

(b) (6), (b) (4)

From: (b) (6) @assigndmb.com>
Sent: Wednesday, September 19, 2018 1:10 AM
To: (b) (6), (b) (4)
Cc: (b) (6) @celerion.com;
(b) (6) @celerion.com
Subject: VLN Cigarette Consumption Study/CA24914: First SAE of patient 001-023_Initial
Attachments: VLNC-Study-001-023-01_Initial_18SEP2018.pdf

Dear Sir and/or Madam,

please find enclosed the initial information concerning the first SAE "Subarachnoid hemorrhage" of subject 001-023 comprising the initial SAE report and the Medical Monitor's assessment (see below).

The onset date of event was on (b) (6), the day when the patient was admitted to hospital. The event was assessed as severe and no information was provided regarding the SAE outcome at the time of the initial report.

The information regarding the assessment of causality and action taken with study product was not provided.

The prepared case narrative and queries (#1-#3) have already been sent to the Medical Monitor. After the Medical Monitor reviewed and approved the narrative and queries, we will forward these documents to you.

There are still some open issues which had to be clarified (stop date of event, date of discharge, SAE outcome, assessment of causality, action taken with study product, medical history in order to determine different causes) with query handling.

Response of Medical Monitor:

The initial narrative contains the bare minimum to confirm that this is, in fact, an SAE. I confirm:

- at least one SAE criterion is met – hospitalization and life-threatening
- the relationship to study product is reasonable according to the definition in the protocol – unable to determine relationship. Temporal relationship is not evident from information. I would also want to see medical history to determine differential causes.
- the action taken towards SAE/study procedure by the PI are appropriate. – patient under the care of treating/admitting physicians. PI has promptly reported the event upon learning of it.

In case of any questions please do not hesitate to contact us.

Thank you very much in advance.

Kind regards,

(b) (6)

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