## **SCIENTIFIC LETTER**



## Triple Infection by Betaherpesviruses (HCMV, HHV-6 and HHV-7) in a Child with Acute Liver Failure

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To the Editor: We report here the first description of a triple infection by betaherpesvirus before liver transplantation. A 1y-old female patient developed acute liver failure (ALF) of unknown etiology. The patient had ALF Grade II encephalopathy and coagulopathy with an international normalized ratio (INR) of 4.2. Enzymes aspartate aminotransferase (AST) and alanine aminotransferase (ALT), alkaline phosphatase (ALP) and total Bilirubin (TB) were increased, hematologic analysis showed severe anemia. The patient had no history of hepatotoxic drugs and symptoms of childhood exanthematic disease. The patient underwent a liver transplantation and peripheral blood and liver explant samples were collected. After 47 d of transplant procedure, the patient died. The etiology investigation of the ALF was performed by polymerase chain reaction (PCR) for virus hepatitis (A, B, C, and E) and Epstein-Barr virus. The sample was positive to Pan PCR herpesviruses (DPOL gene). The infection was confirmed by quantitative PCR (qPCR) and nested PCR specific for each virus detected. The regions UL54, U56 and U37 were targeted to detect and amplify human cytomegalovirus (HCMV), human herpes virus (HHV)-6 A/B and HHV-7 respectively [1]. The triple infection was detected and confirmed by sequencing and viral load of  $4 \times 10^3$  copies/mL,  $7 \times 10^5$  copies/mL and  $1 \times 10^5$ copies/mL to HCMV, HHV-6 and HHV-7 respectively. The etiology of ALF cases is often unknown, and some of these cases are not related to classical etiologies. The HCMV activation is more frequent after liver transplantation than before

liver transplantation [2]. The HCMV positivity in the pretransplant period, found in this study, also raise the hypothesis that viral activation might begin before liver transplantation. Several cases of ALF were associated with primary HHV-6 infection, mainly in infants and HHV-6B infection had yet identified in a liver sample [3, 4]. The indirect effects potentially attributed to HHV-7 have been less characterized, the reactivation of HHV-7 may increase the risk of HCMV infection [2, 5]. Betaherpeviruses can be associated with ALF. Therefore, after the exclusion of other etiologies. It should be included in the diagnostic differential to start therapy and even to avoid death in cases without defined etiology.

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## **Compliance with Ethical Standards**

**Ethics Committee Approval** Ethical permission for the collection and testing of samples was provided by the FIOCRUZ Ethical Committee (440.614).

Conflict of Interest None.

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