

# Oligodendrogliomas

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Most "benign" of gliomas! - never grade 4

## EPIDEMIOLOGY

- 4-19% of all intracranial tumors.
- 2-25% of all gliomas (only 6% in children).
- most commonly - young and middle-aged adults (median age 25-50 yrs).

## CLASSIFICATION

1. **Oligodendroglioma** (WHO grade 2) ≈ 80%; median survival 6-10 yrs.
2. **Anaplastic (malignant) oligodendroglioma** (WHO grade 3) ≈ 15-20%; median survival 2.2-4 yrs.

N.B. there is no grade 4 oligodendroglioma.

## GENETICS, MOLECULAR MARKERS

Definitive diagnosis (a must mutations!) - **IDH1/2 mutation + 1p19q co-deletion** (assay by FISH)  
 N.B. most of low grade oligodendrogliomas are positive for IDH1 R132H mutation with intact ATRX nuclear staining\*.  
 – if histology looks like oligo, but IDH-wild type – call astrocytoma!  
 N.B. deletions of both 1p36 and 19q13 = greater response to chemotherapy.  
 N.B. it has to be deletion of both (co-deletion!)

**100% response to chemotherapy with 1p 19q LOH.**

\*ATRX remains present (vs. astrocytoma).

## LOCATION

- single lesion in **cerebral hemispheres** (white matter):  
**FRONTAL** > PARIETAL, TEMPORAL > OCCIPITAL lobe (3:2:2:1 ratio).
- rarely, in cerebellum, brain stem, spinal cord.
- 10% tumors disseminate through CSF.

## PATHOLOGY

- Low-grade oligodendroglioma** (grade 2)
- grossly **well demarcated** (but generally infiltrative); 20% are cystic.
  - **very cellular** - monotonous side-by-side collection of homogeneous, compact, rounded cells with distinct borders and clear cytoplasm surrounding dark uniform central nucleus ("fried egg appearance"). No conspicuous fibrillary background!
  - may **infiltrate diffusely into cortex** around normal neuronal elements (without causing loss of function) → may extend to leptomeninges.
  - neoplastic cells may tightly surround neurons (**perineuronal satellitosis**).
  - within tumor, branching blood vessels (**delicate network of anastomosing capillaries**) are highly characteristic - divide cells into discrete clusters - "chicken-wire" capillary pattern.
  - **microcalcification** may be extensive.
  - many oligodendrogliomas have some component of astrocytoma within them;
    - it is difficult to distinguish neoplastic astrocytes from reactive astrocytes.
    - some tumors are truly mixed **OLIGOASTROCYTOMAS** (both cell types arise from common precursor - *oligodendrocyte type-2 astrocyte, s. O2A cell*); minimum proportion of astrocyte is 10-25%.



