



# The Brazilian Journal of INFECTIOUS DISEASES

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## Letter to the Editor

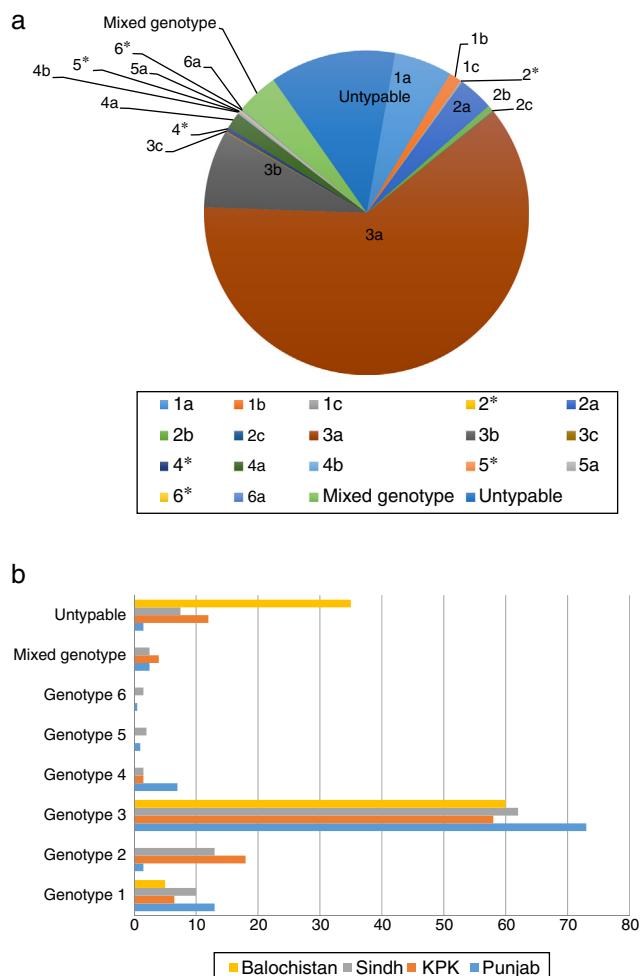
### Recent HCV genotype changing pattern in the Khyber Pakhtunkhwa province of Pakistan; is it pointing out a forthcoming problem?



Dear Editor,

A recent article by Gul et al.<sup>1</sup> highlights an important issue of Hepatitis C Virus (HCV) subtypes sero-prevalence in the Khyber Pakhtunkhwa (KPK) province of Pakistan. It is the first report of its kind on the changing epidemiological pattern of HCV genotypes in the province. The results of the study showed an increase in mixed HCV subtypes (22.9%) and genotype 1a (11.6%) and decrease in 3a genotype (45.5%).<sup>1</sup> The outcome of interferon plus ribavirin therapy is viral genotype specific, some genotypes/subtypes being good responders while others are not. Keeping in view the importance of sero-prevalent viral genotype regarding the management of patients, we analyzed the previous reports (26 published studies) from Pakistan on prevalence of HCV genotypes. The existing data showed that genotype 3a (57.37) is the most common genotype in the country while the prevalence of 1b subtype is only 1.985% and 1a is 5.93% (Fig. 1a), which support the study by Gul et al.<sup>1</sup> The analysis of previous published results regarding province wise distribution of the viral genotypes showed that 3a is the major circulating viral genotype in all four provinces of the country but the distribution of other viral clades is differential (Fig. 1b). By comparing the previously published results with the current report by Gul et al. it is evident that the pattern of HCV subtypes is changing in the region.

Due to the error prone nature of viral polymerase, HCV has many genotypes and subtypes. The interferon therapy response is genotype dependent and different viral genotypes showed varied level of sustained virological response (SVR). Among all viral genotypes 3a has shown to be a good responder to interferon therapy.<sup>2</sup> Pakistan is a resource constrained country with very low per capita income and considerably low expenditure on health by the Government (just 2.7% of GDP).<sup>3</sup> It ranked second in the world in terms of HCV infection with more than 10 million infections.<sup>3</sup> As the data indicates that 3a, supposedly to be a good responder to therapy, is the most prevalent genotype in the country most of the treated individuals with interferon therapy should attain SVR. Accordingly,



**Fig. 1 – Relative frequency distribution of HCV subtypes in Pakistan. \* Subtype not defined (a) and province-wise distribution pattern of major HCV genotypes (b).**

a most recent report (2014) from Pakistan regarding response of different HCV genotypes to interferon therapy showed 95% SVR in 3a infections in contrast to only 34% of 1b infections.<sup>4</sup> The current shift in the circulating viral genotype enlighten an important upcoming danger. Health care workers may face problems to successfully treat patients as it will be more difficult for patients to achieve SVR. For achieving higher rates of SVR antiviral therapy will have to be extended thus increasing the economic burden on society and psychology trauma on the patients and their families. SVR rates are greatly increased with direct acting antivirals (DAA) therapy but the high cost is a major obstacle.<sup>5</sup> In Pakistan, the allocated budget for health is already pretty low and the country is also endemic for other viral infections like polio and dengue which would result in shifting of priorities. By viewing the forthcoming giant of HCV therapy resistance, it is needed to properly monitor the pattern of HCV genotypes in the country. A well developed surveillance system is highly needed to timely tackle the upcoming problem.

## Role of the funding source

There is no role of any funding agency in this study.

## Conflicts of interest

The authors declare no conflicts of interest.

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Received 14 December 2015

Accepted 28 December 2015

Available online 8 March 2016

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<http://dx.doi.org/10.1016/j.bjid.2015.12.011>