TOOLBOX

Scientists aim to create scores of diverse mouse strains

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Founding fathers: Using just eight strains of mice, the Collaborative Cross initiative is planning to generate thousands of strains of genetically diverse mice.

Researchers are creating a population of inbred lab mice, called the **Collaborative Cross**, with the potential to produce thousands of genetically diverse strains. Experiments using the first of these mice are published in the August issue of *Genomic Research*^{1,2,3}.

By maximizing the genetic diversity of mouse strains, researchers can identify the gene variants responsible for various phenotypes and behaviors. Studying multiple variants can help researchers understand complex multi-gene disorders such as autism.

Researchers typically use inbred mouse strains, which have identical genes on each chromosome and whose genetics are well established. However, available strains are **strikingly genetically identical** to one another, greatly limiting the amount of genetic variation that researchers are able to study.

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To address this issue, in 2004, researchers launched a project called the Collaborative Cross to create 1,000 inbred mouse strains from 5 lab strains and 3 derived from wild populations⁴. The project involves interbreeding these strains to maximize their genetic variability. Further breeding among the Collaborative Cross strains has the potential to produce unlimited numbers of genetically diverse mice.

The project has already produced approximately 30 fully inbred strains⁵, with 40 to 50 projected for the end of the year and 100 by the end of 2012. Researchers are also conducting experiments on many of the existing 'nearly inbred' strains of mice.

For example, researchers tested 371 mice from 66 new strains for their susceptibility to the fungus *Aspergillus fumigatus*, which can also infect humans. Several mice show remarkable resistance to infection, allowing the researchers to hone in on the gene variants responsible for resistance³.

In another study, researchers documented the various phenotypes and behaviors visible in Collaborative Cross strains. These strains show a much wider variety of phenotypes, such as body length or level of anxiety, than those documented for BXD mice, an existing inbred lab strain².

The researchers also show that classic lab mouse strains have been selected for domesticity, by documenting wild-like behaviors in those bred from the wild-derived strains.

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