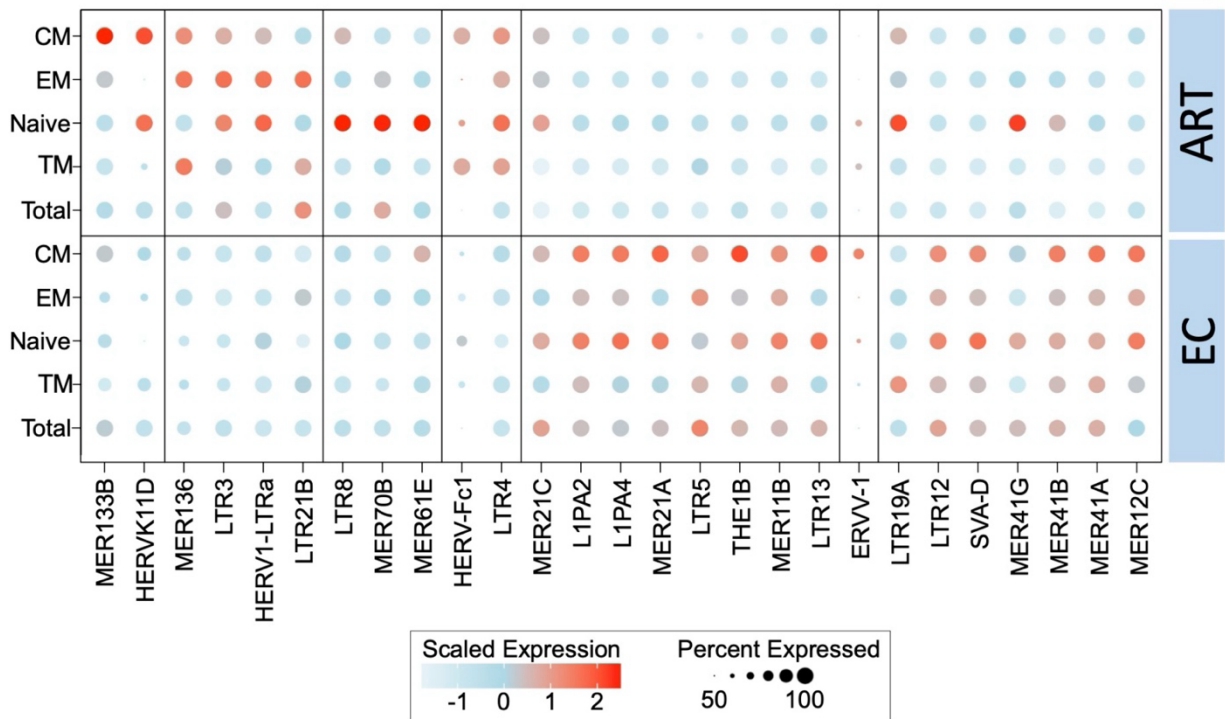


617 **Supplemental Figures**



618

619 **Supplementary Figure 1. Unique retrotranscriptome of ECs vs PLWH-on-ART**

620 Dot plot illustrating the intensity and abundance of the differential expression of TE

621 families between ECs and PLWH-on-ART, separated by CD4<sup>+</sup> T cell subtype. Coloration

