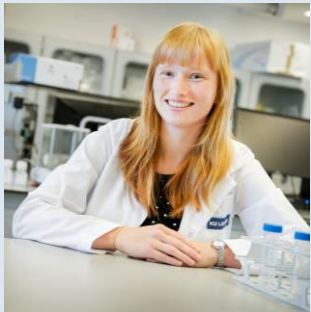


# Targeted single-cell transcriptomics for studying genotype-phenotype relationships

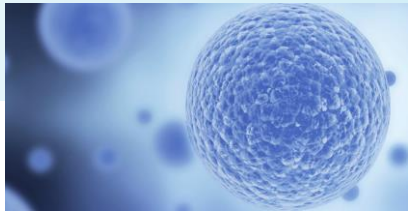


**Lies Van Horebeek<sup>1</sup>**, Klara Mallants<sup>1</sup>, Nina Dedoncker<sup>1</sup>, Suresh Poovathingal<sup>3</sup>, Bénédicte Dubois<sup>1,2</sup>, An Goris<sup>1</sup>

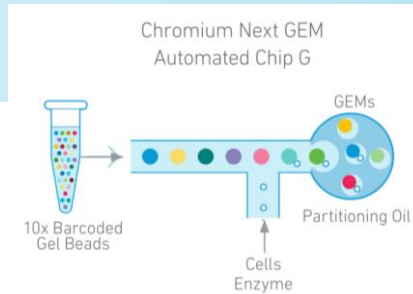
<sup>1</sup> Laboratory for Neuroimmunology, KU Leuven, BE; <sup>2</sup> University Hospitals Leuven, BE; <sup>3</sup> Laboratory of Computational Biology, VIB - KU Leuven, BE