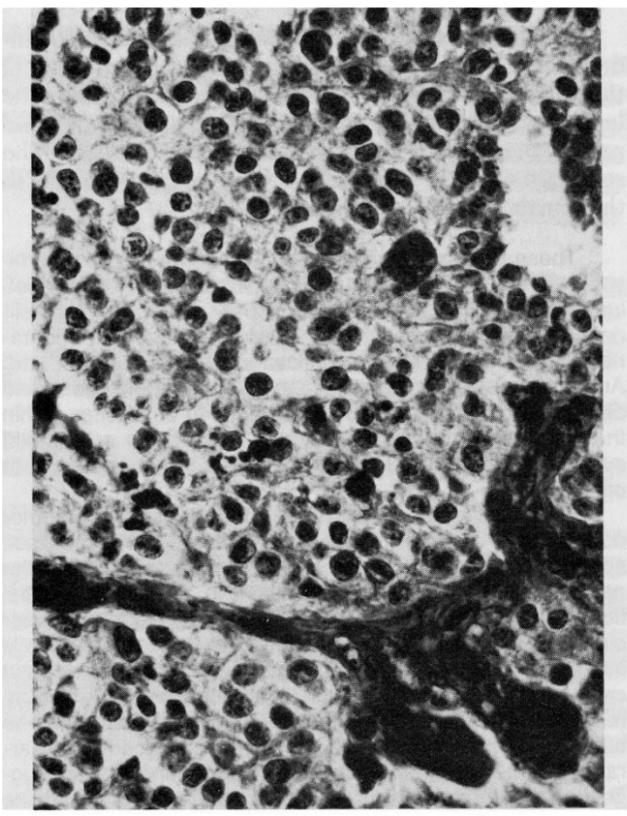
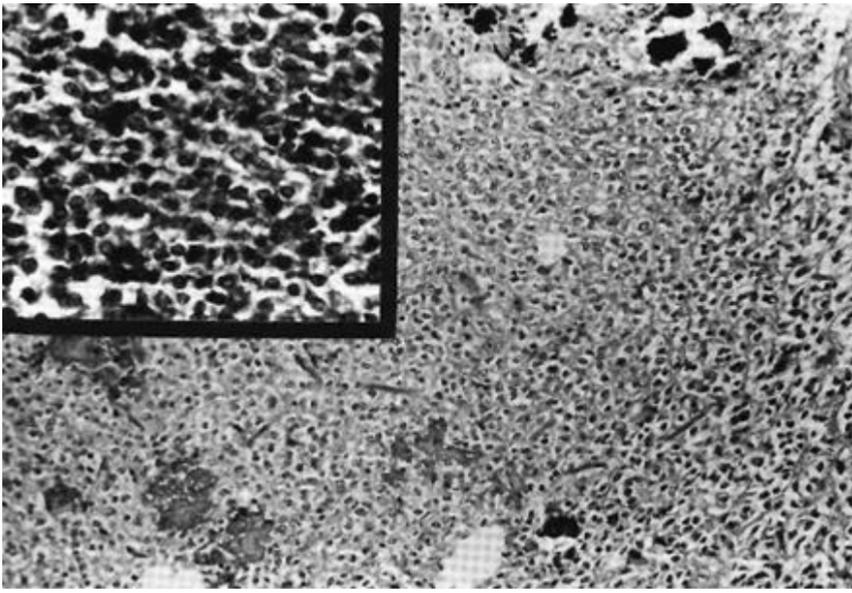
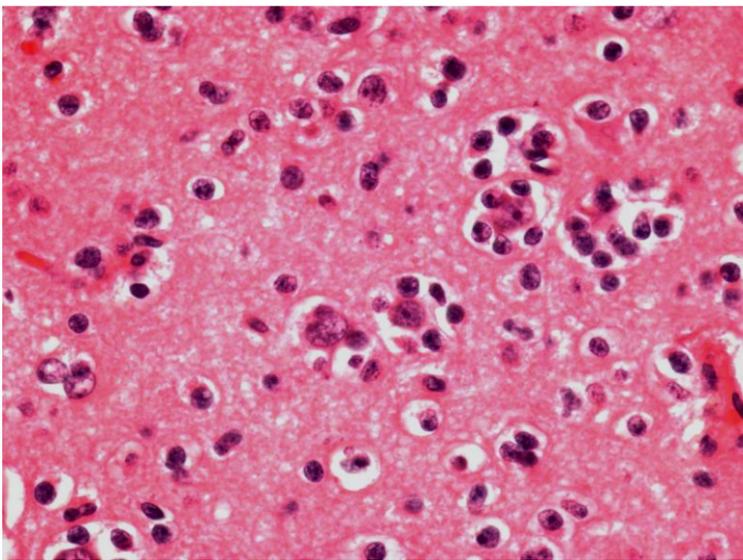