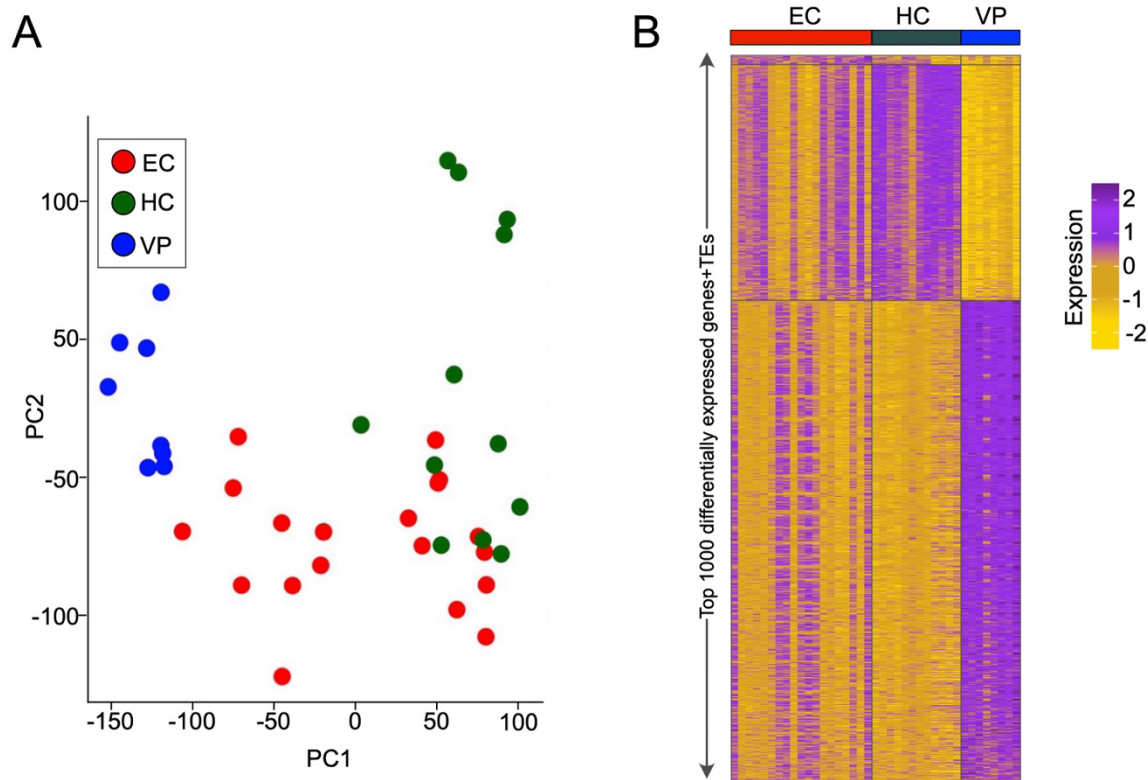
622 is scaled from lower (white) to higher (red) expression. Point size is directly proportional

623 to the percentage of that sample group expressing the given TE family. Data source:

624 [Jiang et al., 2020](#).

