

GUIDELINES ON THE MANAGEMENT OF SEIZURES IN PATIENTS WITH BRAIN TUMOURS (PRIMARY OR SECONDARY). Dr. Yvonne Hart, Dr. Allyson Parry

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The diagnosis and management of patients with seizures is complex and usually requires specialist neurological expertise. Where possible, oncologists are encouraged to refer or discuss the patient with a neurologist.

Who to contact for advice

JRH	Dr Yvonne Hart– ext. 31891, Dr Allyson Parry – ext. 31892
	Neurology SpR on call – JRH switchboard
Other hospitals	The local consultant neurologist

How do you establish the diagnosis of a seizure?

This is dependent on an experienced review of the history obtained from the patient and any witnesses.

What are the possible causes of a seizure in this patient group?

- i) Presence of a space occupying lesion (seizure often focal onset)
- ii) Metabolic derangement e.g., Low Na, Ca
- iii) Accelerated metabolism of anti-epileptic drug by another medication
- iv) Effect of RT
- v) Chemotherapy related (usually with encephalopathy)
- vi) Vascular – arterial or venous cerebral thrombosis
- vii) Non-compliance with pre-existing anti-epileptic medication
- viii) Immunosuppressed patient with an opportunistic cerebral infection

Should the patient be started on anti-epileptic medication?

In general, most (but not all) patients start treatment. However, this decision, along with the particular choice of medication, may be complex. *It is recommended that this is done in conjunction with a neurologist.*

What anti-epileptic drug (AED) should be started?

Drugs favoured by the authors:

Focal onset seizure – carbamazepine, lamotrigine,
Primary generalised seizure – sodium valproate, lamotrigine

N.B. (i) Sodium valproate and phenytoin can be given *intravenously*, (if the patient cannot swallow safely or is not able to absorb oral medication adequately), at the same dose as the patient's usual oral dose (*Important - see BNF regarding the rate of intravenous administration*).

(ii) Phenytoin can be useful when rapid seizure control is required, or if the AED must be given intravenously. However, the use of phenytoin as a long-term AED is not recommended by the authors. Ideally, the patient should be discharged with a plan to switch to an alternative medication, under appropriate supervision.

Summary of commonly used anti-epileptic drugs

Each of these four drugs can be used for both partial (focal onset), and generalised (tonic-clonic) seizures. All drugs are potentially teratogenic. **This table must be used in conjunction with the BNF, particularly regarding the rate of dose escalation.**

Drug	Start dose/day	Common maintenance dose/day	Dosage interval	Interactions*	Common Side-effects
Carbamazepine <u>Slow Release</u>	200mg	400-1200mg	bd	Enzyme Inducer	Dizziness, diplopia, nausea, rash
Phenytoin	200mg	250-450mg	od	Enzyme Inducer	Rash, gum hypertrophy, hirsutism
Valproate	600mg	600-2000mg	bd	Enzyme Inhibitor	Weight gain, tremor, hair loss, <i>platelet dysfunction</i> ++
Lamotrigine	25mg (except in patients on valproate, when use 25mg alt day)	100-400mg	bd	Probable enzyme inducer	If rash - discontinue , fatigue, headache, dizziness, diplopia

*Enzyme inducers may reduce the plasma concentration of other hepatically metabolised drugs e.g., oral contraceptive pill, corticosteroids. Enzyme inhibitors may increase the concentration of other hepatically metabolised drugs, e.g., valproate increases the plasma concentration of temozolomide and lamotrigine.

++ *Sodium valproate can cause platelet dysfunction and/or thrombocytopenia.* Platelet function tests should therefore be performed prior to any surgery.

Is fairly rapid seizure control required (NOT status epilepticus)?

Patients can be loaded with phenytoin (15mg/kg/24 hours) either orally or iv, before starting a maintenance dose. Valproate therapy can also be initiated quickly (BNF)

Special note on the non-linear pharmacokinetics of phenytoin

Small increases or decreases in the dose of phenytoin can result in large increases or decreases in the plasma concentration. This can result in either neurotoxicity (nausea, ataxia, dysarthria, mental slowing and nystagmus), or a decline in seizure control, respectively. After a daily dose of 250mg, do not alter the daily dose of phenytoin by more than 50mg/increment. It is recommended that a phenytoin level is then checked a week later before further dose alterations are made.

Monitoring of drug levels

With the exception of phenytoin (see above), monitoring of plasma drug levels is not recommended, unless clinically indicated.

The patient who has been started on AED (usually pre-operatively), with NO history of seizures. When do you stop the AED?

We recommend that AED are stopped 2 weeks post-operatively. However, it is important that this decision is discussed with the relevant neurosurgeon.