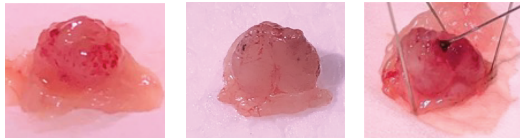


Supplementary Figure 1

A



UPFL2

UPFL6

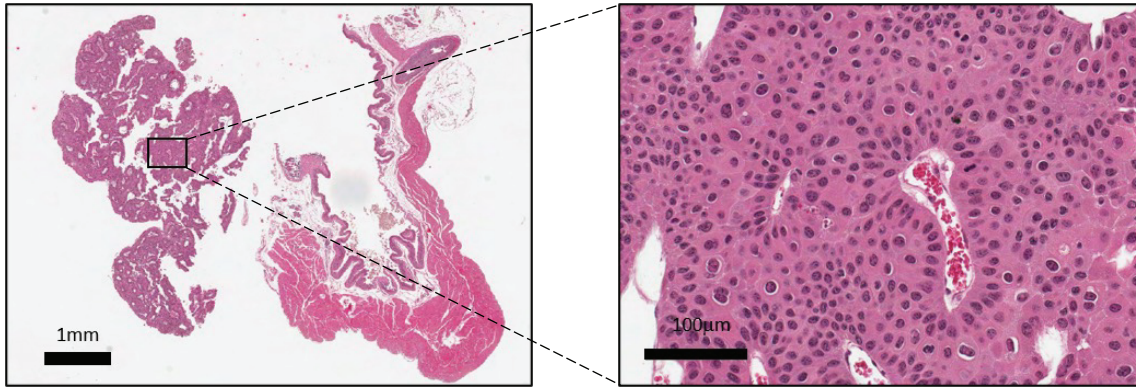
UPFL7

Representative images of UPFL tumors.

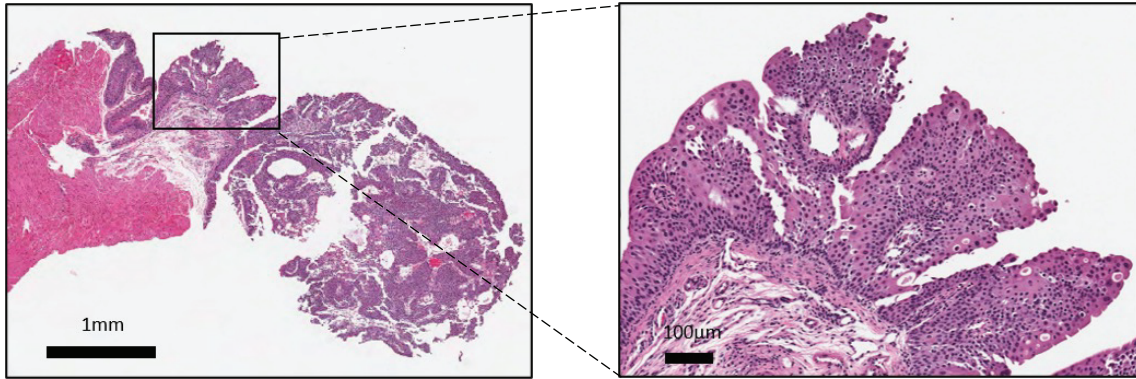
(A) Gross images of UPFL tumor bearing bladders.

(B-E) Photomicrographs of four (4) representative H&E stained UPFL bladder tumors. Each tumor is shown at low magnification with an inset box indicating the area of the related high magnification image.

