

Tuesday, 11 February 2014

Dear Dr.

WJG 5550

Epidemiology of Malignant Digestive NeuroEndocrine Tumours (MD-NET)

Thank you for your reviewers' comments and your editorial notes. We are pleased to submit a revised manuscript (attached) with the following point-by-point responses.

As requested a "core tip" of less than 100 words as been added.

Referee: 1

- 1. Please write a paragraph in the discussion section regarding the potential etiology of CAV based on epidemiologic studies if possible.**

Etiological factors, due to the scarcity of these tumors, are not clearly indentified at the present time. We already had a paragraph in the discussion regarding this problematic.

« Few data are available about the role of environmental factors. Smoking is an identified risk factor^[18]. However the sex ratio of around 1.5 is not characteristic of cancers related to smoking. Obesity was also reported as a risk factor^[19]. It could explain at least part of the increasing trend reported in some areas. It was also suspected that cholecystectomy increased the risk of cancer of the ampulla of Vater^[20]. »

The rôle of endemic distomatose in South East Asia has been discussed. However it is not a risk factor in Europe.

2. Please discuss why the incidence of CAV showed a substantial increase in men over the study period.

This trend is difficult to explain. No data is available on risk factors according to gender.

Referee: 2

Incidence and management of cancer of the ampulla of Vater can be tabulated on the basis of years of study and the advancement of the understanding of the disease, by incorporating the reference numbers as a last column on the in all the Tables. This will not only make the data quickly readable and citing the back references will precisely add an invaluable effort to this article but will also ensure increased readability of this journal by other investigators. Although it appears that Authors have collected enormous evidences to support the article, it will be necessary to increase the number of situations, in order to make a real effort in the form of an original article.

On the whole, the manuscript is worth accepting and promises a large number of readability.

Minor Corrections: Please check the English and Grammar throughout the article before resubmitting.

We welcome this positive evaluation.

As suggested the table 1 as been modified. Unfortunately we cannot increase the number of situations, because of the small number of cases.

The paper has been reviewed by a native English speaker.