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Scope: Format: Amount: GEO accession:

Series GSE108989

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Status Public on Oct 29, 2018

Title Lineage tracking reveals dynamic relationships of **T cells in colorectal cancer**

Organism [Homo sapiens](#)

Experiment type Expression profiling by high throughput sequencing

Summary T cells are central players in cancer immunotherapy¹, yet some of their fundamental properties such as development and migration within tumours remain elusive. The enormous T cell receptor (TCR) repertoire, required for recognising foreign and self-antigens^{2,3}, could serve as lineage tags to track these T cells in tumours⁴. Here, we obtained transcriptomes of 11,138 single T cells from 12 colorectal cancer (CRC) patients and developed STARTRAC (Single T-cell Analysis by Rna-seq and Tcr TRACking) indices to quantitatively analyse dynamic relationships among 20 identified T cell subsets with distinct functions and clonalities. While both CD8⁺ effector and ?exhausted? T cells exhibited high clonal expansion, they were independently connected with tumour-resident CD8⁺ effector memory cells, implicating a TCR-based fate decision. Of the CD4⁺ T cells, the majority of tumour-infiltrating Tregs showed clonal exclusivity, whereas certain Treg clones were developmentally linked to multiple TH clones. Notably, we identified two IFNG⁺ TH1-like clusters in tumours, the GZMK⁺ TEM and CXCL13⁺ TH1-like clusters, which were associated with distinct IFN-?-regulating transcription factors, EOMES/RUNX3 and BHLHE40, respectively. Only BHLHE40⁺ CXCL13⁺ TH1-like cells were preferentially enriched in tumours of microsatellite-unstable (MSI) patients, which might explain their favourable response rates to immune-checkpoint blockade. Furthermore, we found IGFLR1 to be highly expressed in both BHLHE40⁺CXCL13⁺ TH1-like and CD8⁺ exhausted T cells and possessed co-stimulatory functions. Our integrated STARTRAC analyses provided a powerful avenue to comprehensively dissect the T cell properties in CRC, which could shed new insights into the dynamic relationships of T cells in other cancers.

Overall design **T cells from CRC patients were sorted, profiled by Smart-seq2 and sequenced on HiSeq4000.** Based on FACS analysis, single cells of different subtypes, including CD8⁺ T cells (CD3⁺ and CD8⁺), T helper cells (CD3⁺, CD4⁺ and CD25⁻), and regulatory T cells (CD3⁺, CD4⁺ and CD25^{high}) were sorted to perform RNA sequencing. The categories ?""sampleType"" column in the SAMPLES section? contain PTC(CD8⁺ T cells from peripheral blood), NTC(CD8⁺ T cells from adjacent normal colonrectal tissues) ,TTC (CD8⁺ T cells from tumor), PTH(CD3⁺, CD4⁺ and CD25⁻ T cells from peripheral blood), NTH(CD3⁺, CD4⁺ and CD25⁻ T cells from adjacent normal colonrectal tissues), TTH(CD3⁺, CD4⁺ and CD25⁻ T cells from tumor), PTR(CD3⁺, CD4⁺ and CD25^{high} T cells from peripheral blood), NTR(CD3⁺, CD4⁺ and CD25^{high} T cells from adjacent normal colonrectal tissues), TTR(CD3⁺, CD4⁺ and CD25^{high} T cells from tumor), PTY(CD3⁺, CD4⁺ and CD25mediate T cells from peripheral blood), NTY(CD3⁺, CD4⁺ and CD25mediate T cells from adjacent normal colonrectal tissues), TTY(CD3⁺, CD4⁺ and CD25medate T cells from tumor), PP7(CD3⁺, CD4⁺ T cells from peripheral blood), NP7(CD3⁺, CD4⁺ T cells from adjacent normal colonrectal tissues), TP7(CD3⁺, CD4⁺ T cells from tumor).

Raw data access provided at: European Genome-phenome Archive (EGA) under accession EGAS00001002791

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Platforms (1) [GPL20301](#) Illumina HiSeq 4000 (Homo sapiens)
Samples (12) [GSM3356219](#) P0123
[More...](#) [GSM3356220](#) P0215
[GSM3356221](#) P0309

Relations
BioProject [PRJNA429424](#)

Listing of Individual Cells header descriptions

UniqueCell_ID
Patient_ID
majorCluster
sampleType

Data table

UniqueCell_ID	Patient_ID	majorCluster	sampleType
NTH5-20180123	P0123	CD4_C01-CCR7	NTH
NTH64-20180123	P0123	CD4_C01-CCR7	NTH
NTR57-20180123	P0123	CD4_C01-CCR7	NTR
PP7-100-20180123	P0123	CD4_C01-CCR7	PP7
PP7-106-20180123	P0123	CD4_C01-CCR7	PP7
PP7-109-20180123	P0123	CD4_C01-CCR7	PP7
PP7-114-20180123	P0123	CD4_C01-CCR7	PP7
PP7-118-20180123	P0123	CD4_C01-CCR7	PP7
PP7-129-20180123	P0123	CD4_C01-CCR7	PP7
PP7-134-20180123	P0123	CD4_C01-CCR7	PP7
PP7-137-20180123	P0123	CD4_C01-CCR7	PP7
PP7-145-20180123	P0123	CD4_C01-CCR7	PP7
PP7-152-20180123	P0123	CD4_C01-CCR7	PP7
PP7-160-20180123	P0123	CD4_C01-CCR7	PP7
PP7-162-20180123	P0123	CD4_C01-CCR7	PP7
PP7-171-20180123	P0123	CD4_C01-CCR7	PP7
PP7-178-20180123	P0123	CD4_C01-CCR7	PP7
PP7-182-20180123	P0123	CD4_C01-CCR7	PP7
PP7-185-20180123	P0123	CD4_C01-CCR7	PP7

Total number of rows: **11138**

Table truncated, full table size **417 Kbytes**.

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Format

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Supplementary file	Size	Download	File type/resource
GSE108989_CRC.TCell.S10805.norm.centered.txt.gz	368.5 Mb	(ftp) (http)	TXT
GSE108989_CRC.TCell.S11138.TPM.txt.gz	351.6 Mb	(ftp) (http)	TXT
GSE108989_CRC.TCell.S11138.count.txt.gz	69.7 Mb	(ftp) (http)	TXT

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Raw data not provided for this record