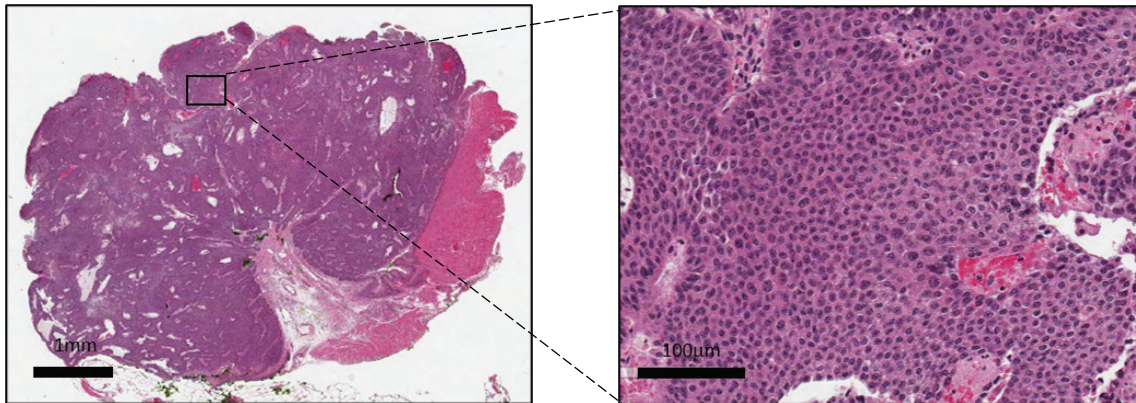
B



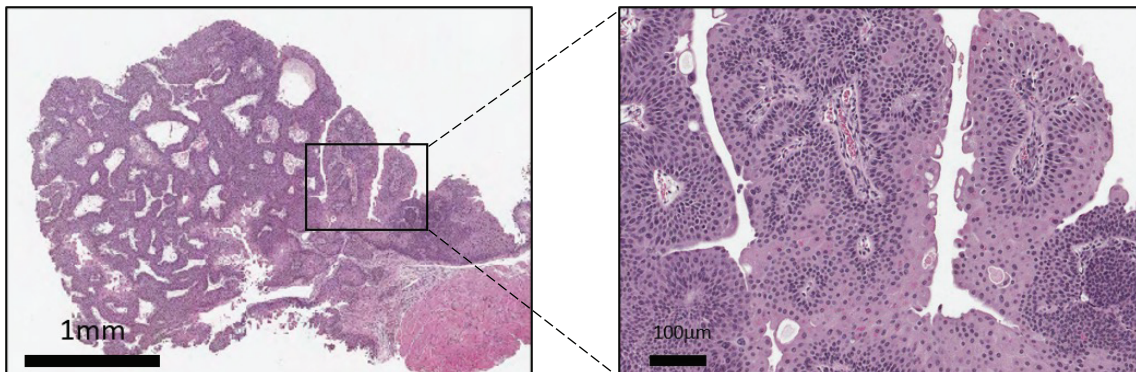
C



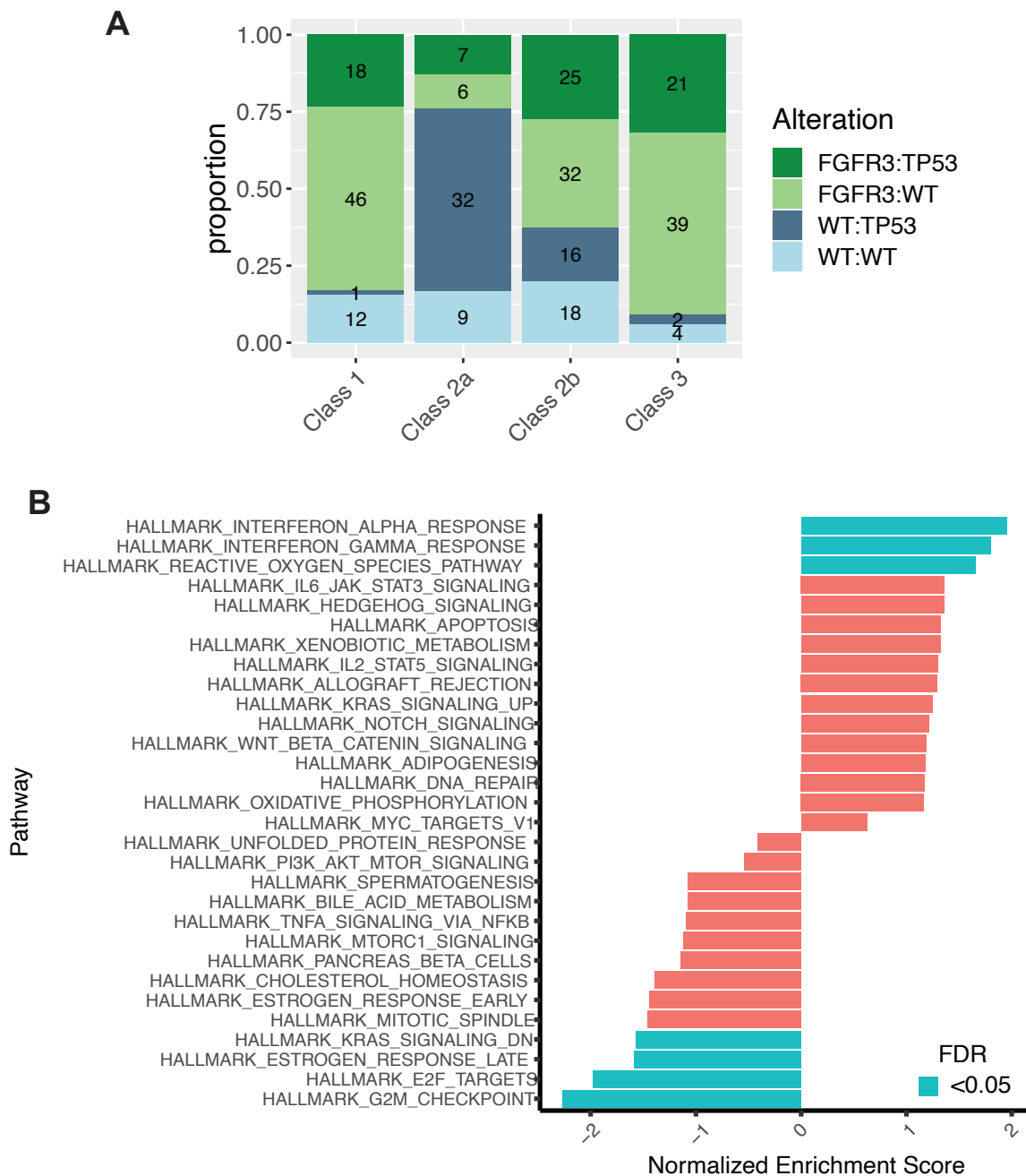
D



E



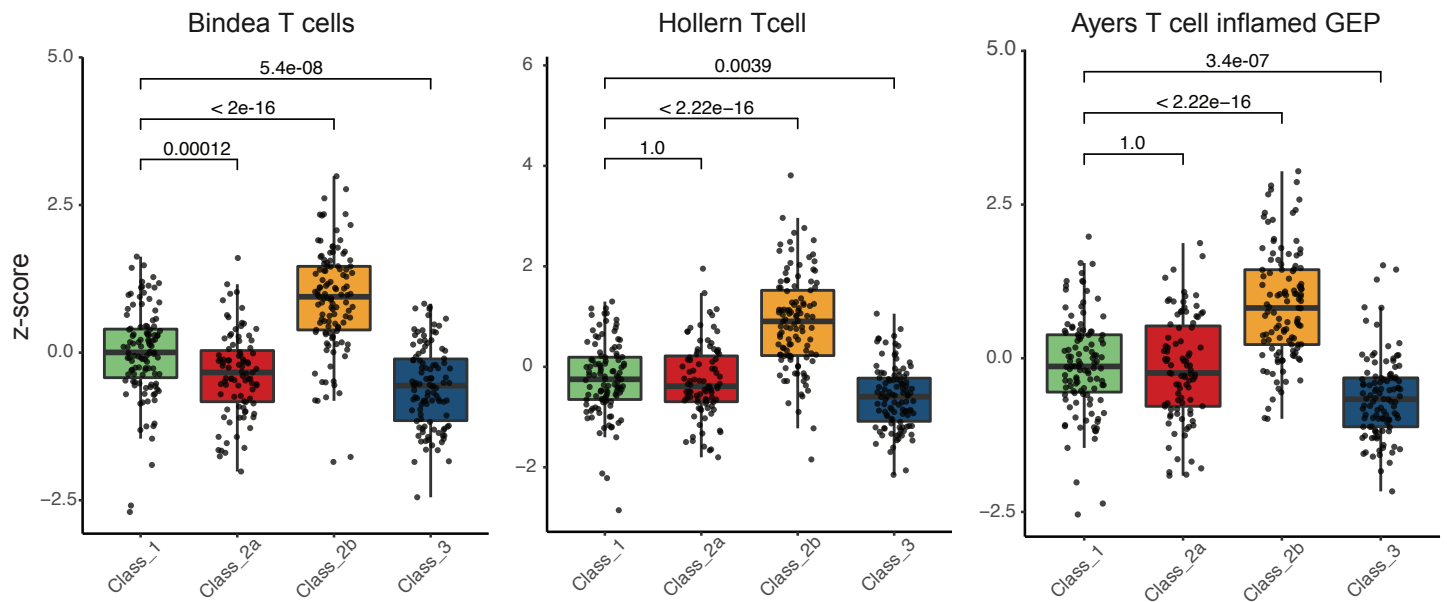
Supplementary Figure 2



Genomic and transcriptomic features of UPFL tumors - (A)

Co-occurrence rates of *FGFR3* DNA calling based mutations and RNA based *TP53* Pathway alterations for the UROMOL cohort. (B) Gene set enrichment analysis (GSEA) for the Hallmark expression signatures were performed on the log₂ - fold change between UPFL and UPPL tumors. The top ranking signatures by normalized enrichments score for UPFL and UPPL were plotted. Signature with an FDR < 0.05 are indicated by teal bars.

Supplementary Figure 3

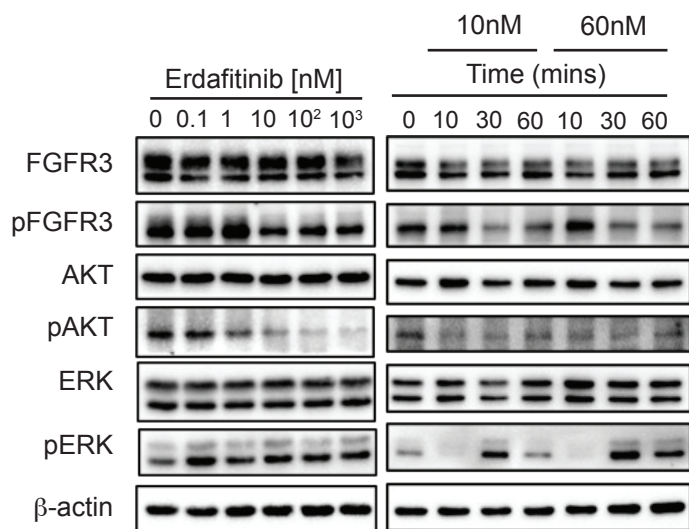


T cell signature immune gene signatures scores.

Signatures representing T cells were calculated for the UROMOL cohort and plotted by NMIBC class. All boxplots are represented by the IQR and midline at the median. Error bars represent $Q1/Q3 \pm 1.5 \cdot IQR$.

