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To Examine the Role of Caveolin-1 in Taiwanese Hepatocellular Carcinoma at DNA, RNA, and Protein Levels

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Backgrounds: Hepatocellular carcinoma (HCC) is one of the most common types of malignant tumor worldwide, for which the prevalence and mortality rates are very high in Taiwan, Caveolin-1 (CAV-1) is a main structural protein of caveolae and plays a regulatory role in signaling pathways and tumorigenesis. High expression of Cav-1 in mouse HCC is positively correlated with higher cell invasive capacity, but the contribution of CAV-1 genetic variants during HCC progression is still largely unknown. In this study, we investigated the contribution of CAV-1 variant to the risk of HCC from the analyses of DNA, RNA and protein. Materials and Methods: We enrolled 298 patients with HCC patients and 298 cancer-free controls frequency matched by age and gender in this case-control study. Firstly, the associations of six single nucleotide polymorphisms (SNPs) of the Cav-1 gene at C521A (rs1997623), G14713A (rs3807987), G21985A (12672038), T28608A (rs3757733), T29107A (rs7804372), and G32124A (rs3807992) with HCC risk in a Taiwanese population were evaluated. Secondly, thirty HCC tissue samples with variant genotypes were tested to estimate CAV-1 mRNA expression by real-time quantitative reverse transcription. Finally, the HCC tissue samples of variant genotypes were examined by western blotting to estimate their CAV-1 protein expression patterns. Results: There were significant differences between the HCC and control groups in the distributions of the CAV-1 G14713A genotypes (p=0.0124), and these carrying AG and AA genotypes had a higher risk for HCC, compared with those with the GG genotype (odds ratio=1.51 and 1.94, respectively). Patients with CAV-1 G14713A AG or AA genotype had higher levels of mRNA (p=0.0001) and protein (p=0.0019) than those with GG genotype. Conclusion: Our multi-approach findings at the DNA, RNA and protein levels suggest that CAV-1 may play a critical role in HCC carcinogenesis, and serve as a target for HCC therapy.

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