

## A Raw, latent loadings

$$\lambda \in \mathbb{R}^{(N \times K \times T)}$$

$N = 400,000$  individuals;  $K = 21$  signatures;

$T = 52$  timepoints

Softmax, across  
 $k$  signatures

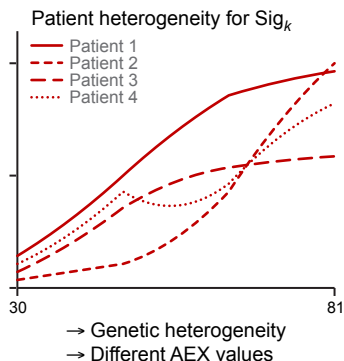
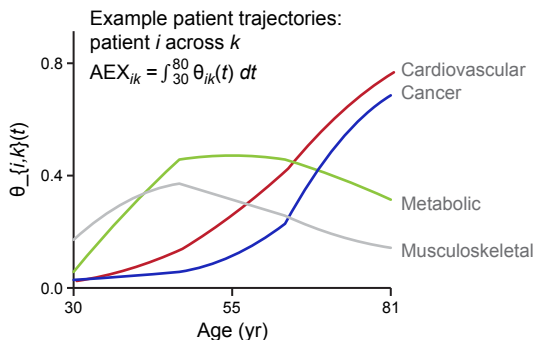
## B Signature weights

$$\theta \in [0, 1]^{(N \times K \times T)}$$

$$\sum_{k=1}^K \theta_{\{i,k,t\}}; \forall \text{ individual } i, \text{ timepoint } t$$

Temporal  
integration

## C Individual signature trajectories and area under curve calculation



## D AEX matrix for genetic discovery

$$AEX \in \mathbb{R}^{(N \times K)}$$

Quantitative phenotypes for GWAS analysis

Each  $AEX_{ik}$  represents individual  $i$ 's lifetime exposure to signature  $k$