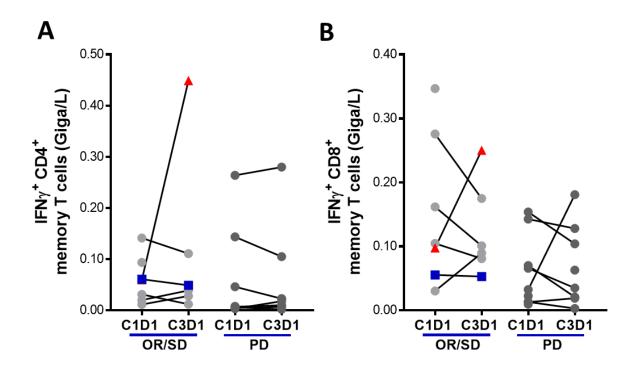


Supplementary Figure S1: At inclusion, patients present blood T and B cell subsets lymphopenia

At inclusion time, absolute number of total Lymphocytes based on clinical blood counts (**A**), total CD4⁺ (**B**), total CD8⁺ (**C**), memory CD4⁺ (**D**) and CD8⁺ (**E**) T cells, B cells (**F**) and Treg cells (**G**) based on FACS analysis for healthy donors (HD), patients who presented a stable disease or responded (OR/SD) or progressed (PD) during treatment. Patients who responded to treatment were highlighted as blue square for partial responder (PR) (**02-003**) and red triangle for complete responder (CR) (**01-005**). Statistical analysis: one-way ANOVA (* p<0,05; *** p<0,005; *** p<0.001, **** p<10⁻⁴).



Supplementary Figure S2: Evolution of the absolute numbers of IFN γ -producing CD4 $^+$ and CD8 $^+$ T cells

Quantification of absolute number of memory CD4⁺ ($\bf A$) and CD8⁺ ($\bf B$) T cells that produce IFN γ after PMA ionomycin short term reactivation.

Supplementary Table 1 – Adverse Events

No. (%) of patients with at least one	All	Grade ≥2	Grade ≥3	Grade ≥ 4
	Grades			
AE Related to Pembrolizumab	14 (70.0%)	4	3	0
AE Related to Cyclophosphamide	16 (80.0%)	11	8	1
Serious AE	3 (10.0%)	1	2	0
IrAE	0 (0.0%)	-	-	-
Grade ≥ 2 related AE (Preferred term)				
Lymphocytes count decreased		11	5	1*
PAL increase		2	1	0
Abdominal pain		1	1	0
Fatigue		1	0	0
ASAT increase		1	1	0
GGT increase		1	1	0
Cystitis		1	0	0
IRR		1	0	0

AE: adverse events, irAE: immune-related AE, PAL: Phosphatase alkaline, ASAT: aspartate aminotransferase, IRR: infusion related reactions, related: to at least one study drug, * Severe toxicity reported during the safety run