

Supplementary Figure 1. Correlation of epithelial seam degeneration, cell death, and *Tgfb3* and *Mmp13* expression in E14.5 mice. (**A-C**) Histological analysis of wild-type embryos reveals a fully formed epithelial seam which is beginning to degenerate (**A**). Similarly, in *Irf6*^{+/R84C};*Jag2*^{+/ Δ DSL} embryos, regions of epithelial degeneration between the palate and the tongue are apparent (arrowed in **B**). Further posteriorly, epithelial degeneration is more extensive and epithelial triangles are present (arrowed in **C**). (**D-F**) Immunofluorescence analysis for activated caspase 3 in adjacent sections to those depicted in A, B and C, respectively reveals extensive cell death in the midline epithelial seam of wild-type embryos (**D**) and in the region of epithelial degeneration in *Irf6*^{+/R84C};*Jag2*^{+/ Δ DSL} littermates (arrowed in **E** and **F**). (**G-L**) At a slightly earlier time-point than shown in **A-F**, when the epithelia are still intact, *Tgfb3* and *Mmp13* are expressed in the midline epithelial seam of wild-type embryos, *Tgfb3* and *Mmp13* are expressed in regions of presumptive epithelial degeneration (arrowed in **H,I,K,L**). p, palate; t, tongue. Scale bars: 100 µm.