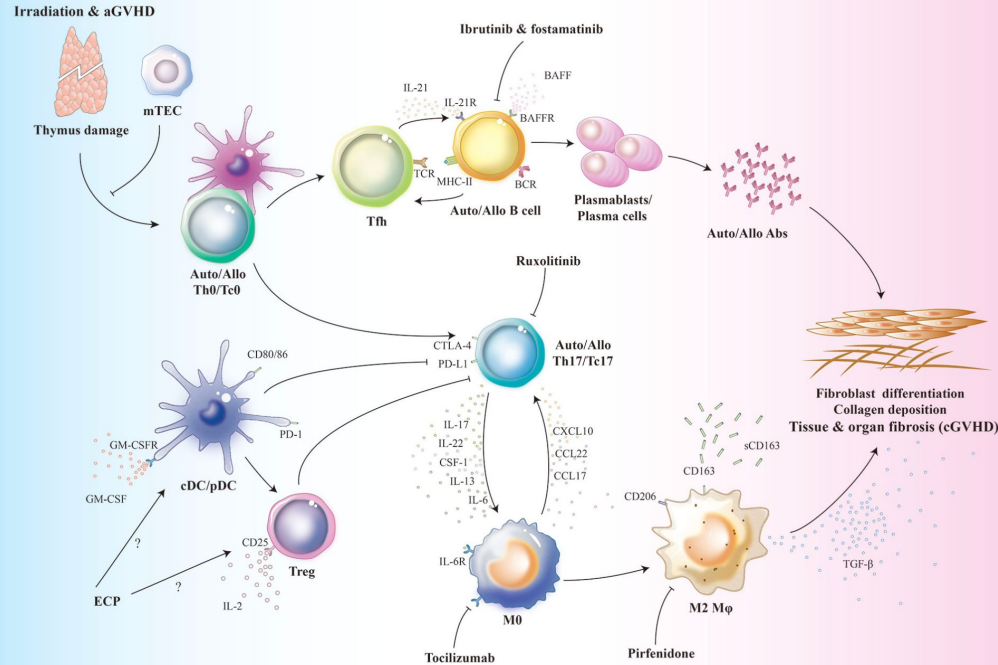


TripToFun

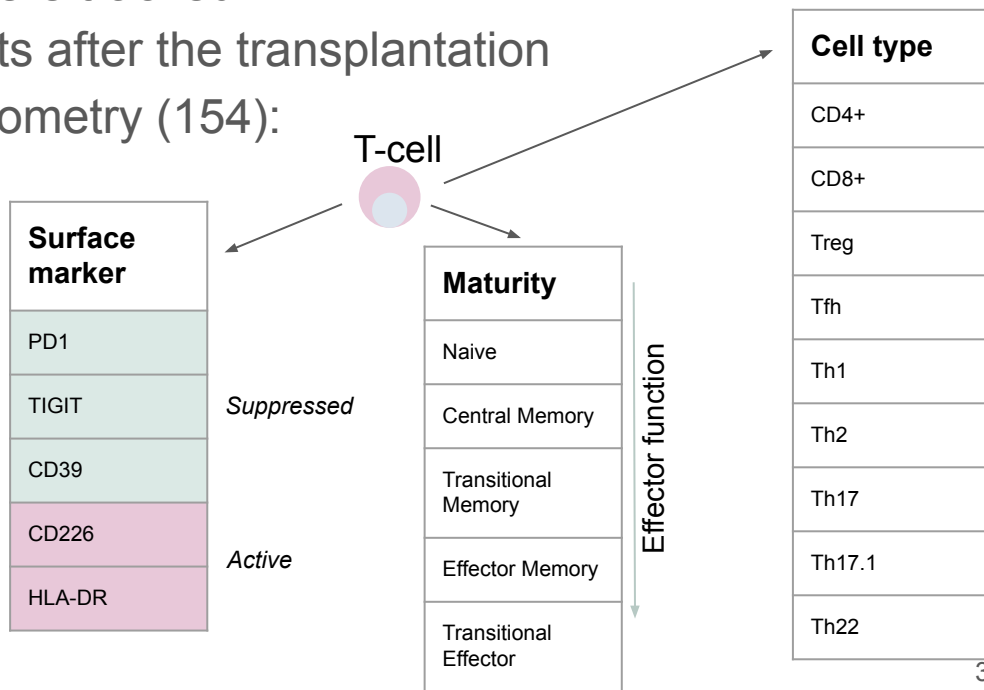
Elena Ocheredko
Nadezhda Lukashevich
Ekaterina Nesterenko
Anna Shchetsova

