

Scientists slam Chinese CRISPR babies research after manuscript released

December 3 2019



Excerpts from the manuscript were published by the MIT Technology Review for the purpose of showing how Chinese biophysicist He Jiankui, pictured here, ignored ethical and scientific norms in creating the twins Lula and Nana

The gene editing performed on Chinese twins last year meant to



immunize them against HIV may have failed in its purpose and created unintended mutations, scientists said Tuesday after the original research was made public for the first time.

Excerpts from the manuscript were released by the MIT Technology Review for the purpose of showing how Chinese biophysicist He Jiankui ignored ethical and scientific norms in creating the twins Lula and Nana, whose birth in late 2018 sent shockwaves through the scientific world.

He made expansive claims of a medical breakthrough that could "control the HIV epidemic," but it was not clear whether it had even been successful in its intended purpose—immunizing the babies against the virus—because the team did not in fact reproduce the gene mutation that confers this resistance.

A small percentage of people are born with immunity because of a mutation in a gene called CCR5 and it was this gene that He had claimed he had targeted using a powerful editing tool known as CRISPR which has revolutionized the field since bursting on the scene in 2012.

Fyodor Urnov, a genome-editing scientist at the University of California, Berkeley told the MIT Technology Review: "The claim they have reproduced the prevalent CCR5 variant is a blatant misrepresentation of the actual data and can only be described by one term: a deliberate falsehood.

"The study shows that the research team instead failed to reproduce the prevalent CCR5 variant."

While the team targeted the right gene, they did not replicate the "Delta 32" variation required, instead creating novel edits whose effects aren't clear.



Moreover, CRISPR remains an imperfect tool because it can lead to unwanted or "off-target" edits, making its use in humans hugely controversial.

Here, the researchers claimed they had searched for such effects in the early-stage embryos and found just one—but this glossed over the fact that it would be impossible to carry out a comprehensive search without inspecting each of the embryo's cells and thus destroying it.

Further ethical concerns included that the parents of the babies might have wanted to take part for the wrong reasons.

The father was HIV positive, which carries a significant social stigma in China and makes it almost impossible to have access to fertility treatment, even though a well-established technique known as "sperm washing" prevents the infection from being passed on to unborn children.

Their lack of access to any kind of fertility treatment may have motivated them to take part in the experiment despite the huge risks to their children, Jeanne O'Brien, a reproductive endocrinologist at Shady Grove Fertility told the MIT Technology Review.

The authors also appeared to have taken steps to make it hard to find the family, like leaving the names of the fertility doctors off the paper, and including a false date of birth (Hu claimed November 2018 while multiple reports have indicated it was in fact October 2018).

He attempted to shop his manuscript around to prestigious journals including *Nature* and *JAMA*, but it remains unpublished.

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