Reviewer's report

Title: Identification of a region required for TSC1 stability by functional analysis of TSC1 missense mutations found in individuals with tuberous sclerosis complex

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Reviewer: Teguh Haryo Sasonoko

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I change my opinion regarding the fact the E51D does not bring any pathogenic effect to TSC1 function. In fact, the authors have scientifically clarified that most of the missense mutations within N-terminal region they studied have hampered the function of TSC1.

Although their current and previous studies have shown some evidence closer to the whole of TSC1 N-terminal (AA 1-270) evidence is still lacking for the regions upstream of AA no.50. Therefore, the conclusion should be more precise mentioning at least the specific range of N-terminal region in which they found pathogenic missense mutation that hampers TSC1 function.

Level of interest: An article whose findings are important to those with closely related research interests

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