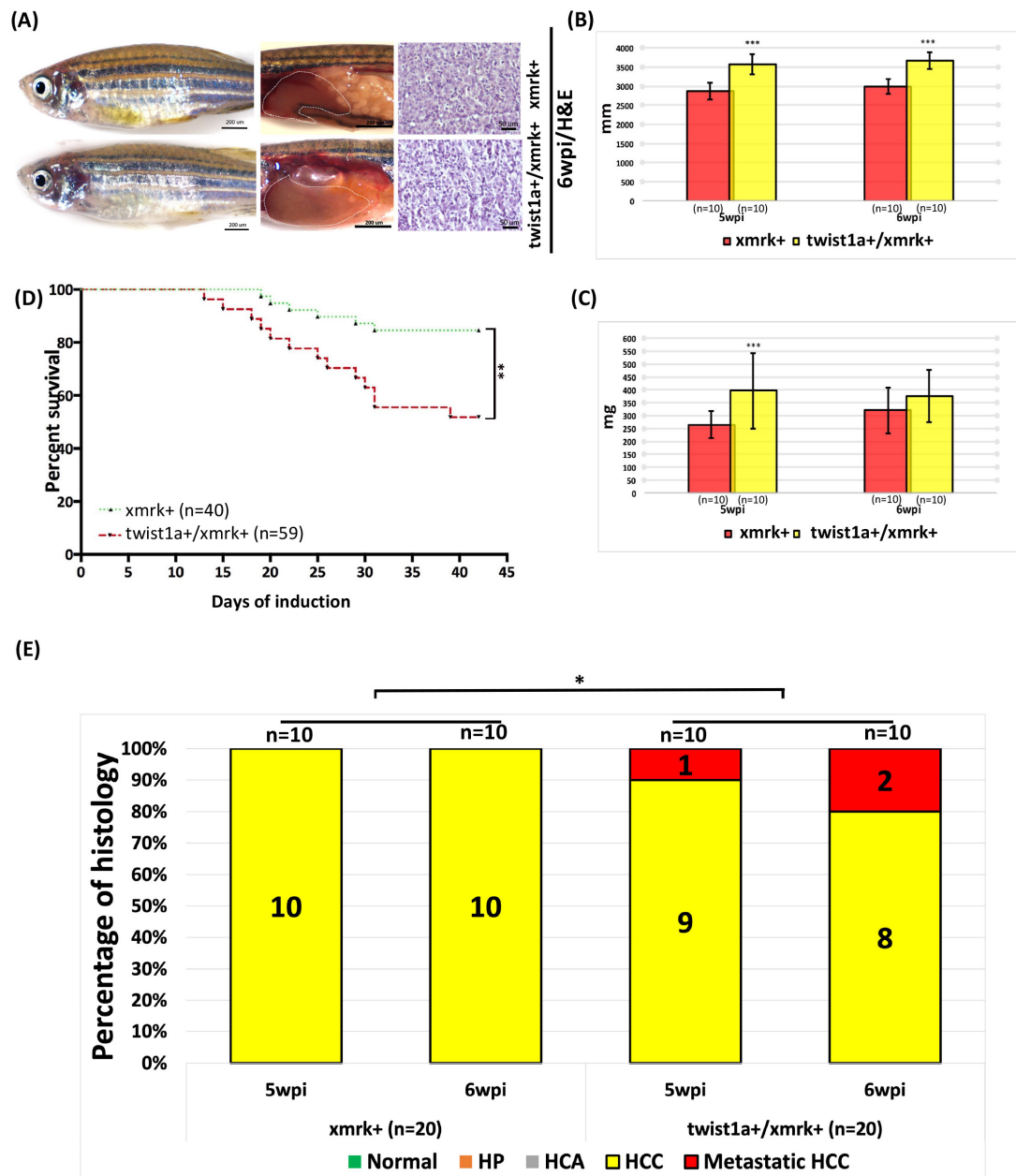


**Figure S1.** Fluorescent images of *twist1a*<sup>+</sup> transgenic zebrafish and a timeline of experimental design with the schedule used for specimen collection. **(A)** Fluorescent images of *twist1a*<sup>+</sup> transgenic larva at 7 dpf and adult zebrafish at 5 mpf. The red denotes fluorescent *twist1a*-expressing hepatocytes in the liver. **(B)** Experiment design and specimen collection schedule for long-term treatment.



**Figure S2.** Liver tumor status and metastasis in *xmrk*<sup>+</sup> and *twist1a*<sup>+/xmrk</sup> transgenic zebrafish: (A) *Twist1a*<sup>+/xmrk</sup> and *xmrk*<sup>+</sup> control transgenic zebrafish were treated with 60 µg/ml Dox for three weeks, at which point the dose was reduced to 30 µg/ml Dox for a further three weeks. Note that 1 µg/ml 4-OHT was maintained throughout the course of the experiment to ensure the sustained induction of *twist1a*. Sampling was performed at 5 and 6 wpi. The left column displays the external appearance, the middle column shows internal abdominal organs with the livers outlined, and the right column depicts H&E staining of liver sections. Scale bar: 50 or 200 µm. In the *twist1a*<sup>+/xmrk</sup>

transgenic zebrafish, we observed significant differences in **(B)** body length and **(C)** body weight at 5 and 6 wpi, compared with the *xmrk*<sup>+</sup> group at the same timepoints. **(D)** Kaplan-Meier survival curves showing days post-induction plotted against percentage survival to 6 wpi. **(E)** Histological examination confirmed that *twist1a*<sup>+</sup>/*xmrk*<sup>+</sup> transgenic zebrafish developed HCC or metastatic HCC at 5 and 6 wpi at an incidence exceeding that of *xmrk*<sup>+</sup> siblings. Differences among the variables were assessed using Student's t-tests or one-way ANOVA. Statistical significance: \*P<0.05, \*\*P<0.01, \*\*\*P<0.001.