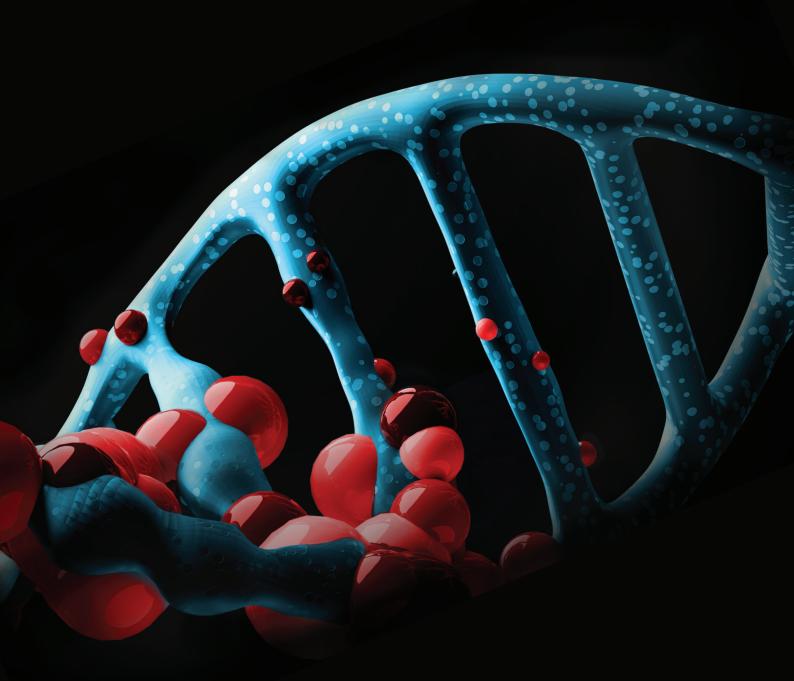


# Simultaneous DNA + Protein + CpG Methylation Analysis on Tapestri®





## Simultaneous DNA + Protein + CpG Methylation Analysis on Tapestri®

A workflow optimized for human samples that reveals clonal architecture, epigenetic states, and differentiation all in the same cells.

### Your Benefits

Bulk assays and plate-based methods miss rare clones and can't connect **genotype**  $\leftrightarrow$  **epigenetic state**  $\leftrightarrow$  **lineage** in the same cells.

Aging and disease reshape hematopoiesis through subtle clonal dynamics that require high cell throughput and multi-omic context to see.

#### With CpG Methylation Profiling on Tapestri, you can:

- Reveal clonal hematopoiesis earlier by pairing single-cell DNA variants/CNVs
  with CpG methylation enabling the ability to detect early heme malignancies.
- Track lineage trajectories retrospectively in human samples with no transgenics required.
- Confidently identify and prioritize new therapeutic or monitoring targets with single-cell multi-omic insights.

# Applications & Clinical Implications for Single-Cell Multi-Omics Methylation Analysis

#### **ONCOLOGY APPLICATIONS**

#### **Tumor Classification & Clonal Subtyping**

 Methylation defined subtypes and associations with BRAF mutation status in colorectal cancers

#### **Prognostic Biomarker Development**

CpG methylation profiling in GBM as a predictive therapy related biomarker

#### **Predicting Therapy Response**

 Profiling of methylation state of therapeutic targets (hypomethylating agents) along with co-occurring genetic lesions that might modulate response

#### **NON-ONCOLOGY APPLICATIONS**

#### **Aging & Longevity Biomarker Research**

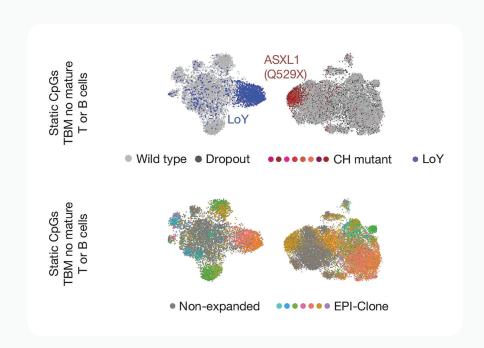
 Tracking CHIP via combined genotyping ("DTA") and epigenetic drift profiling via CpG signatures

#### **Cell and Gene Therapy**

 Ensure engineered cell lines don't retain aberrant methylation patterns linked to malfunction

#### **Immunology & Autoimmune Disorders**

 Linking specific CpG patterns to immune activation states and mutational subgroups in a lineage-specific manner



 $Figure \ 1: Single-cell\ DNA\ and\ CpG\ methylation\ maps\ showing\ clonal\ architecture\ and\ lineage\ composition.$ 

(Top) UMAPs computed per donor on a consensus set of static CpGs, highlighting distinct cell populations containing the specified CH mutations: Loss of Y chromosome (LoY) (A.6, blue) and ASXL1 Q529X mutation (A.7, red).

