



## NewScientist

### '3-parent baby' success

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Exclusive: World's first baby born with new "3 parent" technique

"The mother carries genes for Leigh syndrome, but the child shows no signs of the disease"

-Happy families-

'3-parent baby' success

A healthy boy is the first to be born using a new technique that uses DNA from three people, reports Jessica Hamzelou

IT'S a boy! A 5-month-old baby is the first to be born using a new technique that incorporates DNA from three people. "This is great news and a huge deal," says Dusko Ilic at King's College London, who wasn't involved in the work. "It's revolutionary."

The controversial technique, which allows parents with rare genetic mutations to have healthy babies, has only been <https://www.newscientist.com/article/dn26900-mitochondrial-replacement-vote-what-you-need-to-know/> legally approved in the UK. But the birth of the baby, whose Jordanian parents were treated by a US-based team in Mexico, should fast-forward progress around the world, say embryologists.

The mother carries genes for Leigh syndrome, a fatal disorder that affects the developing nervous system. Genes for the disease sit in DNA in the mitochondria, which provide energy for our cells and carry just 37 genes that are passed down to us from our mothers. This is separate from the majority of our DNA, which is housed in each cell's nucleus.

Around a quarter of the mother's mitochondria have the disease-causing mutation. While she is healthy, Leigh syndrome was responsible for the deaths of her first two children. She and her husband sought out the help of John Zhang (pictured below) and his team at the New Hope Fertility Center in New York City.

Zhang has been working on a way to avoid mitochondrial disease using a so-called "three-parent" technique. In theory, there are a few ways of doing this. The method approved in the UK is called pronuclear transfer and involves fertilising both the mother's egg and a donor egg with the father's sperm. Before the fertilised eggs start dividing into early-stage embryos, each nucleus is removed. The nucleus from the donor's fertilised egg is discarded and replaced by that from the mother's fertilised egg.

But this technique wasn't appropriate for the couple - as Muslims, they were opposed to the destruction of two embryos. So Zhang took a different approach, called spindle nuclear transfer. He removed the nucleus from one of the mother's eggs and inserted it into a donor egg that had had its own nucleus removed. The resulting egg - with nuclear DNA from the mother and mitochondrial DNA from a donor - was then fertilised with the father's sperm.

Zhang's team used this approach to create five embryos. They chose the male embryo that was of the highest quality to implant in the mother. Their baby was born nine months later. "It's exciting news," says Bert Smeets at Maastricht University in the Netherlands. The team will describe <http://www.sciencedirect.com/science/article/pii/S0015028216626705> the findings at the American Society for Reproductive Medicine's Scientific Congress in Salt Lake City in October.

Neither method has been approved in the US, so Zhang went to Mexico instead, where he says "there are no rules". He is adamant that he made the right choice. "To save lives is the ethical thing to do," he says.

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The team seems to have taken an ethical approach with their technique, says Sian Harding, who reviewed the ethics of the UK procedure. They avoided destroying embryos, and used a male embryo, so that the resulting child wouldn't pass on any inherited mitochondrial DNA, she . "It's as good as or better than what we'll do in the UK."

A remaining concern is safety. Last time embryologists tried to create a baby using DNA from three people was in the 1990s, when they injected mitochondrial DNA from a donor into another woman's egg, along with sperm from her partner. Some of the babies went on to develop genetic disorders, and the company performing the technique were advised to halt it. The problem may have arisen from the babies having mitochondria from two sources.

When Zhang and his colleagues tested the baby's mitochondria, they found that less than 1 per cent carry the mutation. Hopefully, this is too low to cause any problems; generally it is thought to take about 18 per cent of mitochondria to be affected before problems start. "It's very good," says Ilic.

Smeets agrees, but cautions that the team should monitor the baby to make sure the levels stay low. There's a chance that faulty mitochondria could be better at replicating, and gradually increase in number, he says. "We need to wait for more births, and to carefully judge them," says Smeets. n

### TWO WOMEN, ONE MAN and a baby

A Jordanian couple have been trying to start a family for almost 20 years. Ten years after they married, she became pregnant, but it ended in the first of four miscarriages.

In 2005, the couple gave birth to a baby girl. It was then that they discovered the probable cause of their fertility problems: a genetic mutation in the mother's mitochondria. Their daughter was born with Leigh syndrome, which affects the brain, muscles and nerves of developing infants. Sadly, she died aged 6. The couple's second child had the same disorder, and lived for 8 months.

Using a controversial "three-parent baby" technique (see main story), their third child was born on 6 April 2016. He is showing no signs of disease.

Jessica Hamzelou

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