

Scientists use stem cells to create models of pre-embryos

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This image, provided by the Institute of Molecular Biology of the Austrian Academy of Sciences, shows a human "blastoid" created in a lab. A new study in Nature on Thursday, Dec. 2, 2021, shows how scientists created this structure that mimics a blastocyst, a ball of cells that form within a week of fertilization, and can be used in research. Credit: Alok Javali, Heidar Heidari and Theresa Sommer/Institute of Molecular Biology of the Austrian Academy of Sciences via AP



Scientists are using human stem cells to create a structure that mimics a pre-embryo and can serve as a research alternative to a real one.

They say these "blastoids" provide an efficient, ethical way to study human development and pursue biomedical discoveries in fertility and contraception.

The latest effort was detailed Thursday in the journal *Nature*. The structures aren't embryos, but scientists nevertheless didn't let them grow past two weeks in deference to longstanding ethical guidelines.

A blastoid is a model for a blastocyst, a ball of cells that form within a week of fertilization and are about the width of a hair. Nicolas Rivron, a researcher at the Austrian Academy of Sciences and one of the authors of the Nature paper, said the models are "a fantastic alternative" to human.embryos for research, partly because donated embryos are hard to obtain and manipulate in the lab.

"It is extremely difficult to use such human embryos to discover any molecules, genes, principles that might allow us to better understand development and also make biomedical discoveries," Rivron said.

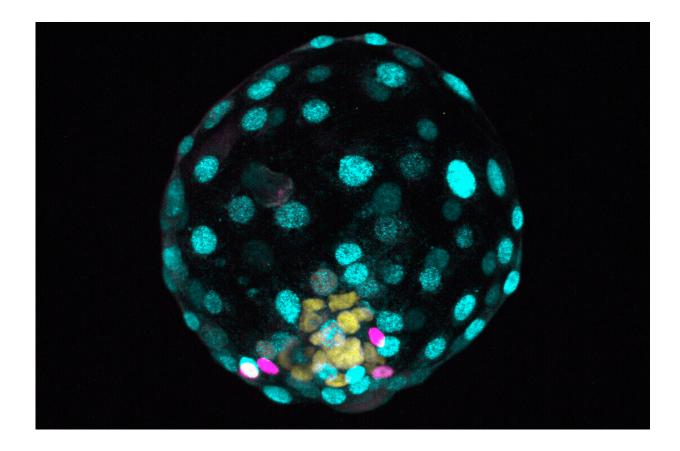
But lab-created stand-ins can be made, altered and studied in big numbers, and would complement embryonic research, he said.

"This unleashes the potential for scientific and biomedical discoveries," he said. For example, what researchers learn studying blastoids could be used to develop contraceptives that don't include hormones.

It's not the first time scientists have created a human blastoid, noted Magdalena Zernicka-Goetz, an expert in <u>stem cell biology</u> at the University of Cambridge who wasn't involved in the latest study. But "every single step is significant," improving efficiency as researchers try



to master the model, she said.



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To create the blastoids, Rivron and his colleagues used two different types of stem cells: either <u>embryonic stem cells</u> from previously established cell lines or stem cells reprogrammed from adult cells, such



as skin cells. No new embryonic cell lines were made for the research.

In the future, the stem cells reprogrammed from adult cells are likely to become the new standard in research, he said, but established embryonic cell lines are necessary now because they "are still the ultimate reference." He said blastocysts were cultured separately to compare them side-by-side with lab-created structures.

The study showed blastoids reliably replicated key phases of early embryo development. When they were placed in contact with cells from the lining of the uterus that had been stimulated with hormones, about half attached and started to grow in the same way blastocysts would.

Rivron said researchers stopped their growth after 13 days and analyzed the cells. At that point, he said, the collection of cells didn't reflect a 13-day-old embryo; they weren't growing enough or organizing as well.

He said ethical concerns also came into play: For decades, a "14-day rule" on growing embryos in the lab has guided researchers. Earlier this year, the International Society for Stem Cell Research recommended relaxing the rule under limited circumstances.

Rivron, part of the working group that updated the society's guidelines, said blastoids aren't subject to the same rule, but he pointed out the guidelines say they should never be transferred into an animal or human.

"It is very clear that blastoids are not <u>embryos</u> ... and if they are not, then why would we apply the 14-days rule to these structures?" he said. Nevertheless, they decided to stop "for the sake of transparency and for making sure things are very well understood by the public."

Dr. Barbara Golder, editor-in-chief of The Linacre Quarterly, the journal of the Catholic Medical Association, said the development of



blastoids shows "how science goes forward." But, she said, it's problematic that embryonic cell lines remain the standard in science.

"The ethical problems will exist as long as there's a connection to the stem <u>cells</u> that are derived from an aborted fetus and as long as we have to correlate one set of stem <u>cell lines</u> against ones that are embryonic stem-cell derived," she said.

More information: Nicolas Rivron, Human blastoids model blastocyst development and implantation, *Nature* (2021). DOI: 10.1038/s41586-021-04267-8. www.nature.com/articles/s41586-021-04267-8

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