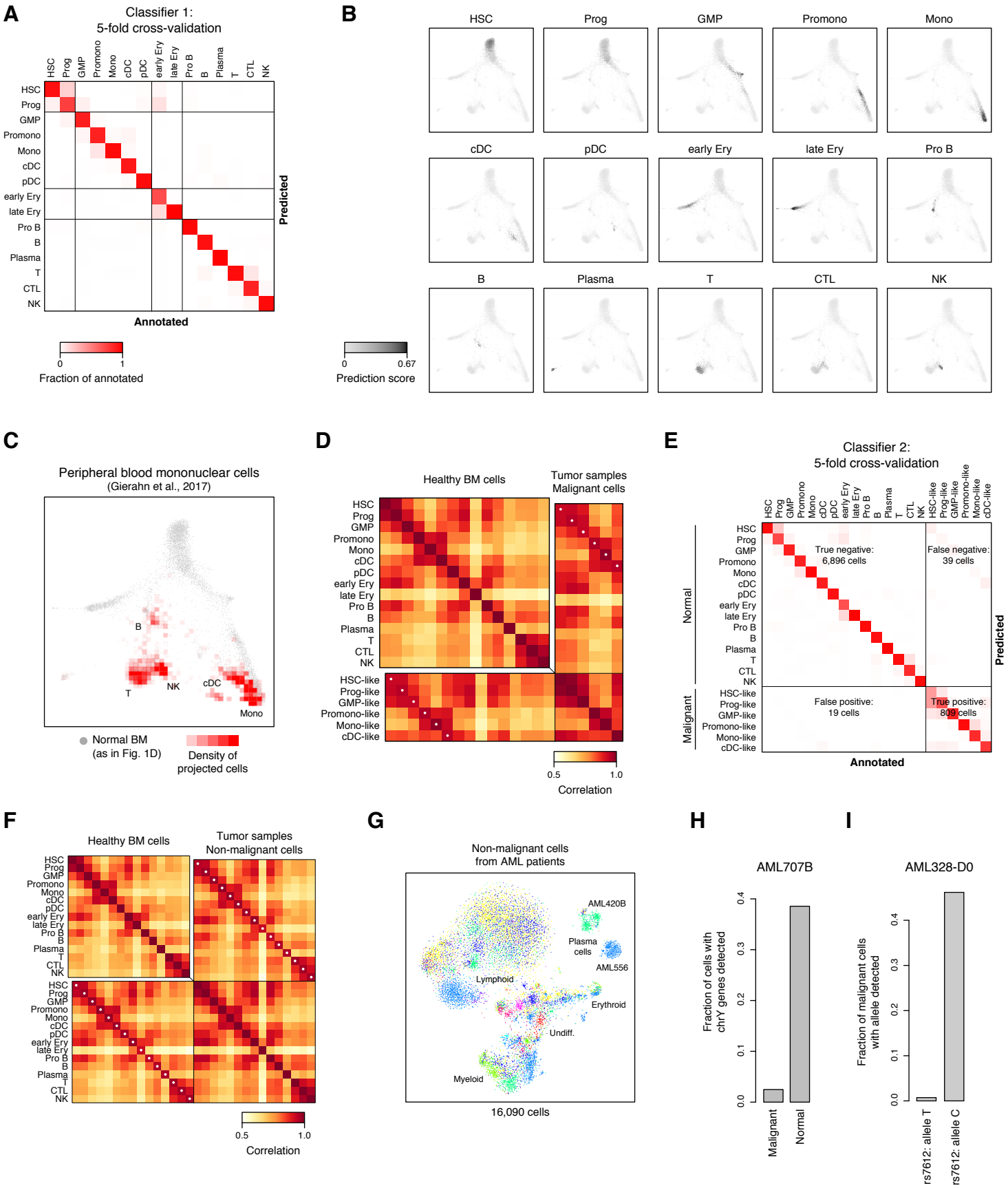Figure S3, related to Figure 4