# Executive Summary: Hepatitis E virus infection in HIV-positive patients (MSS 281 R)

## **Aim and Objectives**

- 1. To elucidate the molecular prevalence and genotype(s) of hepatitis E virus (HEV) infection in HIV-positive patients in Hong Kong.
- 2. To determine the seroprevalence of HEV infection in HIV-positive patients in Hong Kong.

## **Project design**

Plasma samples of 1013 HIV-positive patients were collected from two local hospitals in Hong Kong. Viral RNA was extracted and the presence of HEV RNA and the corresponding viral load was measured using real-time quantitative RT-PCR. The HEV genotype was then determined by amplification of the RdRp gene using RT-PCR and PCR, followed by DNA sequencing and phylogenetic analysis. ELISA was used to determine the seroprevalence of HEV by targeting IgG antibodies in the patient plasma samples.

# **Target population**

People living with HIV/AIDS.

# Main achievements

HEV RNA was detected in 2.17% (22/1013) of HIV-positive samples. Phylogenetic analysis confirmed that the 22 HEV RNA-positive samples were of either HEV genotypes 3 (18 samples) or 4 (4 samples). The ratio of HEV genotype 3 and genotype 4 were 9:2 among the HEV-infected HIV-positive patients. In addition, the viral loads of these HEV RNA-positive samples was shown to range from  $1.94 \times 10^5$  to  $35.32 \times 10^5$  copies/ml.

HEV IgG was detected in 24.67% (250/1013) of HIV-positive samples which was slightly higher than that of the healthy population (20.7%).

Combining the RNA detection and ELISA results, 4 patients (0.39%) were observed to have both HEV RNA and IgG in their plasma samples.

## Conclusion

Our study showed that the seroprevalence of HEV IgG was slightly higher in HIV-positive patients than that of the healthy population in Hong Kong. This result is in line with previous studies which identifies higher seroprevalence of HEV IgG in HIV-positive patients compared to the healthy population. However, our study did not identify a sharp difference amongst the two groups. It is suggested that the humoral response of HIV-positive patients to HEV might not be as high as in the healthy population, hence the impeded immune response resulted in only a small difference in their seroprevalence of HEV IgG.

HEV RNA was detected in 2.17% of the HIV-positive patients with genotype 3 HEV being the predominant type circulating among this group of patients. The source and transmission route of this HEV genotype is not clear as HEV is known to be a foodborne transmitted virus in Hong Kong

and the transmitted HEV genotype is typically genotype 4 via consumption of uncooked pork. It will be interesting to further elucidate the source and transmission route of genotype 3 HEV.

This study has improved our understanding of the epidemiology of HEV infection in HIV-positive patients in Hong Kong and raises our awareness of HEV infection in HIV-positive patients and immunocompromised patients. Moreover, it suggests a change of our clinical practice on the management of HIV-positive patients. Untreated chronic HEV infection can lead to severe liver damage and cirrhosis, hence it is important that regular monitoring is performed for HIV-positive patients with HEV infection. By providing prompt treatment, progression of the disease can be prevented.

### 概要:愛滋病毒(HIV)患者中戊肝病毒(HEV)的感染概況(MSS 281 R)

### 目的

1. 闡明香港 HIV 患者中 HEV 的分子流行特徵及基因表型。

2. 判定香港 HIV 患者中 HEV 的血清流行特徵。

#### 課題設計

收集香港兩家醫院 1013 份 HIV 陽性血漿,提取病毒 RNA, RT-PCR 檢測 HEV RNA 的存在 情況及相應的病毒載量。RT-PCR 結合 PCR 擴增 HEV 陽性樣品中 HEV RdRp 基因片段, 並做測序和系統進化樹分析。同時 ELISA 檢測 HIV 陽性血漿中 HEV IgG 的流行情況。

### 目標人群

HIV/AIDS 患者

#### 主要成果

2.17% (22/1013) 檢測到 HEV RNA。系統進化樹分析證實 18 例為 HEV-3,4 例為 HEV-4, HEV-3 和 HEV-4 的比率為 9:2。病毒載量為每毫升 1.94 x 10<sup>5</sup> 到 35.32 x 10<sup>5</sup> 個。

24.67% (250/1013) 檢測到 HEV IgG, 稍高於健康人群(20.7%)。

結合 RNA 和 ELISA 檢測結果,4(0.39%) 個 HIV 患者中同時有 HEV RNA 和 IgG。

#### 結論

香港 HIV 患者中 HEV IgG 流行率稍高於健康人群, 這與之前發現的 HIV 患者中 HEV IgG 流行率高於健康人群一致, 但該研究未表明 HEV IgG 在這兩個群體中有很大的差別。這說 明在 HIV 感染者中針對 HEV 的體液免疫應答低於正常人群,這種阻礙造成了兩個群體間 HEV IgG 的流行僅有小差別。

2.17% 樣品中檢測到 HEV RNA, 主要為 HEV-3, 但 HEV-3 的來源及傳播途徑並不清楚。 HEV 為食源性傳播,香港流行的 HEV-4 被證實由食用未熟的豬肉獲得,因此進一步闡明 HEV-3 的來源和傳播途徑很有意義。

研究提升了對 HEV 在香港 HIV 及免疫缺陷病人中的流行病學的瞭解和認識,並建議在針對 HIV 患者的臨床干預上做一定改變。未經治療的慢性 HEV 感染可引發嚴重的肝損傷和 肝癌,因此定期的監測 HIV/HEV 共感染患者的肝臟健康狀態有利於提供及時治療預防疾 病進展。