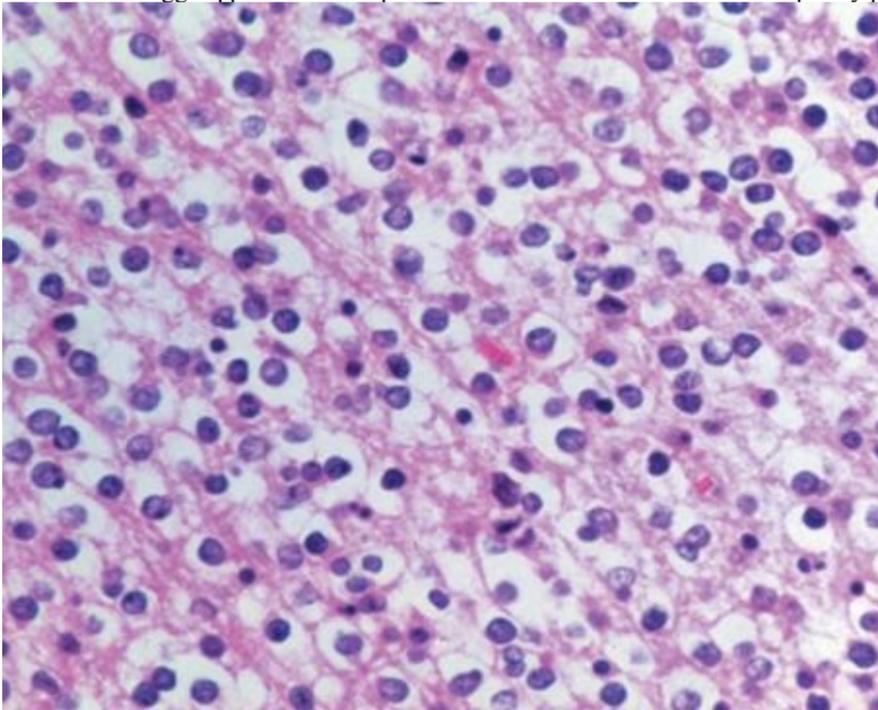
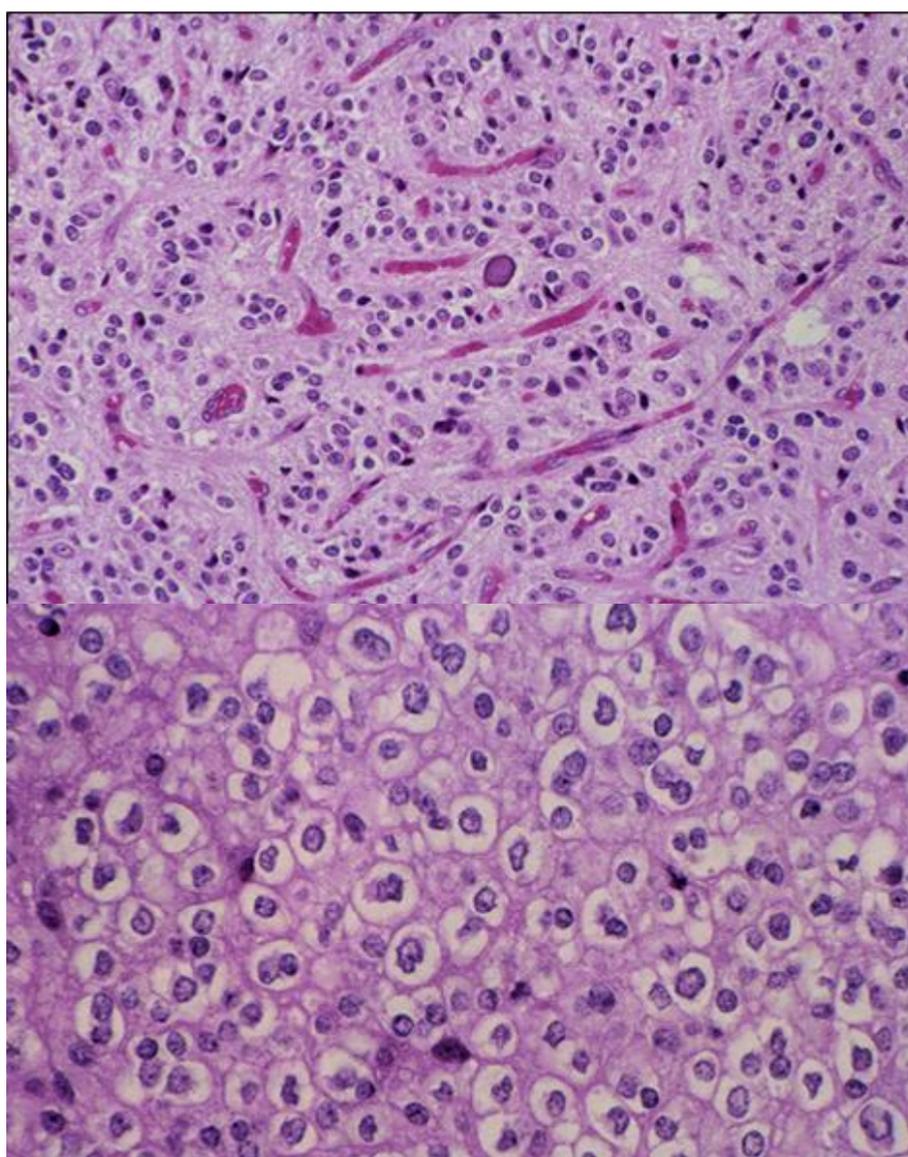


