Reviewer’s report

Title: New case of trichorinophalangeal syndrome-like phenotype with a de novo t(2;8)(p16.1;q23.3) translocation which does not disrupt the TRPS1 gene

Version: 2 Date: 27 March 2014

Reviewer: Dezso David

Reviewer’s report:

The paper by Milena Crippa and co-authors describes clinical and molecular characterization of a second subject with translocation-associated trichorinophalangeal syndrome type I-like phenotype.

The pathogenic 8q23.3 breakpoint was located within LINC00536, at position g.116,981,667_116,981,668, approximately 213 kb downstream from a previously published translocation breakpoint at position g.116,768,309_116,768,310. This paper also points out the importance of the genomic region between the two translocation breakpoints and even further downstream for spatiotemporal regulation of TRPS1 expression.

Unfortunately, without feasible animal models mimicking these translocations we are unable to further dissect the “cis-ruption” effect of these translocations.

Discretionary Revisions

Abstract

Recommend revision of the last sentence of the Conclusions “The clinical and genetic characterisation of the present subject allowed us to make a genetic diagnosis in the context of a known syndrome, contributing to a better comprehension of the complex transcriptional regulation of TRPS1 and TRPS ethiopathogenesis”.

Background

Last paragraph, reconsider “partially overlap with TRPS I” to “mainly or largely” overlap with TRPS I; this might be more accurate.

Clinical report

MEN1 - Include gene name (multiple endocrine neoplasia I).

Additional files

Table S2 (List of additional primers used for amplification of chromosome der(2) and der(8) junction fragments) could be deleted.

I think that repetition of “lincRNA” in the context of “lincRNA LINC00536” is unnecessary.
Level of interest: An article of importance in its field

Quality of written English: Acceptable

Statistical review: No, the manuscript does not need to be seen by a statistician.

Declaration of competing interests:

I declare that I have no competing interests.