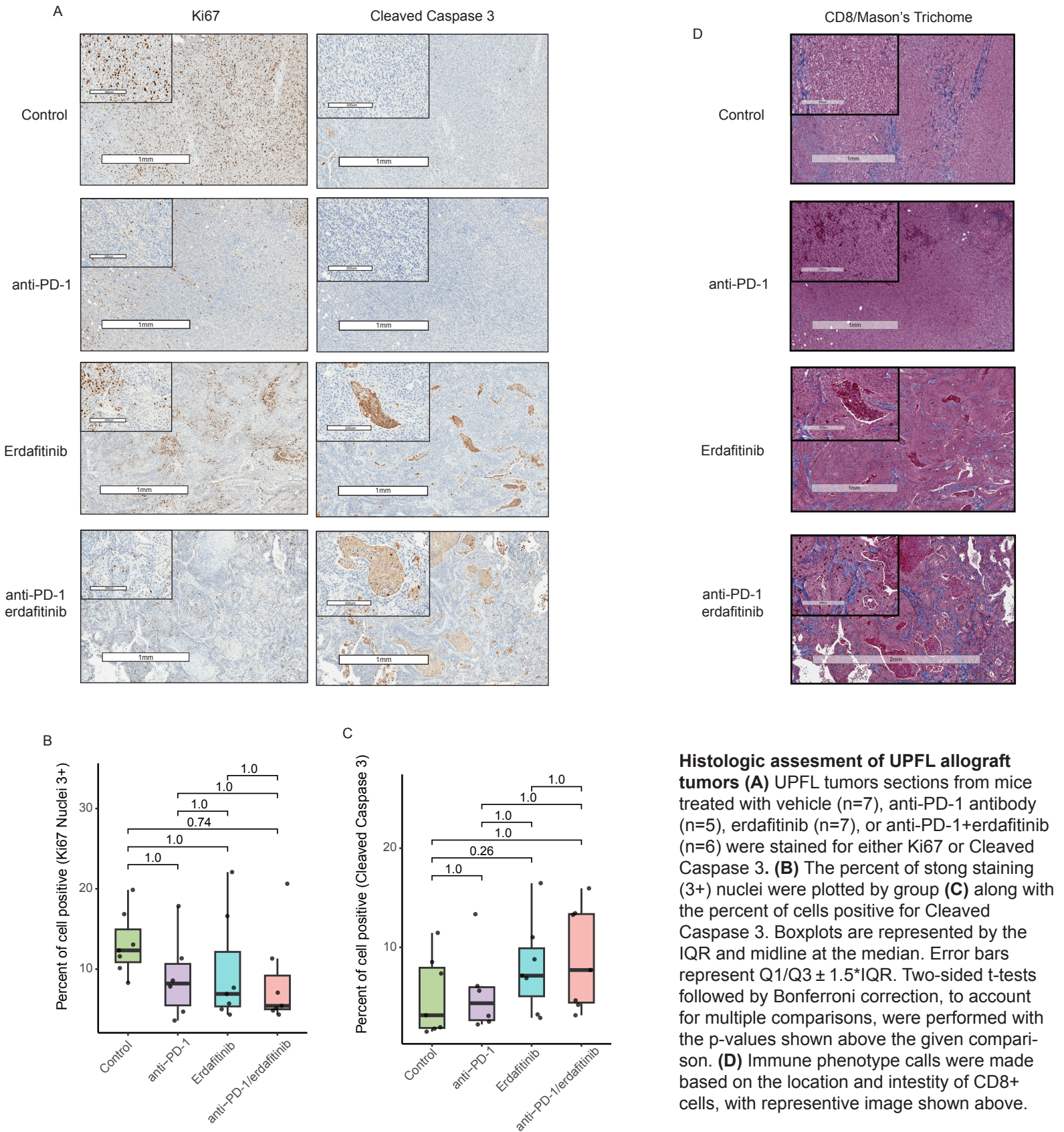
Two-sided t-tests followed by Bonferroni correction, to account for multiple comparisons, were performed with the p-values shown above the given comparison.

