

## Researchers hope a 'robo-nose' could give K-9 officers a break

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Golden retriever Rudy, wearing military dog tags and a U.S. flag, is being trained by the U.S. Army to use his amazing nose to find human remains. Credit: Army photo

Every day, thousands of trained K9 dogs sniff out narcotics, explosives and missing people across the United States. These dogs are invaluable



for security, but they're also very expensive and they can get tired.

Duke <u>researchers</u> have taken the first steps toward building an artificial "robot <u>nose</u>" device made from living <u>mouse cells</u> that officers could use instead of dogs.

The researchers have developed a prototype based on <u>odor receptors</u> grown from the genes of mice that respond to target odors, including the smells of cocaine and explosives. Their work appeared earlier this month in *Nature Communications*.

It turns out there are a couple of very big differences between testing things in a lab dish and testing them in an actual nose.

"This idea of an artificial nose has been present for a long time," said senior study author Hiroaki Matsunami, a professor of molecular genetics and microbiology in the Duke School of Medicine. "The receptors were identified in the 1990s, but there are significant technical hurdles to produce all these receptors and monitor the activity so that we can use that in an artificial device."

"E-noses" that exist now use various chemical compounds to detect smells instead of receptor stem cells, Matsunami said. He said those devices are "not as good as a trained dog."

"The idea is that by using the actual, living receptors, maybe we can develop a device similar to animals," Matsunami said. "Nobody has achieved that yet, but this study is moving toward that goal."

Human, dog and mouse genomes contain around 20,000 genes, which contain instructions to create proteins that smell, taste, feel, move and do everything that our bodies do. About 5 percent of mouse genes have been identified as instructions to make odor receptors, Matsunami said.



In contrast, humans only use about 2 percent of their genes to make odor receptors.

"These animals invest a lot of resources for this purpose," Matsunami said. "Mice and rats are very good smellers; we just don't use mice for detecting explosives in real life. There are some practical problems to do that."

The first step of the study was to identify the best odor receptors to respond to target odors like cocaine or marijuana.

The researchers created a liquid medium primed with molecules that could light up from reactions. Next they copied about 80 percent of the odor receptors from mice, and mixed those receptors with seven target odor chemicals in the medium.

They measured the resulting luminescence and chose the bestperforming odor receptors for the second part of the study, which monitored receptor activation in real time.

Previous research had done this by exposing selected receptors to odor chemicals in a liquid. But there are several differences between the petri dish and the nose. For one, we rarely submerge our noses into liquid baths of odor chemicals. Instead, our noses detect smells from wafting perfumes or stenches borne on the air. And our noses are full of mucus.

So, for the second half of the study, which was supported by the National Institute of Health grants DC014423 and DC016224 and the Defense Advanced Research Project Agency RealNose Project, they attempted to mimic how we use our noses by exposing odorants to vapors and a few enzymes.

The researchers tested the receptors they had identified against two odor



vapors for this study.

"We only tested two of them in the paper, but it's showing the proof of principle of how it can be used," Matsunami said.

The researchers hope they can fine-tune the device to test all receptors against many different smells.

"We have a panel of receptors so we can monitor how different receptors respond differently to various smells, including ones that are similar to each other in <u>chemical structure</u> or ones that might be related to real-world use, like something associated to explosives or drugs," Matsunami said.

The researchers also tested various enzymes that one might find in mucus to see how they aided or impeded reactions. This process is more true-to-life than vapor molecules directly interacting with <u>odor receptors</u>.

"You'd think when we smell a chemical, the chemical would bind to the chemical receptor in the nose, but actually it's not so simple," Matsunami said. "When the chemical dissolves in the nasal mucus before binding to the receptor, it might be converted to another <u>chemical</u> by enzymes in the nasal mucus."

Mucus is an unknown frontier in understanding how we smell. Reconstructing the key components of nasal mucus may be the next step toward building an artificial nose, according to the paper.

"It's not like our paper will be immediately applied to a portable device used in the airport soon, but this is an important step forward to show that it is possible," Matsunami said. "We can more clearly see what kind of hurdles to pass in order for the community to create such a device."



**More information:** Hitoshi Kida et al, Vapor detection and discrimination with a panel of odorant receptors, *Nature Communications* (2018). DOI: 10.1038/s41467-018-06806-w

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