Figure 29–28. Oligodendroglioma. Cells are round and small and have perinuclear halos. (H and E stain.)



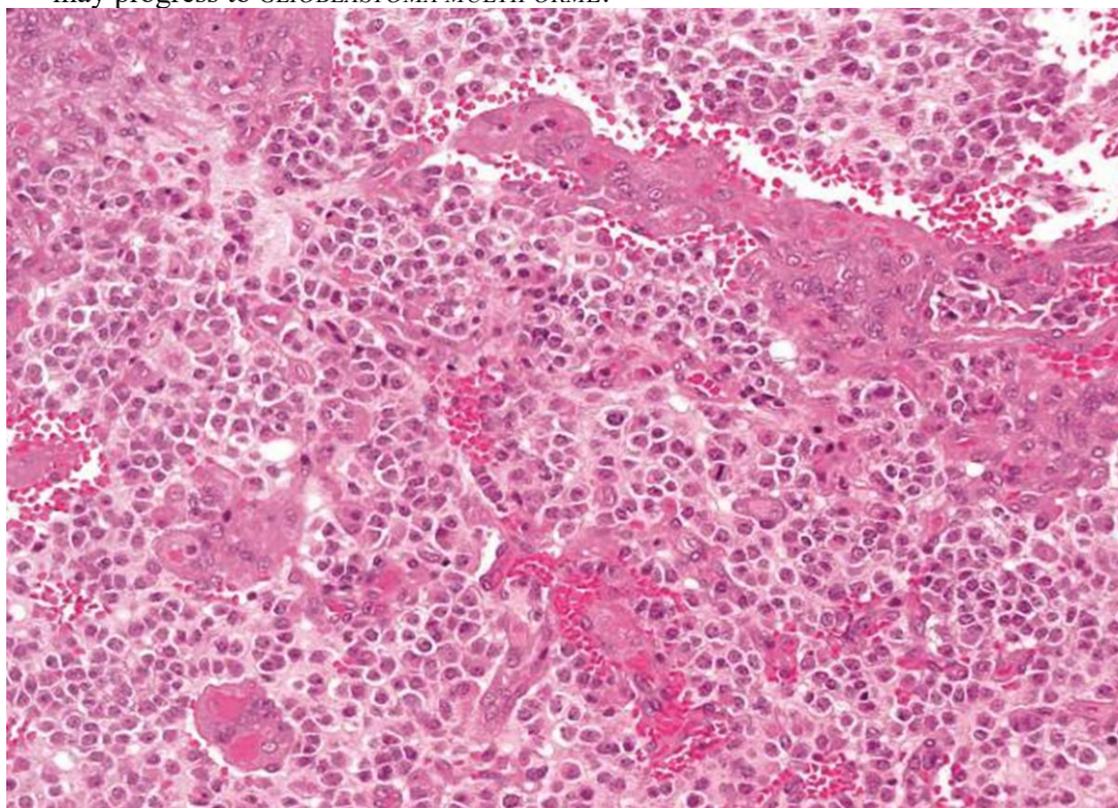
Classic "fried egg" appearance with perinuclear halos and "chicken-wire" capillary pattern:





**Anaplastic (malignant) oligodendroglioma** (grade 3) - increased cellularity, nuclear pleomorphism, endothelial proliferation, mitotic activity, and necrosis.

- may progress to *GLIOBLASTOMA MULTIFORME*.



### CLINICAL FEATURES

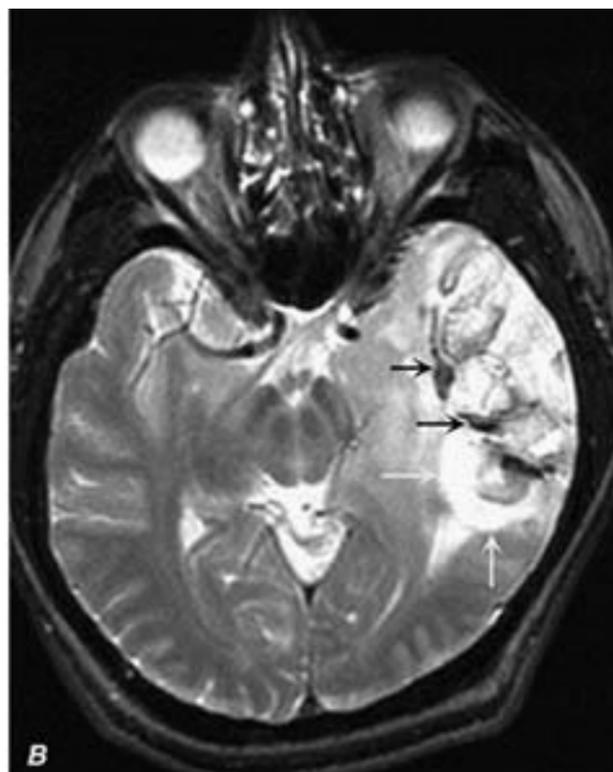
- duration of symptoms before diagnosis averages 7-11 years!
- most common (50%) presenting symptom is **seizure!**; 80% patients have seizures at some time.  
Seizures are more common with oligodendrogliomas than other gliomas!
- **focal cerebral dysfunction**, rarely ICP↑.