Analysis of T-cells immunological landscape and its association with graft-versus-host disease

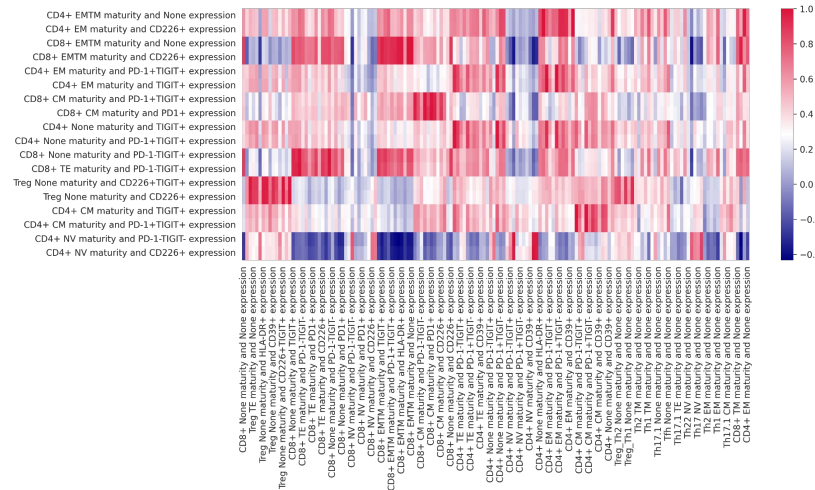


Data

- 67 patients (B-ALL, T-ALL, AML, MPAL), allo-HSCT
- The occurrence of chronic GVHD were tracked
- Blood sampling at multiple timepoints after the transplantation
- T-cell features assessed by flow cytometry (154):



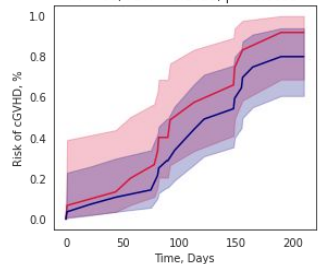
Remove non-uniform data



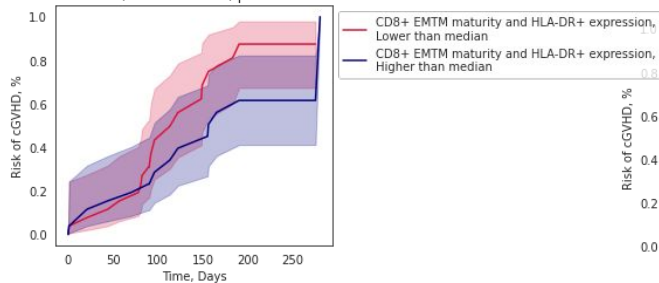
4

Results: Individual T-cell populations are associated with GVHD onset on day 90

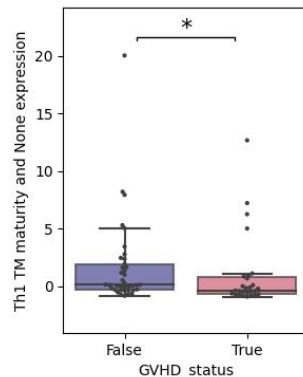
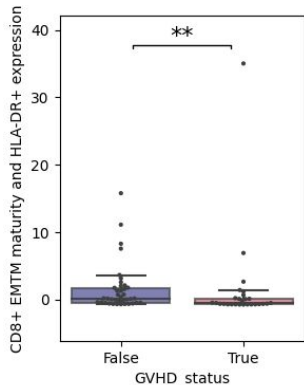
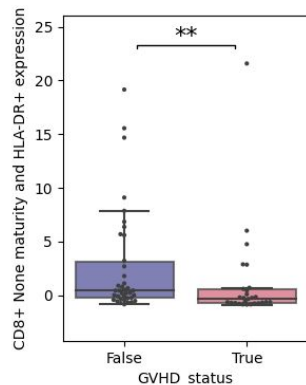
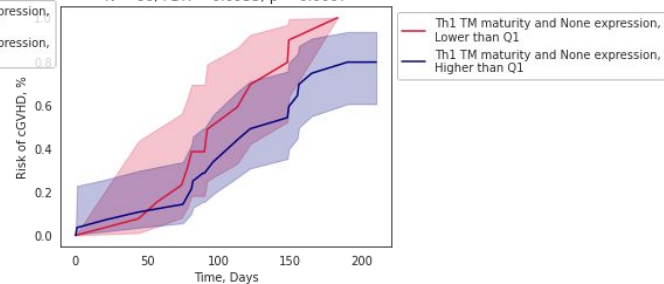
CD8+ None maturity and HLA-DR+ expression,
N = 66, FDR = 0.1421, p = 0.0053



