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HCV in Tissue Culture: Yet Another Confounder

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Hepatitis C virus (HCV) is a very important human pathogen that causes chronic liver disease in a substantial number of infected individuals worldwide. Dr. Robert Purcell, who has headed the NIH's activities on hepatitis viruses, quite often with spectacular success, has been on the leading edge attempting to devise techniques for the cultivation of HCV in tissue culture. Dr. M. Yanagi and co-investigators of his group report on an interesting twist on attempts to grow HCV. This virus has had a long history of reports on such a low level

of replication that reverse transcriptionpolymerase chain reaction (RT-PCR) assays are required for detection.

In a study to refine the cultivation process, the authors discovered yet another complication. When performing the RT-PCR assay with primers deduced from conserved domains of the five prime noncoding regions of HCV, they found appropriately sized cDNA. However, subsequent analysis revealed that the cDNA detected and thought to be associated with HCV was in fact amplified from bovine viral diarrhea virus (BVDV) sequences, which were contained in the lots of commercially available fetal bovine serum (FBS) used in the tissue culture

medium. Six lots of commercially available FBS and one lot of newborn calf serum from three major suppliers in the United States were tested. All were contaminated with BVDV.

The authors recommend that, although there is no evidence of infectious virus being present, bovine sera used in HCV tests or for production of vaccine should be screened for BVDV by RT-PCR.

FROM: Yanagi M, Bukh J, Emerson S, Purcell R. Contamination of commercially available fetal bovine sera with bovine viral diarrhea virus genomes: implications for the study of hepatitis C virus in cell cultures. *J Infect Dis* 1996;174:1324-1327.