Supplementary Figure 4

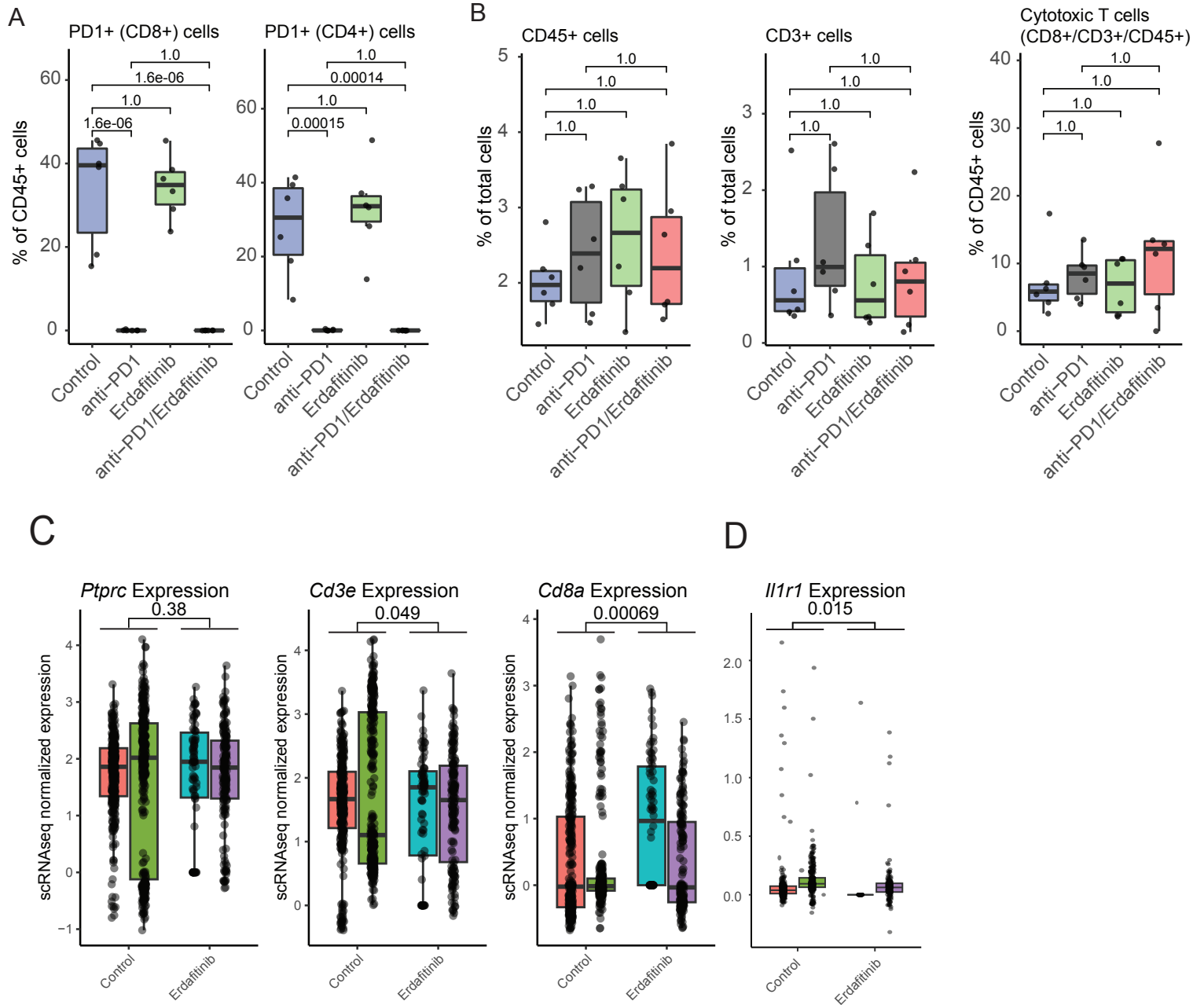


UPFL derived cell line (UPFL3) is sensitive to FGFR inhibition - Immunoblots with the indicated antibodies in whole cell extracts of UPFL3 cells treated with erdafitinib for the indicated dose and time.