# Method: standard 10x + targeted library prep

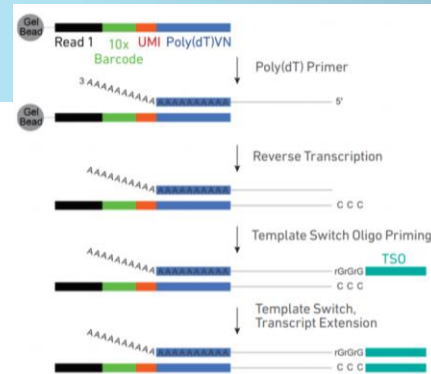
Single-cell sample



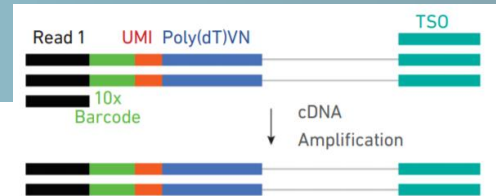
GEM generation



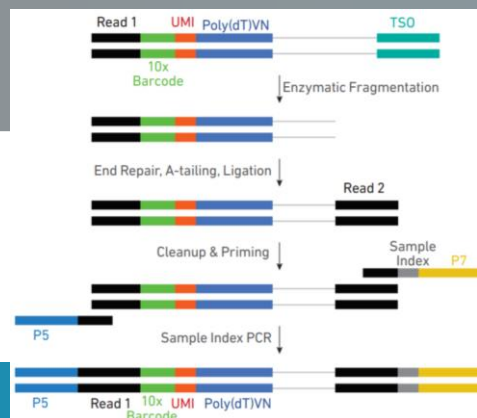
Reverse transcription within GEM



Bulk cDNA amplification



Standard library prep



Targeted library prep\*



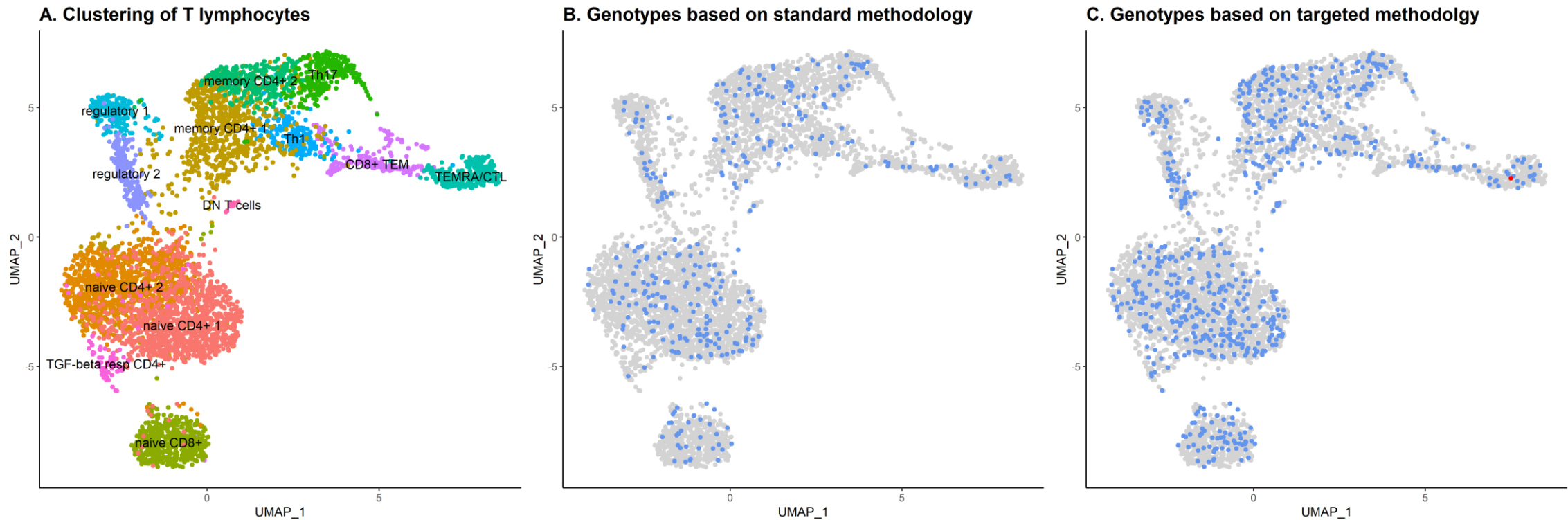
Sequencing & data analysis

Sequencing: Illumina  
Data analysis:

- Cell Ranger
- Seurat
- VarTrix (SC-GT)

