

## Researchers identify micro-RNA that targets oncogenic prostaglandin E2 production in lung cancer cell

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miR-708 and the arachidonic acid pathway. Illustration of miR-708's relationship to the AA signaling pathway. Blue lettering indicates short-lived intermediates and mature signaling molecules within the AA pathway. Red lettering illustrates a negative correlation between miR-708 and AA-related gene's mRNA expression in LUSC patients. Green lettering illustrates a positive correlation between miR-708 and AA-related gene's mRNA expression in LUSC patient. Black lettering illustrates no significant correlation. Solid black lines indicate metabolic steps, while dotted black lines indicate eicosanoid signaling to respective receptors. (\*\*) Represents predicted miR-708 targets from microRNA. org and miRTarBase. Credit: Correspondence to - Carol S. Lutz lutzcs@njms.rutgers.edu

In a study reported in *Oncotarget*, researchers have shown that high miR-708 expression is associated with survival rates in lung squamous cell carcinoma patients. miR-708 also represses PGE2 production by suppressing both COX-2 and mPGES-1 expression in lung cancer cells.

Moreover, miR-708 decreases proliferation,

survival and migration of <u>lung cancer</u> cells, which can be partially attributed to miR-708's inhibition of PGE2 signaling.

Lastly, they identified novel miR-708 predicted targets and possible regulators of miR-708 expression in lung cancer. Collectively, these data demonstrate that dysregulated miR-708 expression contributes to exacerbated PGE2 production, leading to an enhanced pro-tumorigenic phenotype in lung cancer cells.

Dr. Carol S. Lutz at Rutgers Biomedical & Health Sciences in Newark, New Jersey, said, "Lung cancer is the most common cancer, with more than 2.09 million lung cancer cases worldwide in 2018." More importantly, lung cancer is the deadliest cancer in the world, with more than 1.79 million lung cancer-related deaths in 2018.

Lung cancer is a collection of several distinct subtypes, with non-small-cell lung cancer accounting for 85% of all lung tumors. miR-708 also indirectly regulates the expression of genes involved in PI3K signaling, cell cycle progression, <u>epithelial-mesenchymal transition</u> and cancer cell stemness.

In this study, the authors aimed to decipher novel miR-708 targets and resolve the controversy regarding whether miR-708 is an oncogenic or tumor-suppressive mi RNA in lung cancer.

The Lutz research team concluded that collectively, their findings suggest further study of miR-708 in lung cancer. The data paired with previous studies highlights a potential value for miR-708 as a diagnostic in differentiating lung tumors, as well as a potential therapeutic intervention, particularly in lung squamous cell carcinomas.



Their work has identified novel tumor-suppressive miR-708 functions by suppressing oncogenic PGE2 production through targeting of COX-2 and mPGES-1. These findings could be the foundation for identifying novel miR-708 targets, as well as regulators of miR-708 expression in cancer. Moreover, the study highlights the need to better understand lung cancer biology to improve the diagnosis and treatment of lung cancer, ultimately aiming to increase positive patient outcomes.

**More information:** Nicholas J. Monteleone et al, miR-708-5p targets oncogenic prostaglandin E2 production to suppress a pro-tumorigenic phenotype in lung cancer cells, *Oncotarget* (2020). DOI: 10.18632/oncotarget.27614

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