Supplementary Figure 5

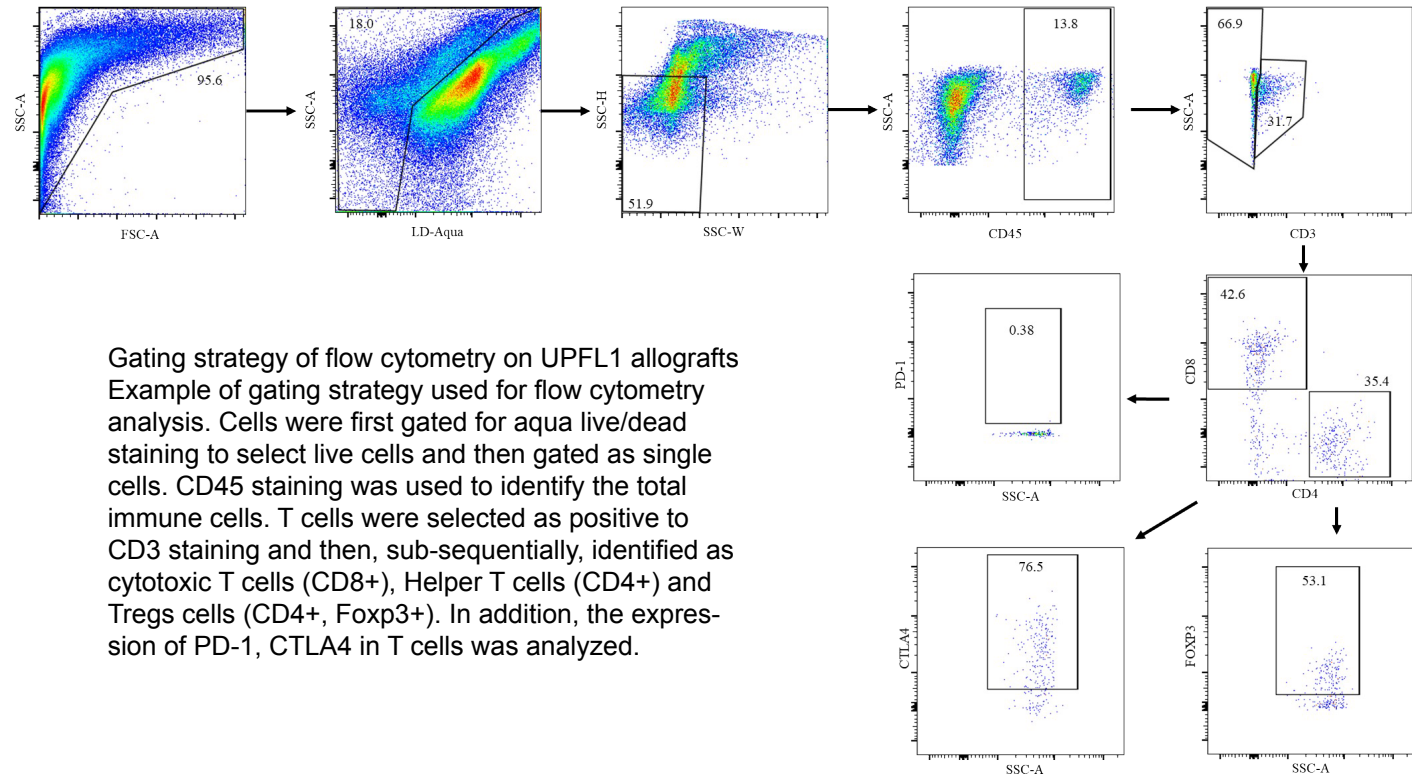


Supplementary Figure 6



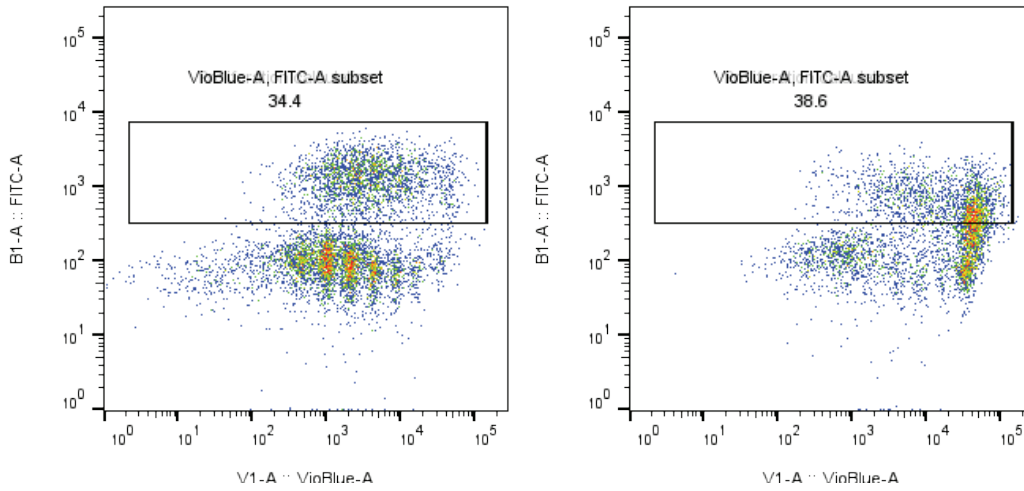
Supplementary Figure 6 (A) Box plots of percentage of CD45+ cells that are PD1+ in CD8+ (left) and CD4+ (right) for the indicated treatment group as well as **(B)** CD45+, CD3+, and CD8+ cytotoxic T cells after 1 week of treatment. **(C)** The T cell subset of cells identified by SingleR from control or erdaftitinib treated tumor were plotted by the scRNA expression values for *Ptprc* (CD45), *Cd3e* (CD3), and *Cd8a* (CD8) **(D)** *Il1r1*. All boxplots are represented by the IQR and midline at the median. Error bars represent Q1/Q3 \pm 1.5*IQR. Two-sided t-tests followed by Bonferroni correction, to account for multiple comparisons, were performed with the p-values shown above the given comparison.

Supplementary Figure 7

