# Diffuse Glioma Testing Resources

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# Purpose

To provide a short list of resources for health care providers who take care of diffuse glioma patients.

# Links to important papers in DropBox

## What we know about glioblastoma

The molecular era of GBM begins

### The rise of IDH1/2

Original work comes from TCGA **GBM** dataset!?

Lower grade diffuse gliomas examined by massive sequencing effort

IDH mutant diffuse and anaplastic astrocytomas: pretty much the same thing

The end of IDH wild type diffuse astrocytoma: WHO update with c-IMPACT-NOW update 3

# Why methylation of MGMT is important

The original New England Journal report Wick's review piece

# In our backyard: Mayo's "Revolution in Brain Tumor Diagnosis"

Mayo stratifies gliomas with 1p/19q, IDH status and TERT promoter methylation

### IDH-WT grade II or III infiltrating astrocytoma: GBM in sheep's clothing

Adult IDH wild type grade II and grade III gliomas should be further stratified Adult IDH wild type grade II and III gliomas belong to GBM molecular class Inventing a new class of diffuse glioma to accommodate the molecular findings Prognostic relevance of genetic alterations to diffuse lower grade gliomas

So you think it might be a "Diffuse astrocytic glioma, IDH-wildtype, with molecular features of glioblastoma, WHO grade IV"... how do you support your impression?

EGFR amplification, +7 and -10 shown to be helpful. . . at least in classification

TERT promoter mutation, +7, -10 shown to improve prognostication in IDH WT diffuse lower grade gliomas

But prognosis and prediction are mostly a matter of IDH mutation and MGMT promoter methylation

#### Additional testing may be useful in some IDH mutant tumors too

CDKN2A loss is associated with shortened survival in IDH mutant grade II and III astrocytomas

#### Other considerations

Diffuse midline gliomas and histone mutations: nomenclature considerations

H3K27M mutants: a plurality in pediatric high grade glioma and DIPG

### In infants and children, see especially...

#### **BRAF** Alterations

Pilocytic astrocytoma and BRAF translocation/duplication

BRAF V600E in context with BRAF translocation/duplications: V600E heralds other tumors too

#### Other uncommon low grade neuroepithelial tumors

Low-grade neuroepithelial tumors: BRAF, FGFR1 and MYB mutations

#### High grade supratentorial pediatric neuroepithelial tumors

Uncommon supratentorial high grade neuroepithelial tumors

### High grade supratentorial pediatric neuroepithelial tumors

Developing ontology: Figure 1 and Table 1 in this paper show how to start dissecting these rare and lethal pediatric tumors

Pediatric gliomas: a review

#### Methylation profiling based classification: what's the fuss?

Time to put away the microscope?

#### Panels for diffuse gliomas come of age.

A diffuse glioma NGS panel covering a variety of targets. (Duesseldorf)

# Important websites:

# Sites for exploring The Cancer Genome Atlas (TCGA) data:

Memorial Sloan Kettering Cancer Center:cBioPortal Broad Institute of MIT and Harvard: Firebrowse

Fred Hutch, Seattle, Oncoscape

# Site for connecting to methylation profiling in Europe:

Heidelberg methylation profiling