

Suppl. Material and Method

EDTA-whole blood samples were collected at Cycle 1, Cycle 5, Cycle 9 and Cycle 13 (in Part B only) prior to treatment (therefore two weeks after the previous dosing) and EOT. Direct blood staining with BD Multitest™ 6-Color TBNK (FITC-labeled CD3, clone SK7; PE-labeled CD16, clone B73.1, and PE-labeled CD56, clone NCAM16.2; PerCP-Cy5.5-labeled CD45, clone 2D1; PE-Cy7-labeled CD4, clone SK3; APC-labeled CD19, clone SJ25C1 and APC-Cy7-labeled CD8, clone SK1) and with BD Multitest™ CD8-FITC (clone SK1)/CD38-PE (clone HB7)/CD3-PerCP (clone SK7)/HLA-DR-APC (clone L343) was performed to enumerate *ex vivo* the absolute counts of T and NK cells populations and activated T CD3⁺CD8⁺ or CD3⁺CD8⁻ cells co-expressing HLA-DR and CD38 in BD Trucount™ tubes with a Lyse/No Wash procedure, avoiding any bias induced by centrifugation steps. Cells were acquired on BD FACSCanto™ II flow cytometer. The paired non-parametric Wilcoxon signed rank test was used to compare the immuno-monitoring results obtained at the different time points to the pre-treatment baseline using JMP® version 12 software.

Suppl. Figure 1 legend

Suppl. Figure 1: Combined treatment effect of T/NK cells compartment.

Absolute counts of circulating CD45⁺ CD3⁺ (T cell), CD45⁺ CD3⁺ CD4⁺ (CD4 T cell), CD45⁺ CD3⁺ CD8⁺ (CD8 T cell), CD3⁺ CD8⁺ HLA-DR⁺ CD38⁺ (Activated CD8 T cells), CD3⁺ CD8⁻ HLA-DR⁺ CD38⁺ (Activated CD4 T cells) in blood samples collected pre-dosing at the start of combined therapy (Cycle 1), at Cycle 5, Cycle 9 and Cycle 13 (for part B only) are shown for each dosing group in Part A and in Part B (top panels). Relative fold changes from the baseline calculated for all cell population are shown (bottom panels) Significant changes assessed by paired non-parametric Wilcoxon signed rank test are displayed with a horizontal bar ($p < 0.05$).