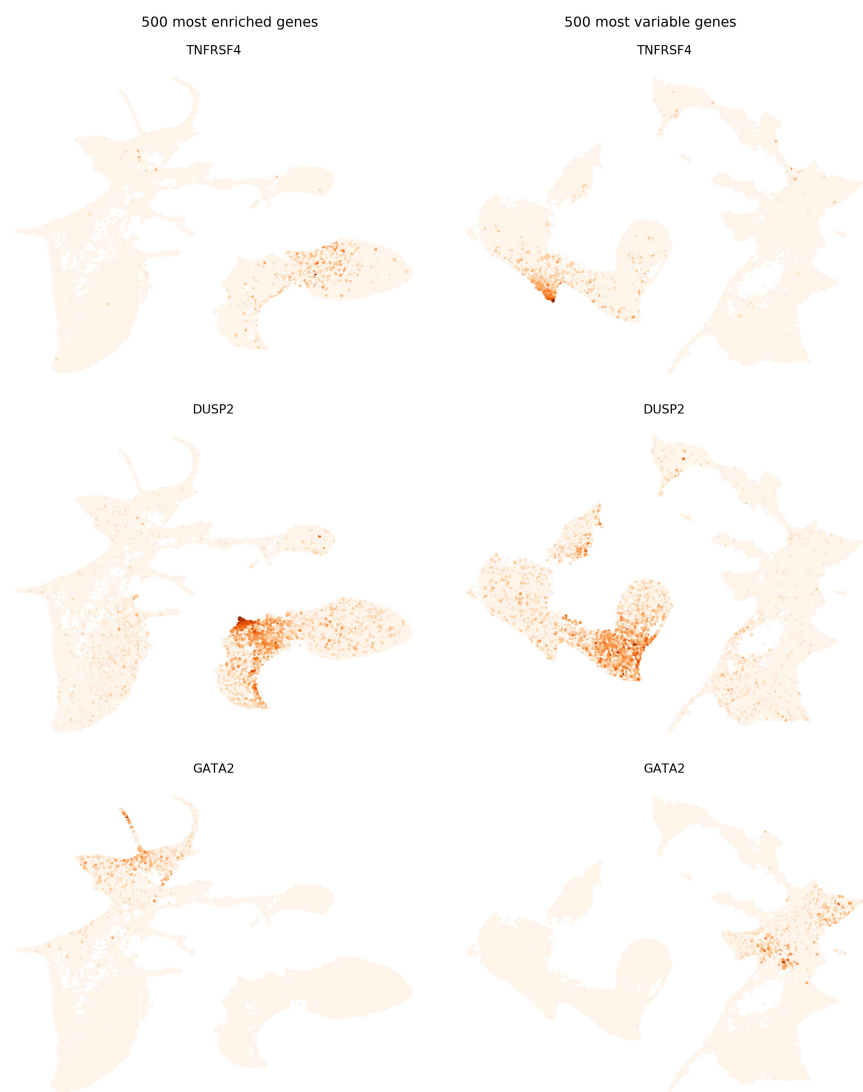


Figure S10: The expression levels of marker genes in 2D UMAPs computed using 500 most enriched genes (left) or 500 most variable genes (right).

SEMITONES identifies cell-specific cis-regulatory elements

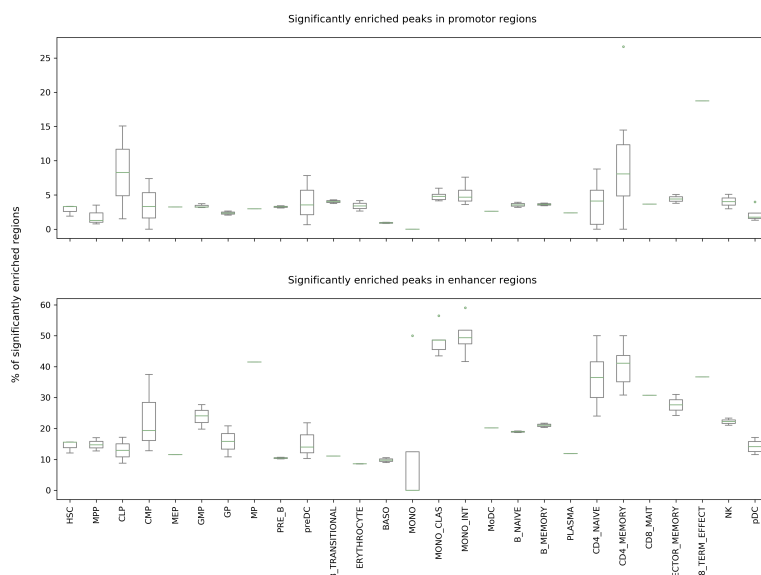


Figure S11: Percentage of significantly positively enriched peaks that fall in promoter (top panel) or enhancer (bottom panel) regions.

Scalability of SEMITONES

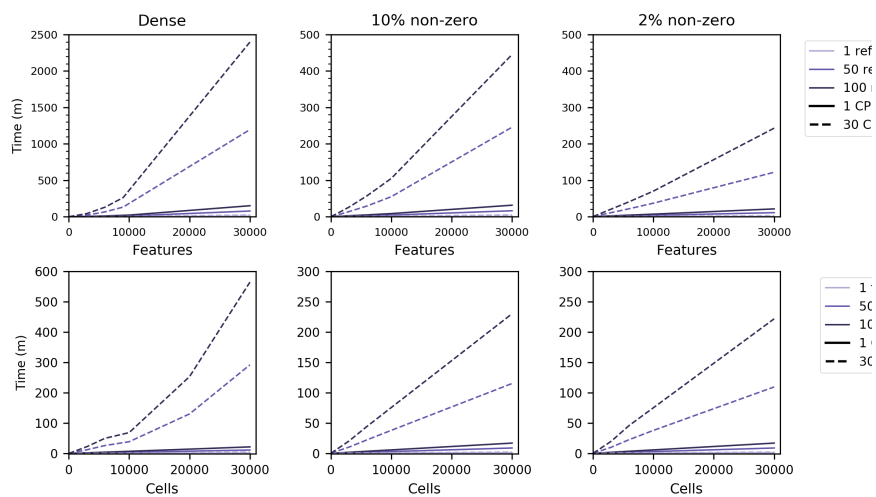


Figure S12: The run time of SEMITONES for 1-30,000 features or reference cells (top row) and 1-100 features or reference cells (bottom row).

Methods

Reference cell selection

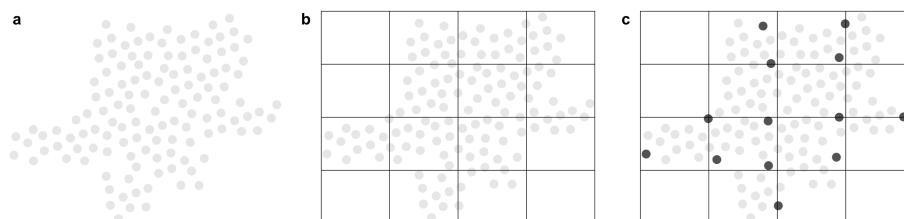


Figure S13: Fixed-grid cell selection. Given a 2D cell embedding (a), we fit a lattice graph of size $n \times n$ (b) and then select cells closest to the intersections of the horizontal and vertical grid lines (c).

References

- [1] Granja JM, Klemm S, McGinnes LM, Kathiria AS, Mezger A, Corces MR, et al. Single-cell multiomic analysis identifies regulatory programs in mixed-phenotype acute leukemia. *Nature Biotechnology*. 2019;37:1458–1465.