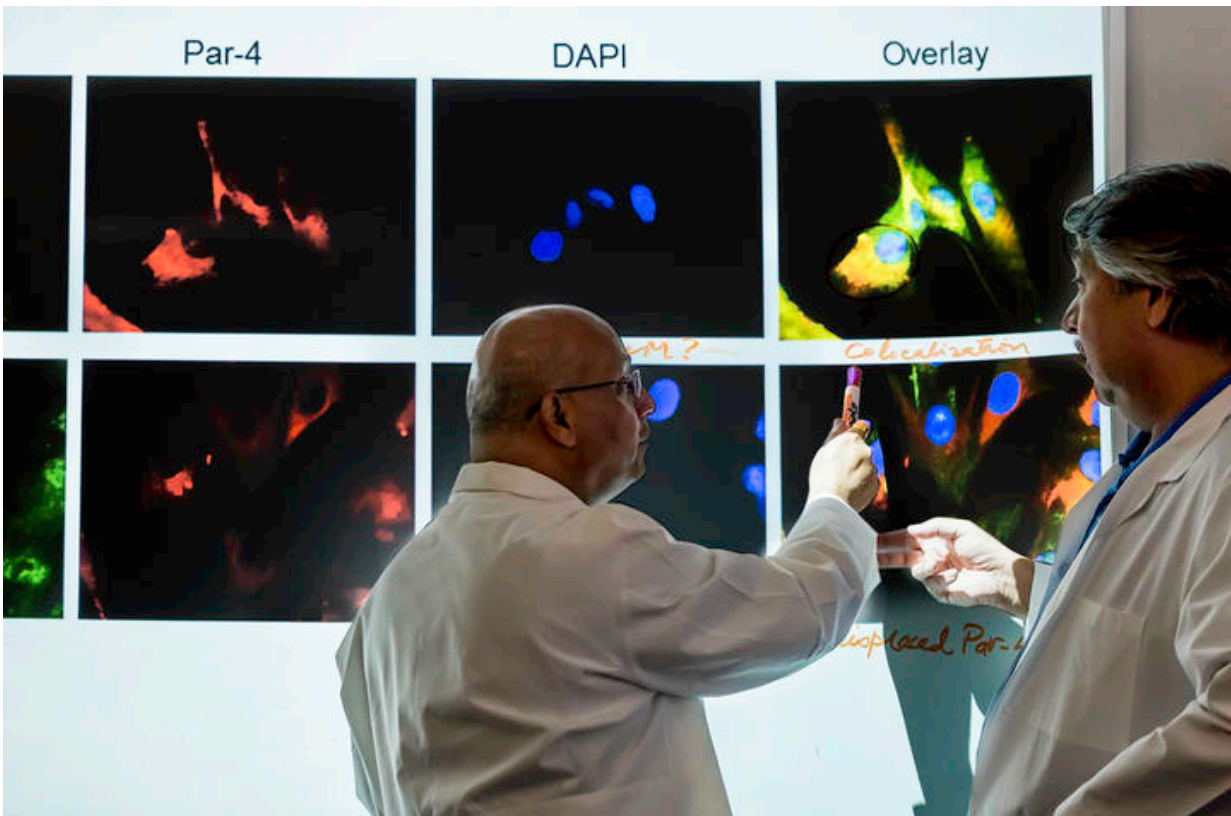


New study discovers "killer peptide" that eliminates resistant cancer cells

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UK's Vivek Rangnekar, left, and Ravshan Burikhanov. Credit: University of Kentucky

A new study by University of Kentucky Markey Cancer Center researchers shows that when therapy-sensitive cancer cells die, they release a "killer peptide" that can eliminate therapy-resistant cells.

Tumor relapse is a common problem following cancer treatment, because primary [tumor cells](#) often contain therapy-resistance cancer cells that continue to proliferate after the therapy-sensitive cells have been eliminated.

In the new study, published in *Cancer Research*, Markey scientists identified a Par-4 amino-terminal fragment (PAF) that is released by diverse therapy-sensitive cancer cells following therapy-induced cleavage of the tumor suppressor Par-4 protein. PAF caused death in cancer cells resistant to therapy and inhibited metastatic tumor growth in mice. Additionally, the PAF entered only cancer cells, not normal cells, keeping healthy tissue intact.

The findings indicate that this naturally generated PAF could potentially be harnessed to target neighboring or distant [cancer cells](#) to overcome metastasis and therapy resistance in tumors.

"This new information could positively impact how physicians plan their treatments, so as to use the sensitive cells in the tumor to release this peptide to help eliminate the [resistant cells](#)," said Vivek Rangnekar, principal investigator and Alfred Cohen Chair in Oncology Research with the UK College of Medicine's Department of Radiation Medicine. "We are developing PAF against therapy-resistant tumor metastasis for which no other treatment options are available."

Markey researchers Nikhil Hebbar and Ravshan Burikhanov from Rangnekar's team were the first two authors on the study, which also involved a partnership with Dr. Kojo Elenitoba-Johnson at the University of Pennsylvania.

Rangnekar's team first announced the generation of the cancer-resistant Par-4 mouse back in 2007. Since then, his team's work has spawned numerous research projects focusing on preventing and treating many

types of cancer, including Markey oncologist Dr. Peng Wang's clinical trial using the anti-malarial drug hydroxychloroquine to induce Par-4 secretion.

More information: Nikhil Hebbar et al. A Naturally Generated Decoy of the Prostate Apoptosis Response-4 Protein Overcomes Therapy Resistance in Tumors, *Cancer Research* (2017). [DOI: 10.1158/0008-5472.CAN-16-1970](https://doi.org/10.1158/0008-5472.CAN-16-1970)

Provided by University of Kentucky

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