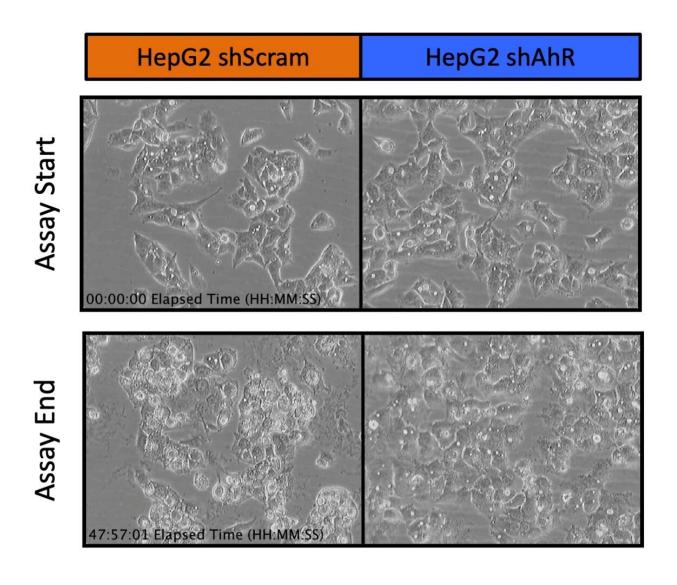


Researchers discover new class of cancerfighting compounds

May 11 2021, by Sean Nealon



A team of Oregon State University scientists has discovered a new class of anticancer compounds that effectively kill liver and breast cancer cells. The findings describe the discovery and characterization of compounds, designated as Select



Modulators of AhR-regulated Transcription (SMAhRTs). The researchers employed two molecular screening techniques to discover potential SMAhRTs and identified a molecule - known as CGS-15943 - that activates AhR signaling and kills liver and breast cancer cells. The images on the right show human hepatocellular carcinoma cells with the expression of the target AhR being killed upon treatment with the identified lead compound. The images on the left are human hepatocellular carcinoma cells without the target AhR. Credit: Oregon State University

A team of Oregon State University scientists has discovered a new class of anti-cancer compounds that effectively kill liver and breast cancer cells.

The findings, recently published in the journal *Apoptosis*, describe the discovery and characterization of compounds, designated as Select Modulators of AhR-regulated Transcription (SMAhRTs).

Edmond Francis O'Donnell III and a team of OSU researchers conducted the research in the laboratory of Siva Kolluri, a professor of cancer research at Oregon State. They also identified the aryl hydrocarbon receptor (AhR) as a new molecular target for development of cancer therapeutics.

"Our research identified a therapeutic lead that acts through a new molecular target for treatment of certain cancers," Kolluri said.

O'Donnell added, "This is an exciting development which lays a foundation for a new class of anti-cancer therapeutics acting through the AhR."

The researchers employed two molecular screening techniques to discover potential SMAhRTs and identified a molecule—known as



CGS-15943—that activates AhR signaling and kills liver and <u>breast</u> cancer cells.

Specifically, they studied <u>cells</u> from human hepatocellular carcinoma, a common type of liver cancer, and cells from triple negative <u>breast</u> cancer, which account for about 15% of breast cancers with the worst prognosis.

"We focused on these two types of cancers because they are difficult to treat and have limited treatment options," said Kolluri, a professor in the Department of Environmental and Molecular Toxicology in the College of Agricultural Sciences. "We were encouraged by the results because they are unrelated cancers and targeting the AhR was effective in inducing death of both of these distinct cancers."

The researchers also identified the AhR-mediated pathways that contribute to the anti-cancer actions of CGS-15943. Developing cancer treatments requires a detailed understanding of how they act to induce anti-cancer effects. The researchers determined that CGS-15943 increases the expression of a protein called Fas Ligand through the AhR and causes <u>cancer</u> cell death.

These results provide exciting new leads for <u>drug development</u>, but human therapies based on these results would not be available to patients for years, the researchers said.

Provided by Oregon State University

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