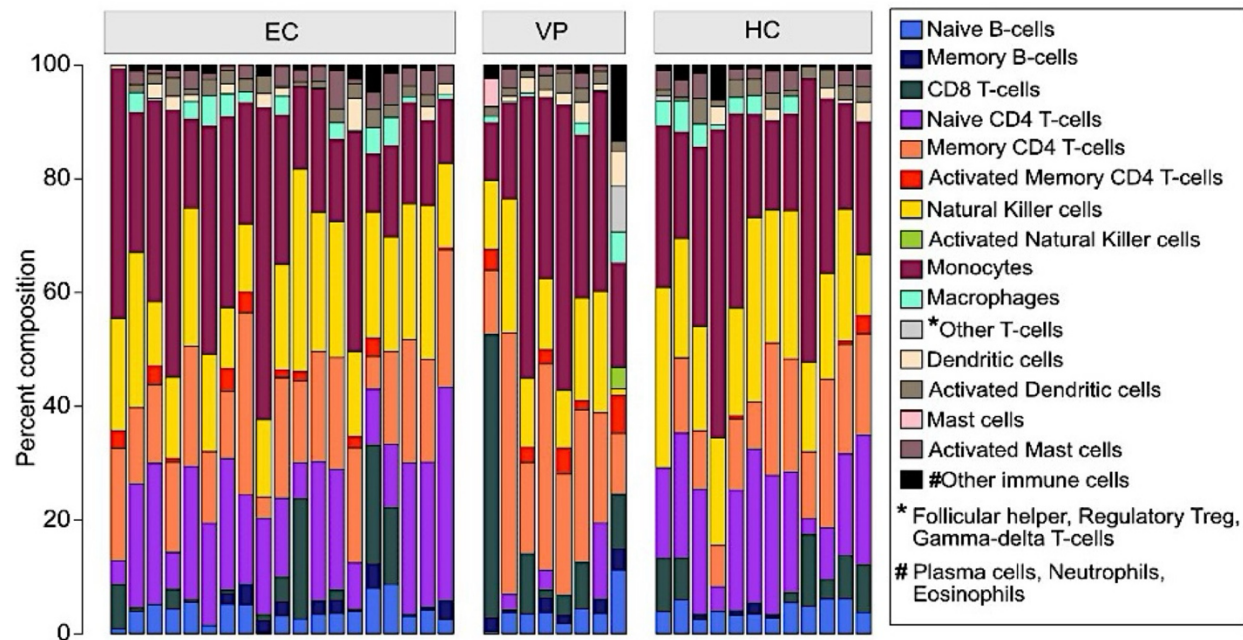
625

626



## Supplementary Figure 2. Heterogeneity of EC (retro)transcriptome

- PCA plot from PBMCs of ECs (red), VPs (blue), and HCs (green), based on the most variably expressed genes and TE families. Every dot is a PBMC sample from an individual. Data source: [Zhang et al., 2018](#).
- Heatmap displaying the scaled expression of genes and TEs distinguishing EC, HC, and VP samples shown in the previous plot. Every row denotes a gene or retroelement. Data source: [Zhang et al., 2018](#).



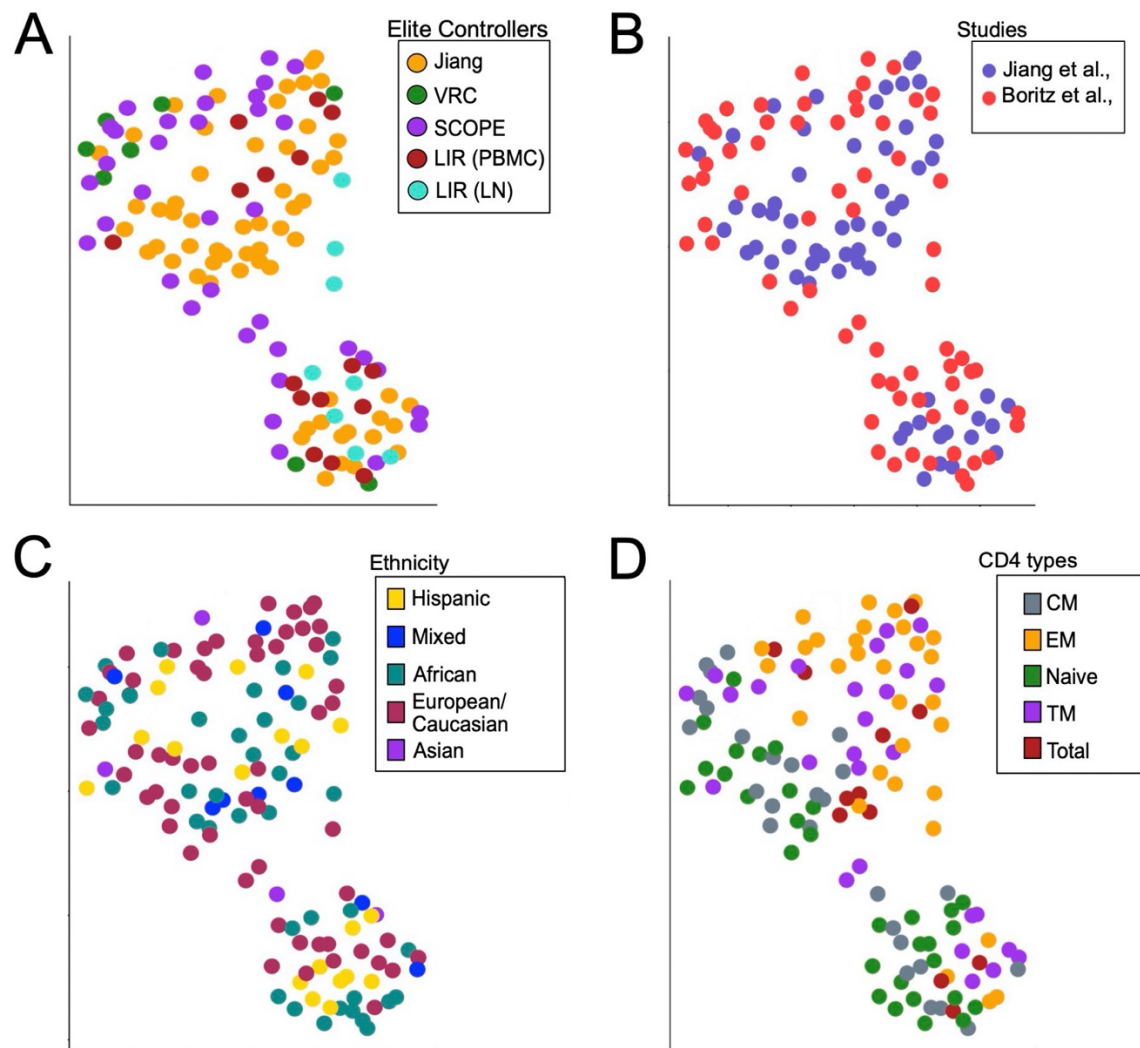
636

### 637 **Supplementary Figure 3. Immune cell profiles of ECs, VPs, and HCs**

638 Stacked barplot showing the immune cell composition of ECs, VPs, and HCs from the  
 639 deconvolution analysis of PBMC RNA-seq data. Data source: [Zhang et al., 2018](#).