(Bottom) UMAPs as above, highlighting clonal populations identified using EPI-Clone.

Figure 1 adapted from Scherer M., Singh, I., Braun, M.M et al. Clonal tracing with somatic epimutations reveals dynamics of blood ageing. Nature 643, 478 (2025).

#### WHAT YOU CAN UNCOVER

- Age-linked epigenetic signatures tied to known and newly detected clonal hematopoiesis (CH) mutations.
- Stable methylation states that illuminate clonal structure beyond transient differentiation signals.
- Emergent SNVs with coordinated methylation + lineage shifts that reveal clone-specific targets to track.

#### WHAT THE DATA COULD REVEAL

- CpG profiles capture age-linked epigenetic signatures, distinguishing donors by age and aligning with both established and newly discovered CH mutations.
- Stable methylation states reveal clonal structure less tied to instantaneous differentiation.
- Analysis of thousands of cells exposes emergent SNVs alongside coordinated methylation and lineage shifts, revealing new targets to track.

## Tapestri Workflow For CpG Methylation

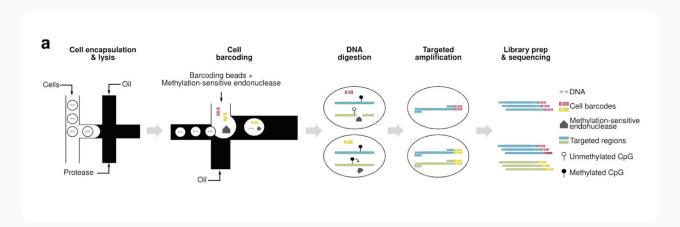
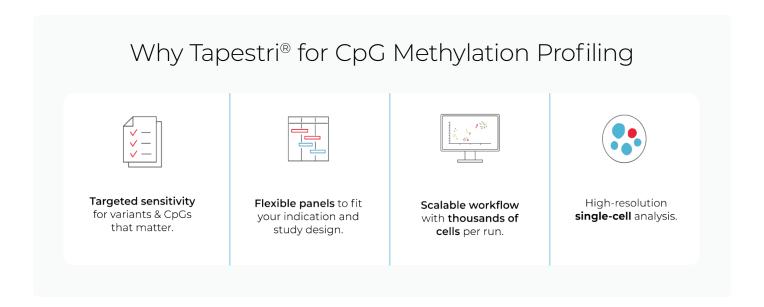


Figure 2: Tapestri CpG methylation workflow overview. Figure 2 adapted from Bianchi, A., Scherer, M., Zaurin, R. et al. scTAM-seq enables targeted high-confidence analysis of DNA methylation in single cells. Genome Biol 23, 229 (2022).

- 1. Prepare single-cell suspensions from human bone marrow or blood.
- 2. Tapestri DNA (+ Protein optional) to call SNVs/CNVs at single-cell resolution.
- **3.** Add methylation-sensitive endonuclease (Single-Cell Targeted Analysis of the Methylome- scTAM-seq) to target informative CpG loci (bisulfite-free).
- **4. Analyze & visualize** by methylation-specific Tapestri Pipeline and ready-to-use dimension reduction and cluster analysis.

### Made for Teams Like Yours

- Translational & clinical research teams monitoring clonal hematopoiesis (CH) /Acute Myeloid Leukemia (AML) / Multiple Myeloma (MM) risk, minimal residual disease (MRD), or therapy response.
- Aging & stem-cell biology labs mapping differentiation potential across life span.
- Drug discovery groups prioritizing clone-specific vulnerabilities.
- Therapy developers targeting the epigenome.



# Ready to see your clones and CpGs together?

Visit support.missionbio.com for additional information.

Contact: support@missionbio.com

### Selected references

- Scherer M. et al., Nature (2025): Clonal tracing with somatic epimutations reveals dynamics of blood ageing.
- Bianchi A. et al., Genome Biology (2022): scTAM-seq enables targeted high-confidence analysis of DNA methylation in single cells.
- Lareau C.A. et al., bioRxiv (2023): Codon affinity in mitochondrial DNA shapes evolutionary and somatic fitness.