CD8+ EMTM maturity and HLA-DR+ expression,
N = 66, FDR = 0.2649, p = 0.0058



Th1 TM maturity and None expression,
N = 66, FDR = 0.0933, p = 0.0007

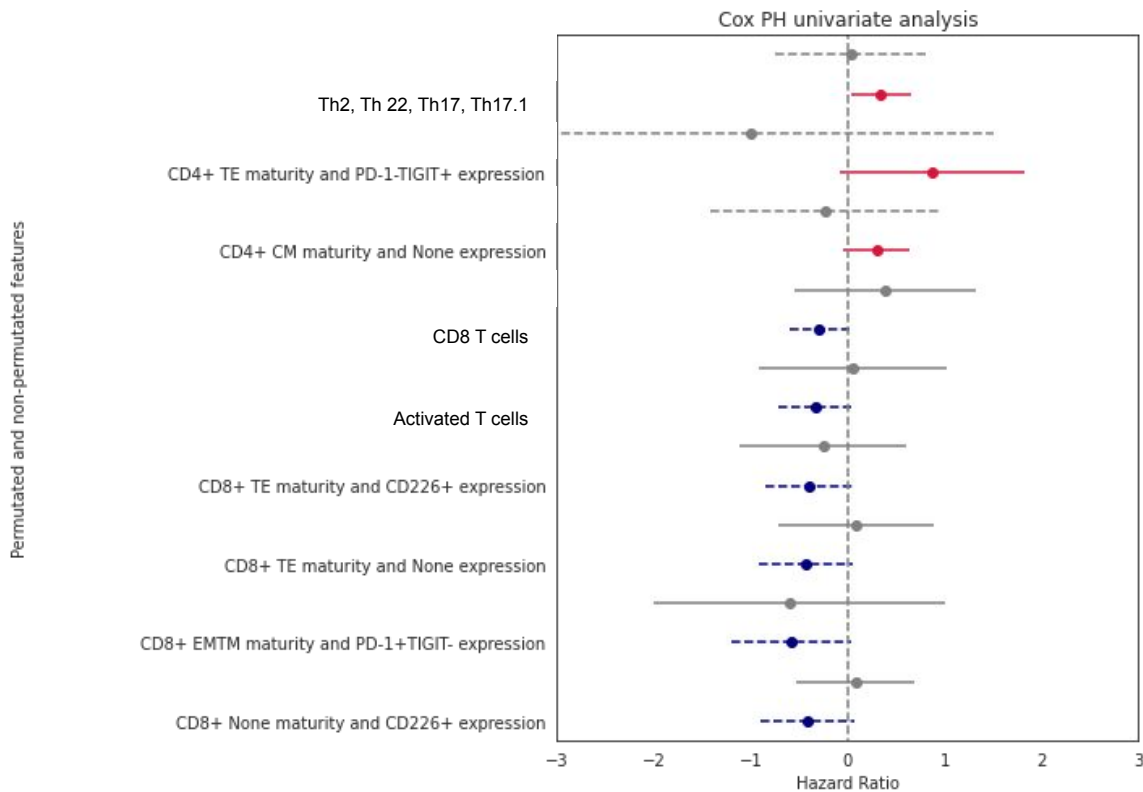


GVHD status:

True - patient developed cGVHD after day 90

False - patient does not have cGVHD after day 90

Results: Individual T-cell populations are associated both with time to GVHD and GVHD onset on day 90