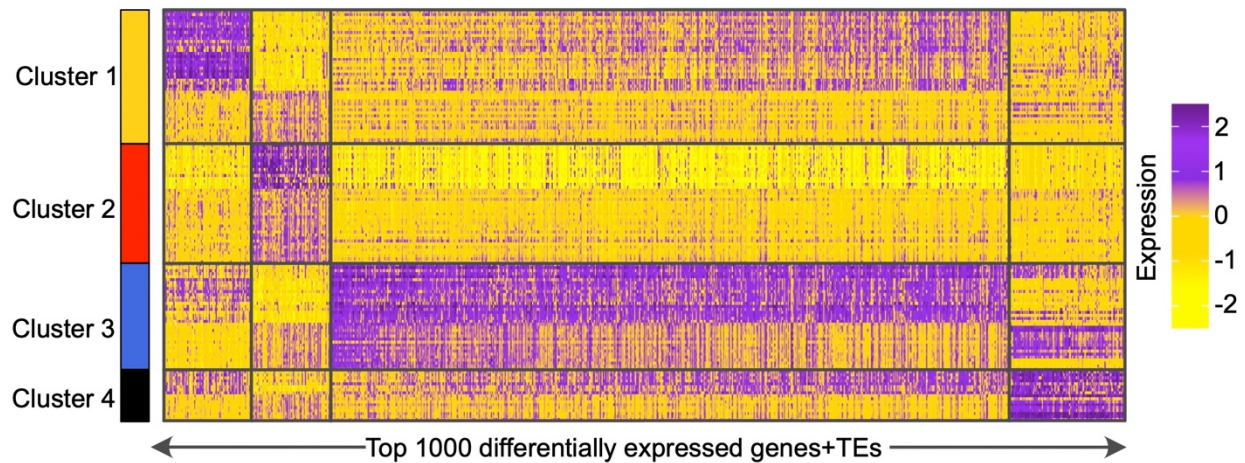
640

641



#### Supplementary Figure 4. Characterization of the EC clusters

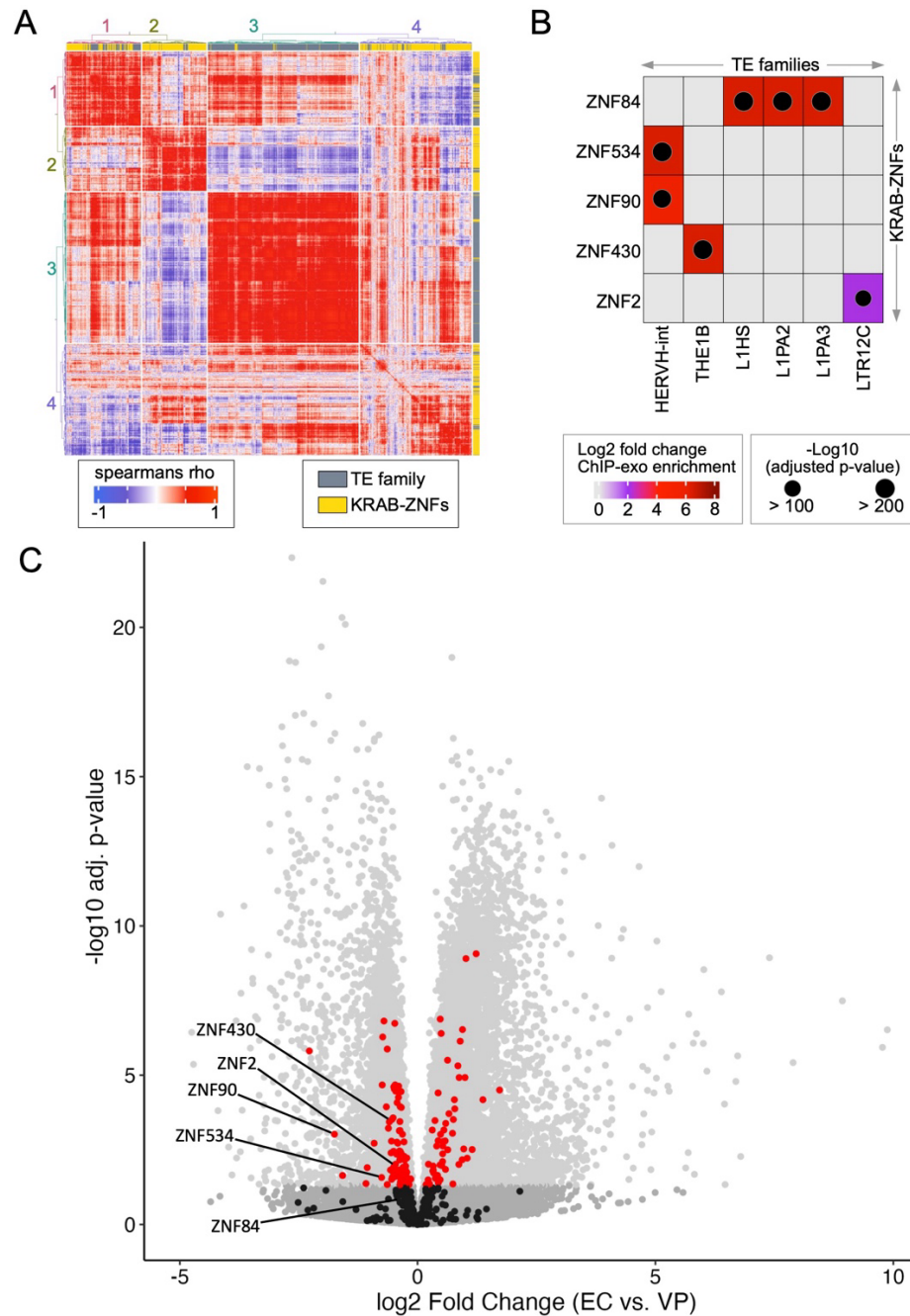
- UMAP plot from Figure 3A with samples color coded based on recruitment location of each patient. Data sources: [Jiang et al., 2020](#) and [Boritz et al., 2016](#).
- UMAP plot from Figure 3A with samples color coded based on the corresponding source study. Data sources: [Jiang et al., 2020](#) and [Boritz et al., 2016](#).
- UMAP plot from Figure 3A with samples color coded based on patient ancestry, inferred by HapMap to make direct variant comparisons. Data sources: [Jiang et al., 2020](#) and [Boritz et al., 2016](#).
- UMAP plot from Figure 3A with samples color coded based on the CD4<sup>+</sup> subtypes: naïve, central memory (CM), transitional memory (TM), effector memory (EM), and total. Data sources: [Jiang et al., 2020](#) and [Boritz et al., 2016](#).



**Supplementary Figure 5. Broad (retro)transcriptomic profile of the EC clusters**

Heatmap of the differentially expressed genes (DEGs) that distinguish the EC clusters from each other. Data source: [Jiang et al., 2020](#) and [Boritz et al., 2016](#).





659

## 660 **Supplementary Figure 6. KZNF dynamics in ECs**

661 **A.** Heatmap showing the correlation matrix of TE families with KZNFs across the  
662 analyzed EC samples. Clustered pairwise correlation matrix generated by  
663 weighted gene co-expression network analysis, using the Spearman's rank  
664 correlation & Euclidean distance. Data sources: [Jiang et al., 2020](#) and [Boritz et](#)  
665 [al., 2016](#).

- 666**      **B.** Heatmap showing the enrichment for binding of a subset of KZNFs to TE families  
**667**            with increased expression in ECs. Coloration is scaled from lower (gray) to  
**668**            higher (red) ChIP-exo enrichment in HEK cells. Point size is directly proportional  
**669**            to the -log10-transformed adjusted p-value. Data source: [Imbeault et al., 2017](#).  
**670**      **C.** Volcano plot displaying the differential expression of genes between EC and VP  
**671**            samples. KZFP genes are in red (p-value < 0.05) or black (p-value > 0.05). All  
**672**            other genes are in gray. Data source: [Zhang et al., 2018](#).