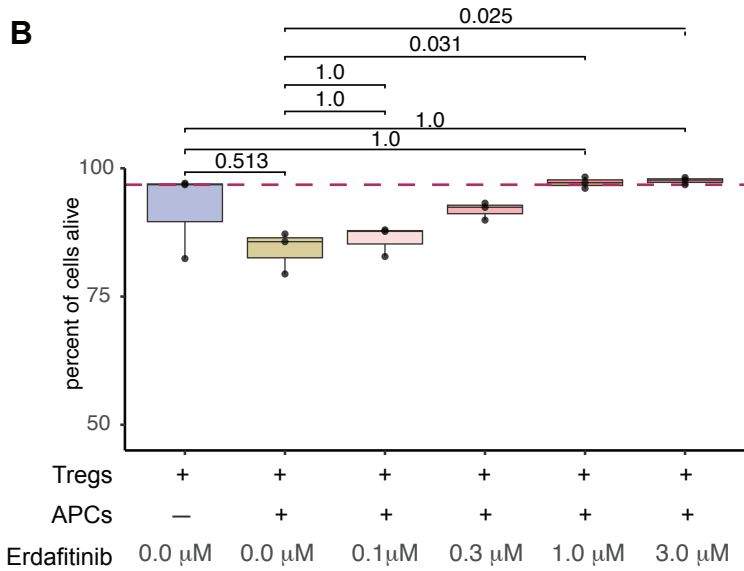


Supplementary Figure 8

A



B



Gating strategy and quantification for the isolation for Treg proliferation assay.

(A) FACS analysis was performed on the CD4⁺/Foxp3-GFP⁺ population. Cells positive for crystal violet tracer (CVT) were identified as the non-proliferative population, whereas CVT⁺ cells were considered proliferative. **(B)** Boxplot of the percent living cell Tregs following the 3 day proliferation assay, regardless of CVT status. **(C)** RNA expression of FGFR1, FGFR2, FGFR3 in T cell subsets. All boxplots are represented by the IQR and midline at the median. Error bars represent Q1/Q3 ± 1.5*IQR. Two-sided t-tests followed by Bonferroni correction, to account for multiple comparisons, were performed with the p-values shown above the given comparison.

C

FGFR family expression by immune cell type (Schmiedel et. al.)