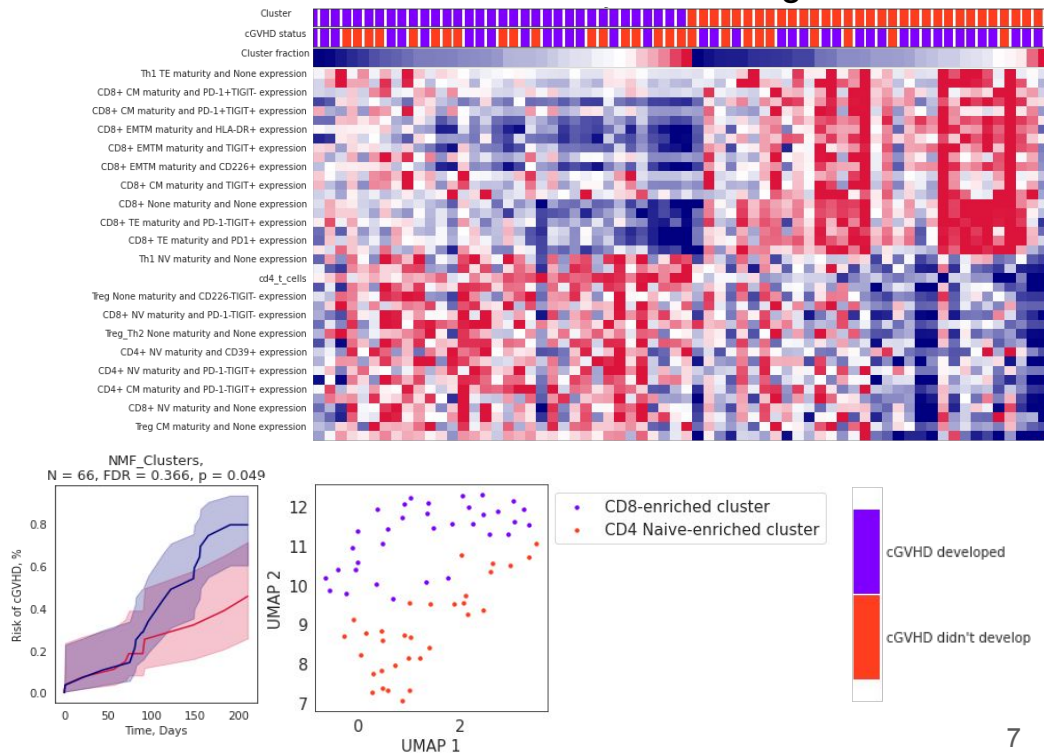
- Higher CD4+ TE maturity PD1- TIGIT+ levels associated with associated with cGVHD onset
- Higher levels of CD8+ cells with different stages of maturity associated with with lack of cGVHD

