## **Description of Additional Supplementary Files**

Supplementary Data 1: DGE on cell lines

A: Results of DGE between cell lines untreated and treated with MEKi for 2 days

B: Genes of the eMEKi signature

Supplementary Data 2: Cell lines subtyping

A: Subtyping classification, according to Moffitt's classification, of CCLE and in house cell lines

B: Subtyping classification, according to Moffitt's classification, of In house cell lines, vehicle and MEKi treated

C: Subtyping classification, according to Moffitt's classification, of PanCuRx samples

D: Subtyping classification, according to Moffitt's classification, of ICGC samples

E: Subtyping classification, according to Moffitt's classification, of TCGA samples

Supplementary Data 3: Annotation of mouse fibroblast clusters A: Differential Gene expression and Enrichment analysis for cluster 0 B: Differential Gene expression and Enrichment analysis for cluster 1 C: Differential Gene expression and Enrichment analysis for cluster 2 D: Differential Gene expression and Enrichment analysis for cluster 3 E: Differential Gene expression and Enrichment analysis for cluster 4 F: Differential Gene expression and Enrichment analysis for cluster 5 G: Differential Gene expression and Enrichment analysis for cluster 6 H: Differential Gene expression and Enrichment analysis for cluster 7 I: Differential Gene expression and Enrichment analysis for cluster 7 L: Differential Gene expression and Enrichment analysis for cluster 9

Supplementary Data 4: Ligand-receptor analysis

A: Ligand-receptor pair data between Epithelial -> Fibroblasts in vehicle and MEKi

B: Ligand-receptor pair data between Fibroblast -> Epithelial cells in vehicle and MEKi

Supplementary Data 5: DGE on fibroblasts cluster from in house sc-RNAseq

A: Results of DGE between cells of the fibroblast cluster from sc-RNAseq data of mice treated either with vehicle or MEKi for 2 days

B: Genes of the sMEKi signature

C: Results of DGE between mapCAFs and other cells of the fibroblast cluster from sc-RNAseq data of vehicle-treated mice

D: Genes of the mouse mapCAF signature

Supplementary Data 6: Annotation of human fibroblast clusters

A: Differential Gene expression and Enrichment analysis (padj<0.05) for cluster 0 B: Differential Gene expression and Enrichment analysis (padj<0.05) for cluster 1 C: Differential Gene expression and Enrichment analysis (padj<0.05) for cluster 2 D: Differential Gene expression and Enrichment analysis (padj<0.05) for cluster 3 E: Differential Gene expression and Enrichment analysis (padj<0.05) for cluster 4 F: Differential Gene expression and Enrichment analysis (padj<0.05) for cluster 5 G: Differential Gene expression and Enrichment analysis (padj<0.05) for cluster 6 H: Differential Gene expression and Enrichment analysis (padj<0.05) for cluster 7 I: Differential Gene expression and Enrichment analysis (padj<0.05) for cluster 7 Supplementary Data 7: Annotation of human fibroblast clusters

A: Results of DGE between mapCAFs and other cells of the fibroblast cluster from human sc-RNAseq data

B: Genes of the human mapCAF signature