### DIAGNOSIS

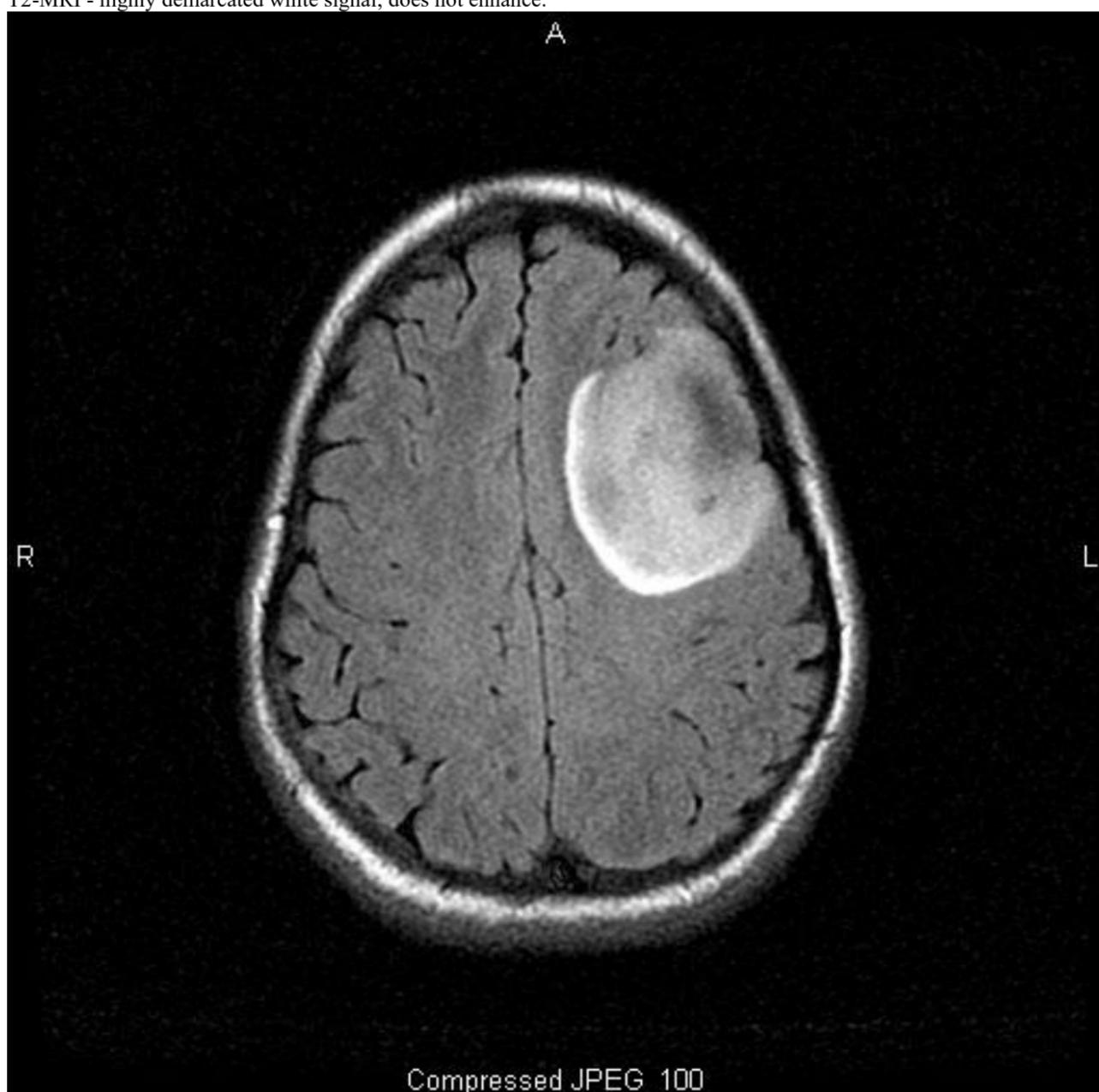
- **CT** - invisible (unless calcified\*). \*calcification fleck on CT may be first clue to neoplasm
- **MRI**: **LOW-GRADE TUMORS** - generally do not enhance (FLAIR is positive), while **ANAPLASTIC OLIGODENDROGLIOMA** does enhance; intratumoral **calcification** is common (≈ 90%).
- definite diagnosis – **biopsy** (almost always possible).

Differentiate **INTRAVENTRICULAR OLIGODENDROGLIOMA** from **CENTRAL NEUROCYTOMA** and **DYSEMBRYOPLASTIC NEUROEPITHELIAL TUMOR** – do not need chemotherapy and radiotherapy!

A. Noncontrast CT - calcified mass in left temporal lobe (*arrows*); mild mass effect but little edema.  
 B. MR-T2 - heterogeneous mass with hypointense signal (*black arrows*) surrounded by higher signal zone (*white arrows*), consistent with calcified temporal lobe mass.

