

Heat waves, food insecurity due to climate change may weaken immune systems

February 4 2019



Researchers at the University of Tokyo are studying the effects of climate change and nutrition on the immune system. Their recent research on influenza is published in the *Proceedings of the National Academy of Sciences*. Credit: University of Tokyo CC-BY

Heat waves can reduce the body's immune response to flu, according to new research in mice at the University of Tokyo. The results have



implications for how climate change may affect the future of vaccinations and nutrition.

Climate change is predicted to reduce crop yields and <u>nutritional value</u>, as well as widen the ranges of disease-spreading insects. However, the effects of <u>heat waves</u> on immunity to influenza had not been studied before.

University of Tokyo Associate Professor Takeshi Ichinohe and thirdyear doctoral student Miyu Moriyama investigated how high temperatures affect mice infected with influenza <u>virus</u>.

Flu in a heat wave

"Flu is a winter-season disease. I think this is why no one else has studied how high temperatures affect flu," said Ichinohe.

The influenza virus survives better in dry, cold air, so it usually infects more people in winter. However, Ichinohe is interested in how the body responds after infection.

The researchers housed healthy, young adult female mice at either refrigerator-cold <u>temperature</u> (4 degrees Celsius or 39.2 degrees Fahrenheit), <u>room temperature</u> (22 C or 71.6 F), or heat wave temperature (36 C or 96.8 F).

When infected with flu, the immune systems of mice in hot rooms did not respond effectively. Most affected by the high heat condition was a critical step between the <u>immune system</u> recognizing influenza virus and mounting a specific, adaptive response.

Otherwise, heat-exposed mice had no other significant changes to their immune system: They had normal reactions to flu vaccines injected



under the skin. Moreover, bacteria living in the gut, which are increasingly becoming regarded as important for health, remained normal in the mice living in hot rooms.

Temperature and nutrition

Notably, mice exposed to high temperature ate less and lost 10 percent of their <u>body weight</u> within 24 hours of moving to the hot rooms. Their weight stabilized by day two and then mice were infected by breathing in live flu virus on their eighth day of exposure to heat.

Mice living in heat wave temperatures could mount a normal immune response if researchers provided supplemental nutrition before and after infection. Researchers gave mice either glucose (sugar) or short-chain fatty acids, chemicals naturally produced by intestinal bacteria.

In experiments at room temperature, researchers surgically connected mice so that body fluids moved freely between underfed and normally fed mice, both infected with influenza. The fluids from normally fed mice prompted the immune systems of underfed mice to respond normally to the flu virus.

"Does the immune system not respond to <u>influenza virus</u> maybe because the <u>heat</u> changes gene expression? Or maybe because the mice don't have enough nutrients? We need to do more experiments to understand these details," said Moriyama.

The results may shed light on the unfortunate experience of getting sick again while recovering from another illness.

"People often lose their appetite when they feel sick. If someone stops eating long enough to develop a nutritional deficit, that may weaken the immune system and increase the likelihood of getting sick again," said



Ichinohe.

Future of infection

An important area of future study will be the effect of high temperature on different types of vaccinations. Flu vaccines injected into the upper arm use inactivated virus, but vaccines sprayed into the nose use live attenuated (weakened) virus.

"The route of delivery and the type of virus both may change how the immune system responds in high temperatures," said Moriyama.

Until more research can clarify what these findings may mean for humans, Ichinohe and Moriyama cautiously recommend a proactive approach to public health.

"Perhaps vaccines and nutritional supplements could be given simultaneously to communities in food-insecure areas. Clinical management of emerging infectious diseases, including influenza, Zika, and Ebola, may require nutritional supplements in addition to standard antiviral therapies," said Ichinohe.

The researchers are planning future projects to better understand the effects of temperature and nutrition on the immune system, including experiments with obese <u>mice</u>, chemical inhibitors of cell death, and different humidity levels.

More information: Miyu Moriyama el al., "High ambient temperature dampens adaptive immune responses to influenza A virus infection," *PNAS* (2019). www.pnas.org/cgi/doi/10.1073/pnas.1815029116



Provided by University of Tokyo

Citation: Heat waves, food insecurity due to climate change may weaken immune systems (2019, February 4) retrieved 20 April 2024 from https://medicalxpress.com/news/2019-02-food-insecurity-due-climate-weaken.html

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