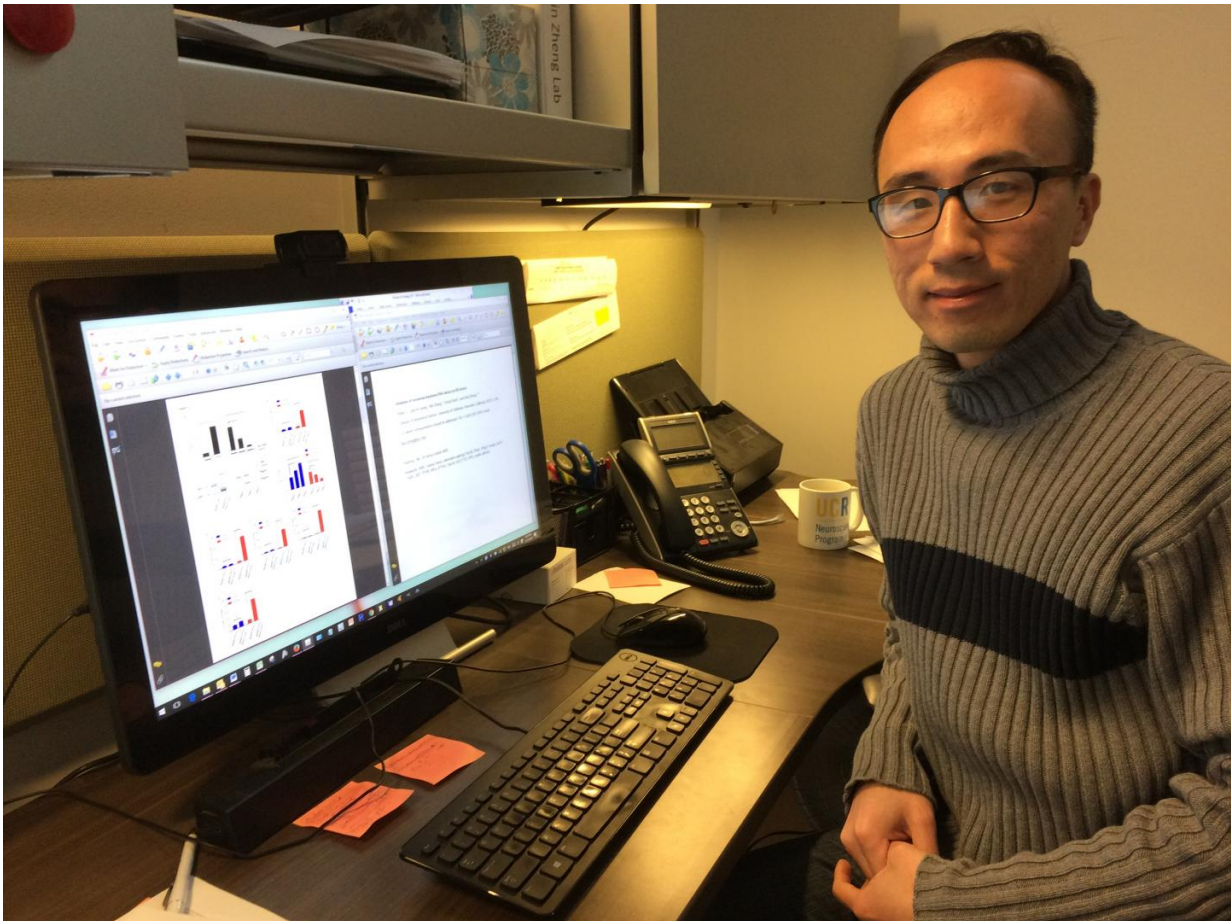


# Method developed by biomedical scientists could help in treatment of several diseases

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Sika Zheng is an assistant professor of biomedical sciences in the School of Medicine at UC Riverside. Credit: I. Pittalwala, UC Riverside.

In cells, DNA is first converted to RNA, and RNA is next converted to proteins—a complicated process involving several other steps. Nonsense-mediated RNA decay (NMD) is a processing pathway in cells that, like a broom, cleans up erroneous RNA to prevent its productive conversion into an aberrant protein, which could lead to disease.

In some diseases, like amyotrophic lateral sclerosis (ALS, also called Lou Gehrig's disease), excessive junk RNA is produced, possibly contributing to the disease. In such instances, more NMD is useful to get rid of the junk RNA.

In other diseases, such as muscular dystrophy and cystic fibrosis, a decrease in NMD is a better option. In these diseases, the NMD pathway eliminates the RNA, resulting in a complete loss of the gene function. But a defective RNA may translate to a semi-functional protein, which is better than no RNA for protein formation in cells. And so, a tuned-down NMD is more useful.

Sika Zheng, an assistant professor of biomedical sciences in the School of Medicine at the University of California, Riverside, and colleagues now [report in the journal \*RNA\*](#) (volume 23, no.3, March 2017 issue) that they have come up with a [method](#) in the lab that detects NMD efficiency inside the cell.

"Our method can screen a host of chemicals and allows us to identify molecules that regulate this efficiency," Zheng said. "We have already identified [thapsigargin](#) as a molecule that indirectly and robustly inhibits NMD."

The new method works by harnessing some normal targets of NMD and turning them into "reporters" of NMD activity. The method relies on the knowledge that NMD is more than a quality-control mechanism; it can also determine the level of some naturally occurring RNA. The lower the

abundance of these natural NMD in a cell, the stronger the NMD activity, and vice versa.

Zheng explained that in the past, measuring NMD efficiency was a cumbersome process and could be performed only on cell cultures, not directly on tissues.

"Our method is far easier and more sensitive, and for the first time can make measurements in all kinds of samples," Zheng said. "This is essential for drug screening. We are going to screen a larger library of chemicals to identify molecules that either boost or weaken NMD, which should help develop better and more targeted drugs for treating ALS, [muscular dystrophy](#) and cystic fibrosis."

Provided by University of California - Riverside

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