

## P-49

**THE DETECTION OF HYPOXIA MARKERS IN NON-HPV AND HPV-ASSOCIATED HEAD AND NECK CANCERS: IMPLICATIONS TO PATIENT SURVIVAL**

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In the microenvironment of growing solid tumors, oxygen level often decreases and a hypoxic state is induced. This can lead to a worse treatment response and poor patient prognosis. One of the hypoxia-responsive genes is aspartate- $\beta$ -hydroxylase (ASPH), whose activity promotes the growth, invasiveness, and metastasis of many solid tumors<sup>1</sup>. Head and neck cancers (HNC) are highly heterogeneous. A proportion of HNC is induced by high-risk human papillomavirus (HPV) infections and is associated with better patient outcomes compared to patients with tumors linked to tobacco and alcohol abuse.

In our study, we analyzed 93 HNC specimens. ASPH and selected endogenous hypoxia markers (HIF1A, HIF2, VEGFA, GLUT1, P4HA1, CA9, MMP9, and MMP13) were detected by multiplex fluorescent immunohistochemistry. The results were correlated with tumor etiology, clinical and pathological characteristics of the patients. The Cox proportional hazards model was used to evaluate the prognostic value of the analyzed markers.

Statistically significant higher protein expressions of ASPH, HIF1A, GLUT1, and MMP13 were detected in the HPV-positive tumor group compared to the HPV-negative group. Except for MMP9/13, higher expression of the markers was detected in the tumor parenchyma compared to the stroma. Increased protein expression of GLUT1 and HIF1A had a positive impact on 5-year overall and disease-free survival in HNC patients, independently of HPV tumor status.

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## REFERENCE

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