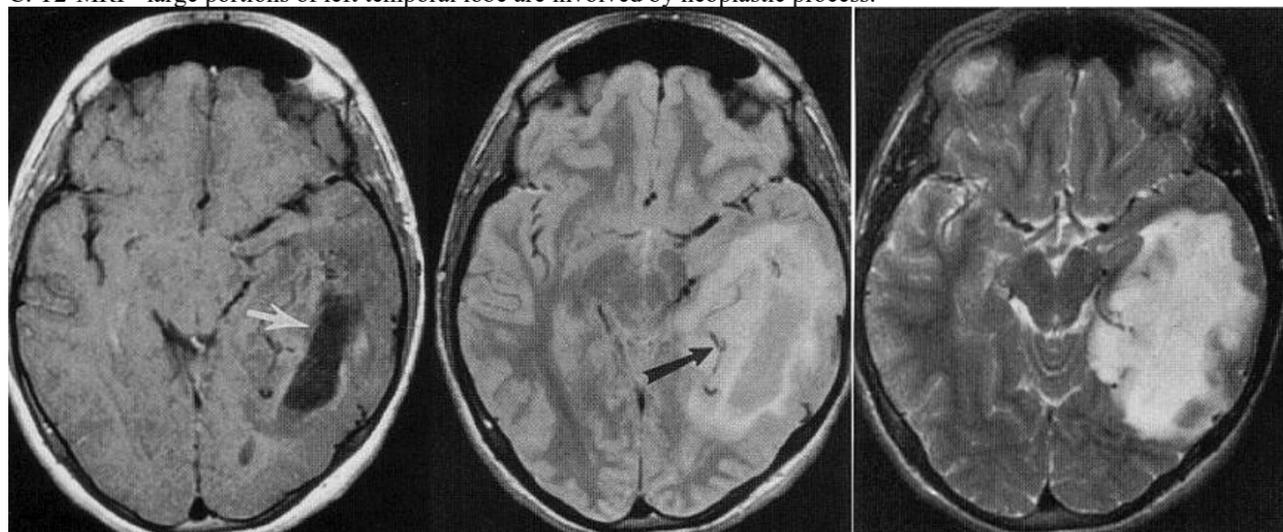


T2-MRI - highly demarcated white signal; does not enhance:



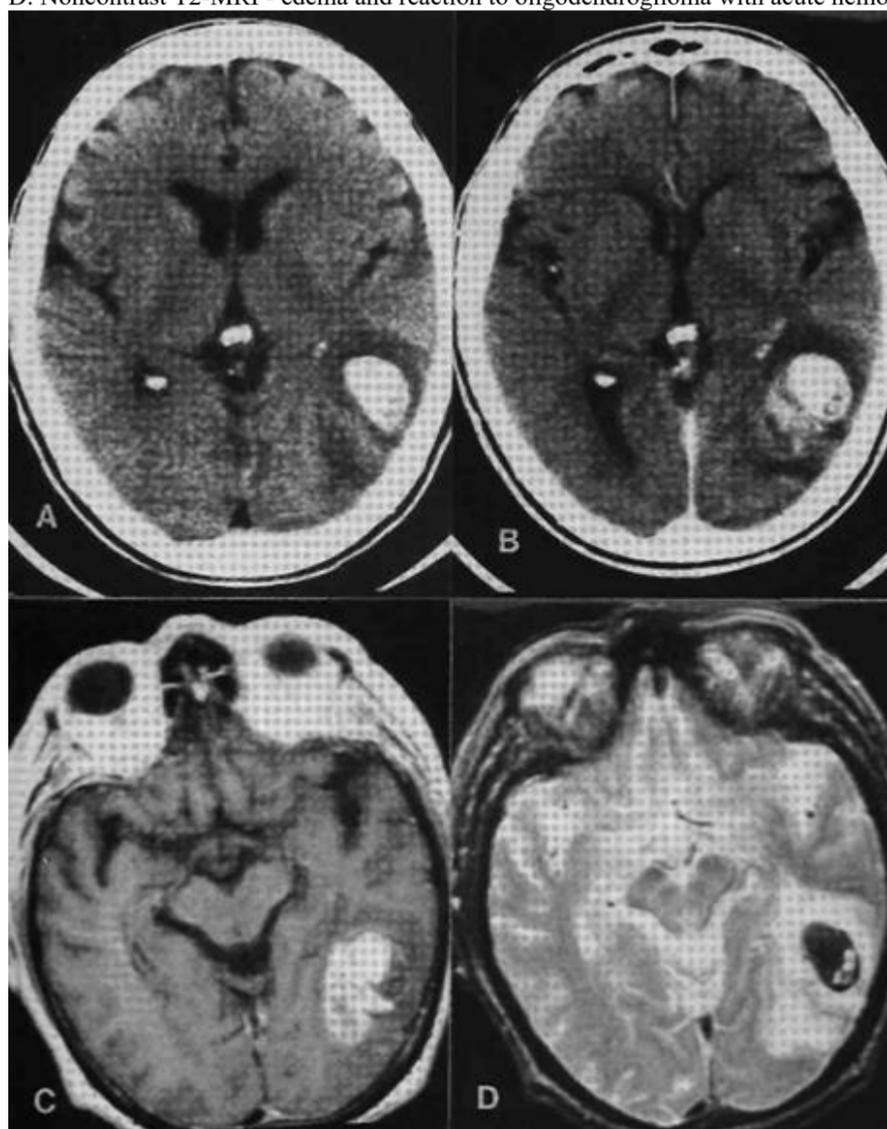
Anaplastic oligodendroglioma:

