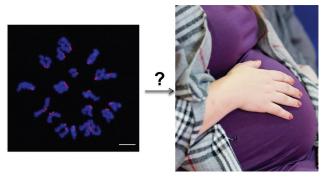


Why do so many pregnancies and in vitro fertilization attempts fail?

13 April 2020, by Todd Bates



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"Modeling efforts such as ours can provide guidelines on, for instance, how many eggs must be collected during a single IVF cycle to ensure there will be at least one chromosomally normal conception," said co-author Karen Schindler, an associate professor in the Department of Genetics and at the Human Genetics Institute of New Jersey.

Chromosomes in an egg. Modeling can help predict how many eggs should be retrieved for in vitro fertilization to lead to a normal, successful pregnancy. Credit: Cecilia Blengini and Kristen Driscoll

Pregnancy loss is extremely common, with nearly 20 percent of clinically recognized pregnancies resulting in miscarriage, and many more unrecognized pregnancies end earlier, the study notes.

Scientists have created a mathematical model that can help explain why so many pregnancies and in vitro fertilization attempts fail.

A leading cause of early miscarriage is called aneuploidy, when eggs have the wrong number of chromosomes, and it's also the main cause of IVF failure. The vast majority of eggs with chromosome problems are linked to errors in female cell division that increase as women age. Understanding how that happens is crucial because the average age at conception is rising in developed countries.

The Rutgers-led study, which may help to improve fertility, is published in the journal *Proceedings of the National Academy of Sciences*.

"Such basic knowledge is required to pave the way for future diagnostic and therapeutic innovations to improve human fertility," the study says.

Mistakes in female meiosis, the cell division process that creates <u>egg cells</u>, result in <u>eggs</u> with an abnormal number of chromosomes (too many or too few). This phenomenon is strongly associated with the repeated loss of pregnancies and the failure of in vitro fertilization (IVF) procedures, as well as <u>developmental disorders</u> such as Down syndrome.

The scientists developed a mathematical model describing all possible abnormal chromosome count issues in eggs due to cell division errors. Using data on 11,157 early stage human embryos (blastocysts), the model revealed previously unknown patterns of errors.

"Our study demonstrates that in the future, mathematical models can be powerful tools for predicting the outcomes of in vitro fertilization for infertility patients and/or provide the basis for considering alternative family planning options, such as adoption," said senior author Jinchuan Xing, an associate professor in the Department of

The model can be used to identify IVF patients who produce an extreme number of abnormal embryos. It's also a powerful tool for understanding why abnormal numbers of chromosomes arise when cells divide and for predicting the outcomes of IVF reproduction. The model potentially could provide



guidance for clinicians on the expected number of IVF cycles needed to get a normal conception for each patient. The modeling framework can also be expanded and adapted to address other processes, such as predicting errors in sperm.

More information: Katarzyna M. Tyc el al., "Mathematical modeling of human oocyte aneuploidy," *PNAS* (2020). www.pnas.org/cgi/doi/10.1073/pnas.1912853117

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