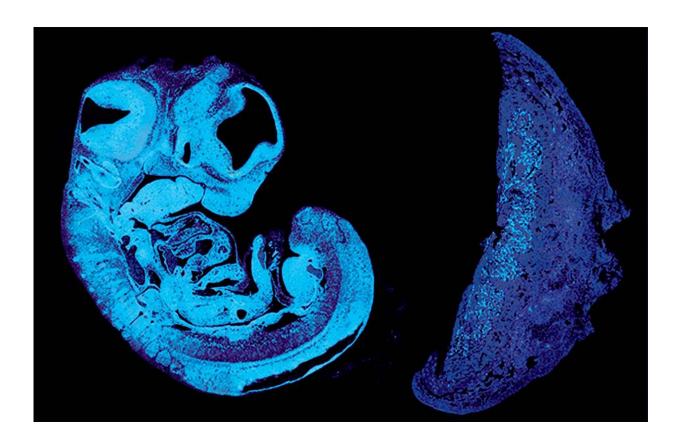


'Battle of the sexes' begins in womb as father and mother's genes tussle over nutrition

December 27 2021



Sections of the fetus and placenta. Credit: Ionel Sandovici

Cambridge scientists have identified a key signal that the fetus uses to control its supply of nutrients from the placenta, revealing a tug-of-war between genes inherited from the father and from the mother. The study, carried out in mice, could help explain why some babies grow poorly in



the womb.

As the fetus grows, it needs to communicate its increasing needs for food to the mother. It receives its nourishment via <u>blood vessels</u> in the placenta, a specialised organ that contains cells from both baby and mother.

Between 10% and 15% of babies grow poorly in the womb, often showing reduced growth of blood vessels in the placenta. In humans, these blood vessels expand dramatically between mid and late gestation, reaching a total length of approximately 320 kilometres at term.

In a study published today in *Developmental Cell*, a team led by scientists at the University of Cambridge used genetically engineered mice to show how the fetus produces a signal to encourage growth of blood vessels within the placenta. This signal also causes modifications to other cells of the placenta to allow for more nutrients from the mother to go through to the fetus.

Dr. Ionel Sandovici, the paper's first author, said: "As it grows in the womb, the fetus needs food from its mum, and healthy blood vessels in the placenta are essential to help it get the correct amount of nutrients it needs.

"We've identified one way that the fetus uses to communicate with the placenta to prompt the correct expansion of these blood vessels. When this communication breaks down, the blood vessels don't develop properly and the baby will struggle to get all the food it needs."

The team found that the fetus sends a signal known as IGF2 that reaches the placenta through the umbilical cord. In humans, levels of IGF2 in the umbilical cord progressively increase between 29 weeks of gestation and term: too much IGF2 is associated with too much growth, while not



enough IGF2 is associated with too little growth. Babies that are too large or too small are more likely to suffer or even die at birth, and have a higher risk to develop diabetes and heart problems as adults.

Dr. Sandovici added: "We've known for some time that IGF2 promotes the growth of the organs where it is produced. In this study, we've shown that IGF2 also acts like a classical hormone—it's produced by the fetus, goes into the fetal blood, through the <u>umbilical cord</u> and to the placenta, where it acts."

Particularly interesting is what their findings reveal about the tussle taking place in the womb.

In mice, the response to IGF2 in the blood vessels of the placenta is mediated by another protein, called IGF2R. The two genes that produce IGF2 and IGF2R are 'imprinted' – a process by which molecular switches on the genes identify their parental origin and can turn the genes on or off. In this case, only the copy of the *igf2* gene inherited from the father is active, while only the copy of *igf2r* inherited from the mother is active.

Lead author Dr. Miguel Constância, said: "One theory about imprinted genes is that paternally-expressed genes are greedy and selfish. They want to extract the most resources as possible from the mother. But maternally-expressed genes act as countermeasures to balance these demands."

"In our study, the father's gene drives the fetus's demands for larger <u>blood</u> vessels and more nutrients, while the mother's gene in the placenta tries to control how much nourishment she provides. There's a tug-of-war taking place, a battle of the sexes at the level of the genome."

The team say their findings will allow a better understanding of how the



fetus, <u>placenta</u> and mother communicate with each other during pregnancy. This in turn could lead to ways of measuring levels of IGF2 in the <u>fetus</u> and finding ways to use medication to normalise these levels or promote normal development of placental vasculature.

The researchers used mice, as it is possible to manipulate their <u>genes</u> to mimic different developmental conditions. This enables them to study in detail the different mechanisms taking place. The physiology and biology of mice have many similarities with those of humans, allowing researchers to model human pregnancy, in order to understand it better.

More information: Miguel Constância, The Imprinted Igf2-Igf2r Axis is Critical for Matching Placental Microvasculature Expansion to Fetal Growth, *Developmental Cell* (2021). <u>DOI:</u> <u>10.1016/j.devcel.2021.12.005</u>. <u>www.cell.com/developmental-cel ...</u> <u>1534-5807(21)00994-1</u>

Provided by University of Cambridge

Citation: 'Battle of the sexes' begins in womb as father and mother's genes tussle over nutrition (2021, December 27) retrieved 25 April 2024 from https://medicalxpress.com/news/2021-12-sexes-womb-father-mother-genes.html

This document is subject to copyright. Apart from any fair dealing for the purpose of private study or research, no part may be reproduced without the written permission. The content is provided for information purposes only.