- A. T1-MRI - minimal heterogeneous contrast enhancement; central area of low signal intensity indicates necrosis (*arrow*).
- B. Spin density - better delineates extent of vasogenic edema and vascular structures within and adjacent to neoplasm (*arrows*).
- C. T2-MRI - large portions of left temporal lobe are involved by neoplastic process.



Spontaneous hemorrhage into mixed oligodendroglioma:

- A. Noncontrast CT - spheroid hematoma in mass with calcification located in left parietal lobe surrounded by zone of decreased attenuation.
- B. Contrast CT - enhancing tumor and relationship of hematoma.
- C. Noncontrast T1-MRI - hemorrhage in tumor and surrounding edema.
- D. Noncontrast T2-MRI - edema and reaction to oligodendroglioma with acute hemorrhage.



## TREATMENT

No intervention ÷ aggressive multimodal treatment

Grade II guidelines - see p. Onc10 >>  
Algorithm - see p. Onc10 >>

- **anticonvulsive therapy** is recommended once oligodendroglioma is diagnosed.
- some small asymptomatic (except for controlled seizures) tumors can be **observed**.
- **surgery** - mainstay of treatment (resection is usually subtotal because of infiltrative nature of tumor - *surgical cure remains unlikely!*);  
total gross resection (esp. in < 40 yo) → **observation** for recurrence; recurrence → **radiotherapy**.  
incomplete removal → **radiotherapy**.

*ANAPLASTIC OLIGODENDROGLIOMA* (regardless of resection extent) → **radiotherapy**.

- use 2-3 cm margin for 54-60 Gy radiotherapy (children – 50 Gy).
- **chemotherapy** - favorable response (most chemosensitive of gliomas)!!! (esp. in **combined loss of 1p/19q**):
  - a) for recurrences
  - b) adjuvant for *ANAPLASTIC OLIGODENDROGLIOMA*
 standard - PCV = **PROCARBAZINE** + **LOMUSTINE (CCNU)** + **VINCRIStINE**  
 for relapses also may be tried - **TEMOZOLOMIDE**

**100% response to chemotherapy with 1p 19q LOH.**

Ino Y, et al. *Clin Cancer Res.* 2001;7:839-845

**VORASIDENIB** – FDA approved, improves progression-free survival vs placebo (27.7 months vs. 11.1 months). see p. Onc3 >>

