Figure S1



Figure S1. Immunohistochemistry (IHC) analysis of H-CAF density and the expressions of Ecadherin and p-STAT3 in HCC patients. (A) E-cadherin was lowly or moderately expressed in the cancer cell cytoplasm, whereas it was highly expressed in well-differentiated or normal tissue (original magnification, $100 \times$). An enlarged view is shown in the lower panel (original magnification, $400 \times$). (B) p-STAT3 was highly or moderately expressed in the cancer cell nucleus, whereas it was lowly expressed in well differentiated or normal tissue (original magnification, $100 \times$). An enlarged view is displayed in the lower panel (original magnification, $400 \times$). (C) A high density of H-CAFs in the samples was associated with low E-cadherin expression, whereas a low density of H-CAFs was associated with high E-cadherin expression (original magnification, $100 \times$). The lower panel shows the enlarged view (original magnification, $400 \times$). (D) IHC scores of E-cadherin were compared in the subgroups with dichotomized H-CAFs density (low VS moderate and high). (E) Correlation of H-CAFs density with p-STAT3 expression. (The p-STAT3 and E-cadherin expression, brown color, green row; H-CAFs, unstained, red row). (F) IHC scores of p-STAT3 were compared in the subgroups of H-CAFs density.



Figure S2. (A) Western blotting and (B) transwell analysis of Hep3B, treated with H-CAFs^{low} or H-CAFs^{moderate/high}, were performed.



Figure S3. TG2 expression was tested in MHCC-97L and Huh7 cell lines by western blotting.



Figure S4. Overexpression of TG2 induced EMT of HCC cells. The error bars represent \pm SEM; * P < 0.05, *** P < 0.001). (A) TG2 was stably overexpressed by CMV promoter in Huh7 cells. An empty vector was used as a control. N-cadherin and E-cadherin were used as indicators of EMT initiation. (B) Wound healing, (C) transwell, and (D) invasive assays were performed using Huh7-Vec and TG2 OE cells. (E) In the nude mouse metastatic tumor model, Huh7-Vec or TG2 OE cells were injected into the spleen of nude mice. After 21 days, the tumor of liver metastases was counted and then confirmed by H&E staining (F). Original magnification, $50 \times$.