



# Innovative Strategies for Liver Cancer: Ribozyme-Mediated RNA Replacement

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## DESCRIPTION

Liver cancer, particularly Hepatocellular Carcinoma (HCC), is a devastating disease with limited treatment options and a high mortality rate. Conventional therapies such as chemotherapy, radiation, and surgical resection often have limited effectiveness and are associated with significant side effects. Therefore, the development of innovative and targeted treatment strategies is significant to improve outcomes for patients with liver cancer. One potential approach is targeted suicide gene therapy, which utilizes the selective killing of cancer cells while sparing healthy cells. This therapeutic strategy involves the introduction of genes that encode enzymes capable of converting non-toxic prodrugs into cytotoxic compounds within tumor cells. This approach offers the potential for targeted and localized treatment, minimizing systemic toxicity. Ribozymes are Ribonucleic Acid (RNA) molecules with catalytic activity that can specifically cleave RNA sequences in a sequence-dependent manner.

This unique distinctive characteristics of ribozymes makes them attractive candidates for gene therapy applications. By designing ribozymes that specifically target and cleave key RNA molecules involved in cancer cell survival and proliferation, it is possible to disrupt critical pathways and induce cell death. The basic principle behind this approach is to introduce engineered ribozymes that can cleave and replace specific RNA molecules within cancer cells, thereby disrupting essential cellular processes and promoting cell death. One key advantage of ribozyme-mediated RNA replacement is its ability to target post-transcriptional regulation, which plays a significant role in cancer progression. By specifically targeting RNA molecules involved in post-transcriptional regulation, such as microRNAs or messenger RNAs, ribozymes can exert specific control over gene expression and protein production within cancer cells. In the context of liver cancer, specific microRNAs and messenger RNAs have been identified as critical regulators of key signaling pathways and cellular processes. By designing ribozymes that can specifically target and cleave miR-21, it is possible to inhibit its oncogenic function and induce cell death. To achieve the ribozyme-mediated

RNA replacement, several steps are involved. First, engineered ribozymes targeting the desired RNA molecule, such as miR-21, are designed and synthesized. These ribozymes are designed to have high specificity and efficiency in cleaving the target RNA. Next, delivery systems are utilized to introduce the ribozymes into liver cancer cells.

Various delivery methods, including viral vectors or nanoparticles, can be employed to ensure efficient and targeted delivery. Once inside the cancer cells, the ribozymes are released and can specifically recognize and cleave the target RNA molecules, such as miR-21. This cleavage disrupts the regulatory function of miR-21, leading to a loss of its oncogenic activity. In addition, the cleavage products generated by the ribozymes can further trigger cellular responses, such as apoptosis or immune-mediated cell death, contributing to the elimination of cancer cells. Furthermore, the versatility of ribozyme-mediated RNA replacement allows for the targeting of multiple RNA molecules simultaneously. By designing ribozymes that target different oncogenic microRNAs or messenger RNAs involved in liver cancer, it is possible to exert a more comprehensive and synergistic effect on tumor cells.

This multi-targeted approach may enhance the therapeutic efficacy and reduce the likelihood of resistance development. Despite the promising results, several challenges and considerations need to be addressed before ribozyme-mediated RNA replacement can be translated into clinical practice. One challenge is the efficient and targeted delivery of the ribozymes to liver cancer cells. Various delivery systems and strategies are being explored to improve delivery efficiency and minimize off-target effects.

Another consideration is the potential for immune responses against the viral vectors or delivery systems used in this therapy. Immune reactions may limit the therapeutic efficacy and pose safety concerns. Strategies to minimize immune responses, such as the use of immunomodulatory agents or modifications of the delivery systems, are being investigated to enhance the clinical applicability of ribozyme-mediated RNA replacement.

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## CONCLUSION

Targeted suicide gene therapy based on ribozyme-mediated RNA replacement through post-transcriptional regulation holds great promise for the treatment of liver cancer, particularly hepatocellular carcinoma. By specifically targeting and cleaving critical RNA molecules involved in cancer progression,

this approach offers a unique opportunity to disrupt key cellular processes and induce cancer cell death.

Further research and clinical trials are warranted to optimize delivery strategies, enhance therapeutic efficacy, and ultimately convert this innovative therapeutic approach into clinical practice for the benefit of liver cancer patients.