\* Similar to: Nam *et al.* Nature 2019

# Applied to somatic *TREX1* variant in MS patient

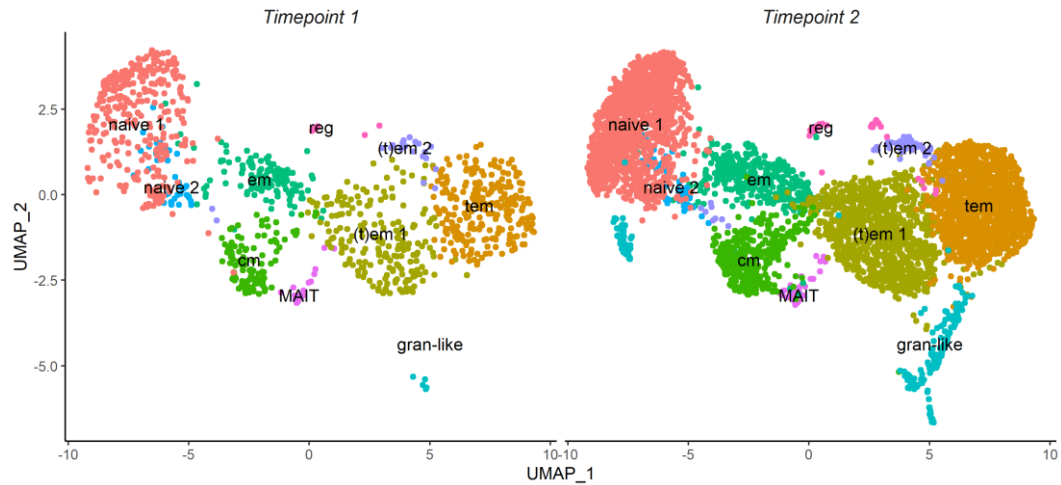


**A. Clusters of T lymphocytes, based on gene expression data.**

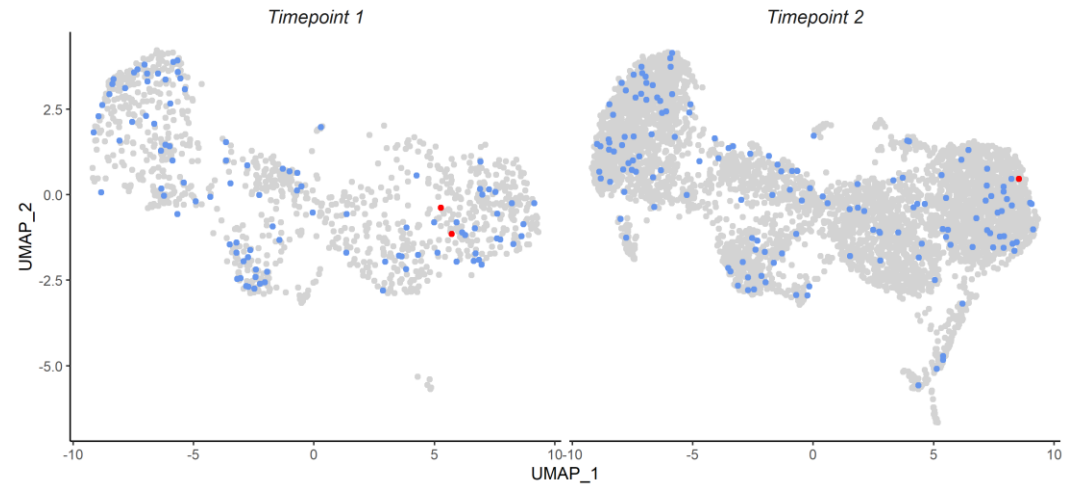
**B and C. Genotypes of cells for the somatic *TREX1* variant based on the standard 10x methodology (B) or the targeted methodology (C). Blue: reference allele only, red: alternate allele only, purple: reference and alternate allele, grey: no coverage of variant region.**

# CD8+ T cells of two timepoints

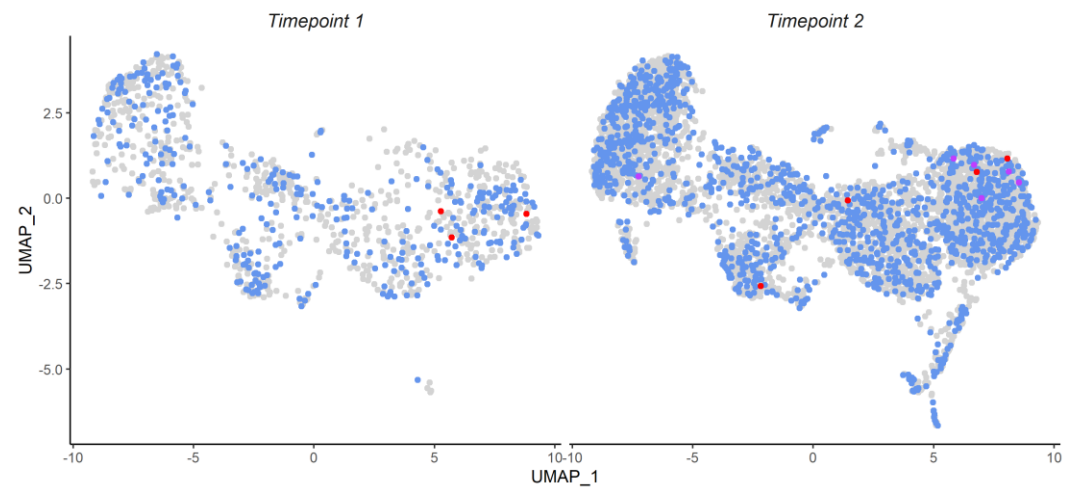
D. Clustering of CD8+ T lymphocytes



E. Genotypes based on standard methodology



F. Genotypes based on targeted methodology



D. Clusters of CD8+ T lymphocytes, based on gene expression data.  
E and F. Genotypes of cells for the somatic *TREX1* variant based on the standard 10x methodology (E) or the targeted methodology (F).  
Blue: reference allele only, red: alternate allele only, purple: reference and alternate allele, grey: no/insufficient coverage of variant region.