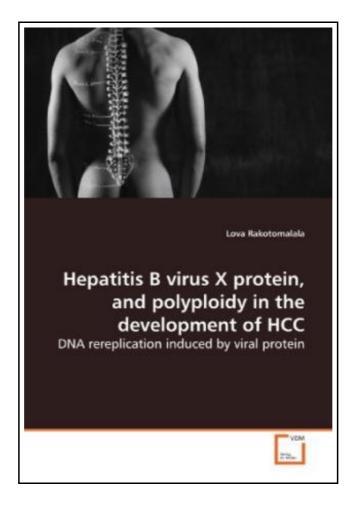
Hepatitis B virus X protein, and polyploidy in the development of HCC



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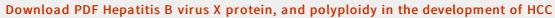
HEPATITIS B VIRUS X PROTEIN, AND POLYPLOIDY IN THE DEVELOPMENT OF HCC



Condition: New. Publisher/Verlag: VDM Verlag Dr. Müller | DNA rereplication induced by viral protein | Hepatitis B Virus (HBV) X protein (pX) is implicatedby an unknown mechanism inhepatocellular carcinoma (HCC) development, characterized by increased rate of chromosomal aberrations. Although most patients clearthe infection and developimmunity, chronic infection in 5 to 10% infected patients lead to HCC development in the 4th to 5th decade of life. I demonstrate that tetracycline-regulated pX expression induces multipolar spindles and polyploidy (4N DNA). To understand the mechanism of pX-mediated polyploidy, I investigated whether pX promotes DNA re-replication. Dual-parameter flow cytometry demonstrates pX-dependent BrdU incorporation in cells with 4N DNA. pX also induces expression of replication initiation factors Cdc6 and Cdtl, while suppressing geminin expression, an egative regulator of re-replication. G2-phase synchronized cells exhibit pX-dependent: i)nuclear Cdc6 and Mcm5 colocalization; ii) absence of nuclear geminin; iii) increased BrdUincorporation; iv) ATRactivation; v) RAD17 and H2AX phosphorylation; and vi) co-localization of -H2AX with the DNA elongation factor PCNA. | Format: Paperback | Language/Sprache: english | 200 gr | 144 pp.



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