Results: GVHD onset predictive models on day 90

Leiden clustering



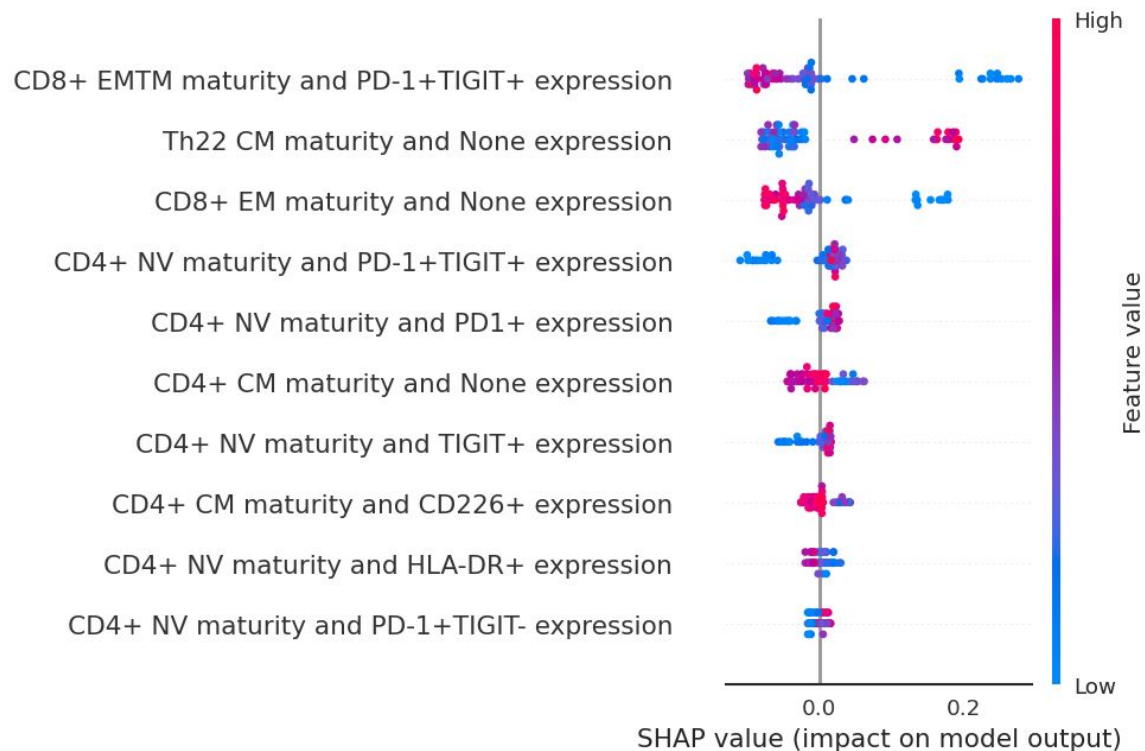
NMF clustering



Results: GVHD onset predictive models on day 90

Accuracy	AUC	Recall	Prec.	F1	Kappa	MCC	TT (Sec)
Random Forest Classifier	0.7525	0.8045	0.7063	0.7321	0.7063	0.4948	0.5087
K Neighbors Classifier	0.7145	0.7304	0.619	0.75	0.6116	0.3983	0.4457
Extreme Gradient Boosting	0.712	0.8063	0.7063	0.6528	0.6723	0.4176	0.425
CatBoost Classifier	0.712	0.7738	0.6111	0.7083	0.6344	0.4025	0.4257
Linear Discriminant Analysis	0.6532	0.6519	0.6508	0.5741	0.6083	0.2977	0.3008
SVM - Linear Kernel	0.652	0	0.2857	0.5238	0.3386	0.186	0.2248
Gradient Boosting Classifier	0.6311	0.6989	0.5556	0.5833	0.5501	0.242	0.2551
Extra Trees Classifier	0.6311	0.7272	0.5079	0.5833	0.5261	0.2326	0.2457
Naive Bayes	0.6299	0.7325	0.6111	0.5563	0.5554	0.2603	0.2769
Decision Tree Classifier	0.6299	0.6204	0.5556	0.6481	0.5556	0.2484	0.2806
Ridge Classifier	0.6127	0	0.5476	0.5333	0.5374	0.2011	0.2028
Logistic Regression	0.6115	0.731	0.5556	0.5238	0.5348	0.2061	0.2091
Ada Boost Classifier	0.6103	0.6458	0.5	0.5595	0.5134	0.191	0.2016
Light Gradient Boosting Machine	0.5919	0.5	0	0	0	0	0
Dummy Classifier	0.5919	0.5	0	0	0	0	0

Results: GVHD onset predictive models on day 90



Conclusions

- CD8+ T-cells (any maturity, predominantly active) are associated with better outcomes after allo-HSCT transplantation
- CD4+ T-cells (low effector function) are associated with chronic GVHD onset

CD8 T Cell



CD4 T Cell



Acknowledgements



MIKHAIL Y. DROKOV

Head of the Hemoblastosis Chemotherapy, Hematopoietic Depression and Bone Marrow Transplantation Research Sector
(NATIONAL MEDICAL RESEARCH CENTER FOR HEMATOLOGY)









BostonGene

Project GitHub:

https://github.com/onion-42/cGVHD_T_cell_populations_BioHackathon_2023

Feature engineering

Ranks	1	2	3	4	5	6
	CD8 T Cell	CD8 T Cell	CD4 T Cell	CD8 T Cell	CD4 T Cell	CD4 T Cell
						

$$\text{CD8 T cells} = \frac{\text{ranks of CD8 T}}{\text{ranks of CD4 T}} * \alpha = \frac{1+2+4}{3+5+6} * \alpha = \text{score}$$