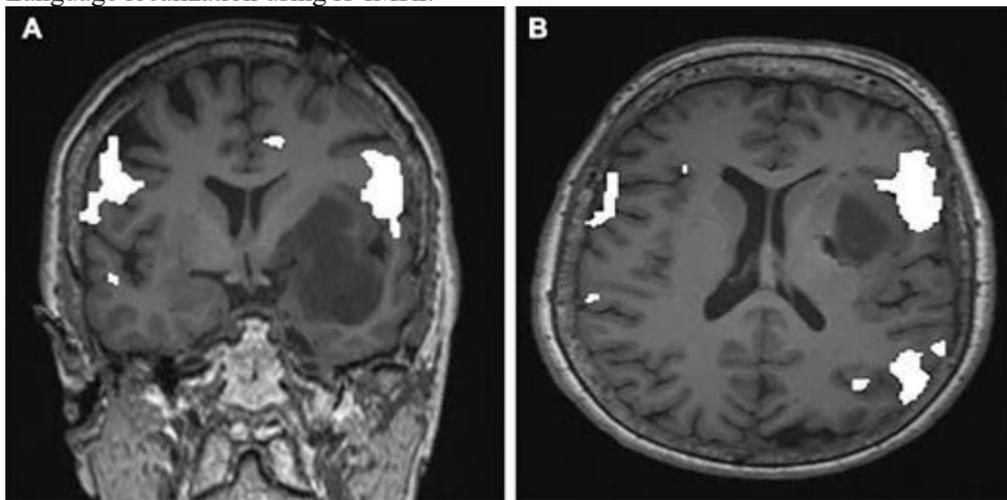
**LITT**

**Insular oligo**

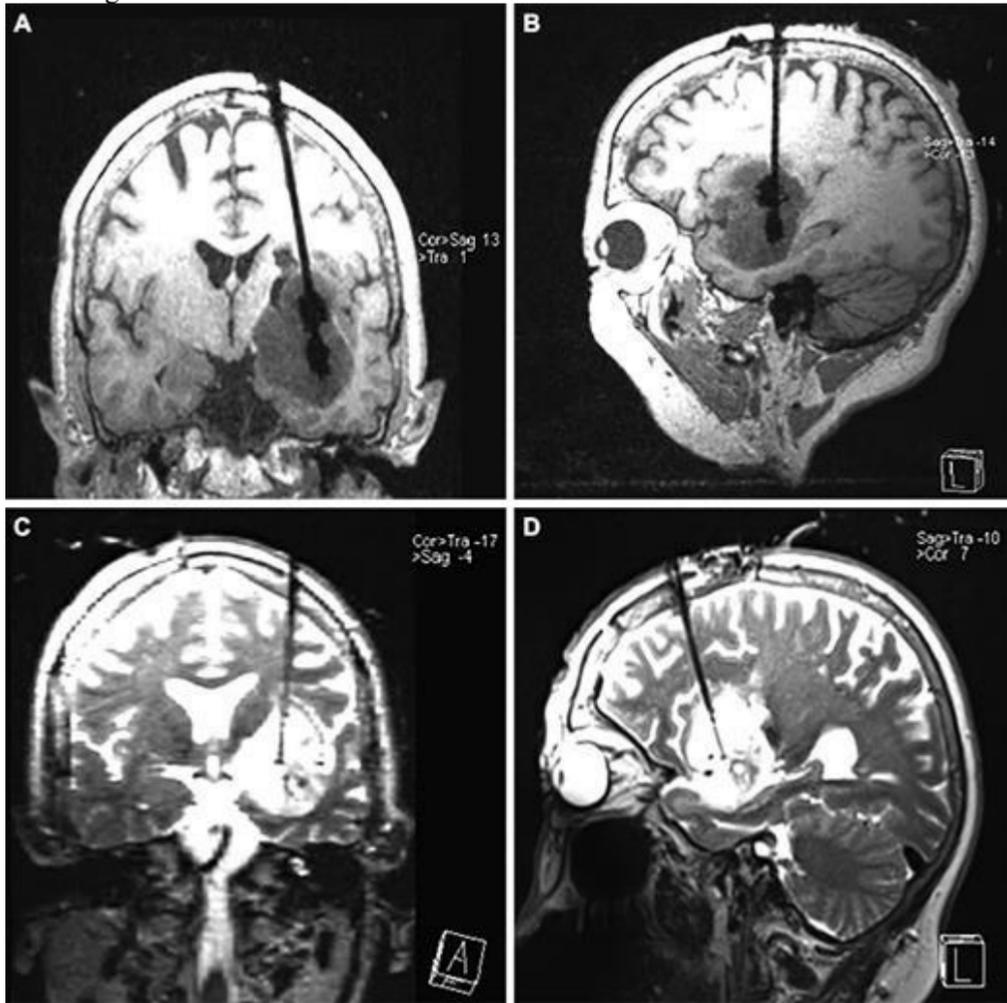
*Daniel M Hafez et al. Staged Laser Interstitial Thermal Therapy (LITT) Treatments to Left Insular Low-Grade Glioma. Neurosurgery, Volume 86, Issue 3, March 2020, Pages E337–E342*

- left-sided insular oligodendroglioma treated in two stages (3 months apart - due to large 5 cm size – to prevent severe edema and seizures) with no permanent clinical deficit (temporary mild difficulty with word repetition) → chemoradiation → near resolution of the tumor at 2 yrs:

Language localization using rs-fMRI:



A-B. Stage 1  
C-D. Stage 2



**PROGNOSIS**

- prognosis is much better than for *ASTROCYTOMAS*!  
 N.B. late progression of disease is common (5-year survival time used to indicate "cure" in other cancers is not relevant for oligodendrogliomas)
- indolent course - patients may survive for many years.
- **combined loss (co-deletion) of 1p/19q** is significant predictor of longer survival in anaplastic oligodendroglioma;  
 vs. **CDKN2A/B deletion** – worse overall survival in in anaplastic oligodendroglioma
- prognosis is worse for **mixed tumors (OLIGOASTROCYTOMAS)**.

BIBLIOGRAPHY for ch. “Neuro-Oncology” → follow this [LINK](#) >>