



# Deciphering Glioma Subtypes: Implications for Prognosis and Treatment

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## INTRODUCTION

Gliomas represent the most common type of primary brain tumor, comprising a heterogeneous group of neoplasms that arise from glial cells within the central nervous system. Traditionally classified based on histological features, recent advances in molecular profiling have revealed distinct glioma subtypes characterized by unique genetic alterations and biological behaviors. Understanding the molecular landscape of gliomas and their prognostic significance is crucial for guiding treatment decisions and improving patient outcomes. One of the most widely recognized glioma subtypes is glioblastoma (GBM), a high-grade glioma associated with aggressive growth, treatment resistance, and poor prognosis. GBM is characterized by genetic alterations such as mutations in the IDH1/2 genes, loss of the chromosome 10q arm (containing the PTEN gene), and amplification of the epidermal growth factor receptor (EGFR) gene. These molecular alterations are associated with distinct biological pathways implicated in tumor growth, invasion, and resistance to therapy, providing valuable prognostic information and potential therapeutic targets.

## DESCRIPTION

In contrast to GBM, lower-grade gliomas (LGGs) are characterized by a slower growth rate, longer overall survival, and a propensity for progression to higher-grade tumors over time. LGGs encompass a spectrum of histological subtypes, including astrocytomas, oligodendrogliomas, and mixed gliomas, each with distinct molecular profiles and prognostic implications. The presence of mutations in the IDH1/2 genes and co-deletion of chromosome 1p/19q are defining molecular features of oligodendrogliomas, which are associated with improved response to chemotherapy and longer progression-free survival. Moreover, advances in molecular profiling have led to the identification of additional glioma subtypes with

unique genetic alterations and clinical behaviors. For example, gliomas with mutations in the H3F3A or HIST1H3B genes are classified as diffuse midline gliomas, often occurring in children and young adults and associated with a poor prognosis. These tumors frequently harbor additional genetic alterations such as TP53 mutations, amplification of the MYCN oncogene, or alterations in the histone methyltransferase genes EZH2 and ATRX, which contribute to their aggressive growth and treatment resistance.

Furthermore, molecular subtyping of gliomas has important implications for guiding treatment decisions and predicting response to therapy. For example, IDH-mutant gliomas are associated with a more favorable prognosis compared to IDH-wildtype tumors and may benefit from less aggressive treatment approaches such as observation or targeted therapies. Similarly, 1p/19q co-deleted oligodendrogliomas are highly sensitive to chemotherapy with alkylating agents such as temozolomide or PCV (procarbazine, lomustine, vincristine), leading to improved progression-free and overall survival compared to other glioma subtypes. Molecular subtyping of gliomas enables the development of targeted therapies that specifically target oncogenic pathways implicated in tumor growth and progression.

## CONCLUSION

Gliomas represent a heterogeneous group of tumors with distinct molecular subtypes characterized by unique genetic alterations and clinical behaviors. Molecular subtyping of gliomas has important prognostic implications, guiding treatment decisions and predicting response to therapy. By integrating molecular profiling into routine clinical practice, clinicians can optimize patient care, develop personalized treatment strategies, and improve outcomes for patients with gliomas. Through continued research and innovation, we strive to unravel the complexities of glioma biology and develop more effective therapies for these devastating diseases.

<b>Received:</b>	29-November-2023	<b>Manuscript No:</b>	IPJNO-24-19624
<b>Editor assigned:</b>	01-December-2023	<b>PreQC No:</b>	IPJNO-24-19624 (PQ)
<b>Reviewed:</b>	15-December-2023	<b>QC No:</b>	IPJNO-24-19624
<b>Revised:</b>	20-December-2023	<b>Manuscript No:</b>	IPJNO-24-19624 (R)
<b>Published:</b>	27-December-2023	<b>DOI:</b>	10.21767/2572-0376.8.4.39

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**Citation** Walker K (2023) Deciphering Glioma Subtypes: Implications for Prognosis and Treatment. Neurooncol. 8:039.

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