

Guidance for the prescribing and monitoring of phenytoin (IV and oral) in Adults

- For **STATUS EPILEPTICUS** management see guideline:
<https://nhs.uk/guidelines/scot.nhs.uk/media/2340/management-of-generalised-convulsive-status-epilepticus-in-adults.pdf>

- For **UNCONTROLLED EPILEPSY**:

❖ If **NEW phenytoin**:

1. Load with phenytoin as per Status Epilepticus guideline¹ (see above)
2. If to continue therapy, commence **maintenance dose** of phenytoin (**3-5mg/kg/day²**) starting **12-24** hours after loading dose (if given)
3. Ensure appropriate monitoring (tables 1 and 2)

❖ **ON phenytoin** prior to admission:

1. Check phenytoin and albumin levels
2. If **albumin** level is **LOW** correct the phenytoin level using the equation¹ below:

$$\text{Corrected phenytoin concentration} = \frac{\text{measured phenytoin concentration}}{(0.9 \times \text{Albumin (g/L)} / 42^*) + 0.1}$$

*Mid-point of albumin range

(Phenytoin is highly protein-bound but only the unbound concentration is active. If serum albumin concentration is low, a higher proportion of the total (measured) phenytoin concentration is unbound and caution is therefore required when interpreting the result- the equation gives a rough estimate and the patient's clinical condition should be the most important consideration. Seek advice from neurology or pharmacy if needed)

3. If **phenytoin** level (or corrected phenytoin level) is **LOW**:
 - Give a top-up dose of phenytoin to achieve a therapeutic level (table 3)
 - Ensure appropriate monitoring (tables 1 and 2)
 - Consider increasing the maintenance dose of phenytoin- usually small increments only (25-50mg) with monitoring (tables 1 and 2), discuss with neurology if required
4. If **phenytoin** level (or corrected phenytoin level) is **NORMAL**:
 - Consider increasing the maintenance dose of phenytoin- usually in small increments only (25-50mg) with monitoring (tables 1 and 2) **OR**
 - Discuss with neurology re addition of other anti-epileptic agents

Table 1- When to check phenytoin levels²

Target concentration- 5-20mg/l- remember to always check albumin level and correct if low
After loading or top-up doses- 2-4 hours post IV dose or 12-24 hours after oral dose
Commencing maintenance dose or maintenance dose change- 3 -5 days after starting maintenance dose then re-analyse after a further 5-10 days (further accumulation can occur)
Routine monitoring- trough concentration (i.e. sample prior to next dose)
Suspected phenytoin toxicity- check level daily until < 20mg/l and/ or clinical signs resolved (table 4)

Guidance for the prescribing and monitoring of phenytoin (IV and oral) in Adults

Table 2- Phenytoin safety and monitoring requirements

- Check drug is prescribed correctly on HEPMA
- Check drug dose is appropriate
- Check drug is being administered and route is appropriate (table 5)
- Check LFT's and renal function
- Always check albumin level when checking phenytoin levels
- If new therapy- monitor for serious side effects e.g. leucopenia, skin reactions (BNF³)
- Consider drug interactions (BNF³)
- Maintain patient on the same **BRAND** of phenytoin

Table 3- TOP-UP dose equation²

Phenytoin sodium Top-up dose (mg) = (20 – measured concentration (mg/l)) x 0.7 x weight (kg)

Dose \ Weight	Approximate concentration increase			
	50kg	60kg	70kg	80kg
250mg	7mg/l	6mg/l	5mg/l	4.5mg/l
500mg	14mg/l	12mg/l	10mg/l	9mg/l
750mg	21mg/l	18mg/l	15mg/l	13.5mg/l

(Approximate increase in phenytoin concentration with 'top-up' doses)

Example- if the patient weighs 70 kg and has a measured concentration of 5 mg/L, a single dose of 750 mg will increase the concentration to around 20 mg/L (5 mg/L + 15 mg/L).

Table 4- Common signs of Phenytoin Toxicity^{3,4}

Drowsiness	Confusion	Slurred speech
Ataxia	Nausea	Nystagmus
Mental changes	Hyperglycaemia	Coma

Table 5- Conversion between formulations^{2, 5}

Phenytoin sodium **100 mg capsules / tablets/ injection** = phenytoin **90mg suspension**

(Patients with **enteral feeding tubes**- administer phenytoin by parenteral injection if possible as enteral absorption is extremely unpredictable- contact pharmacy for further advice if required)

References

1. Management of generalised convulsive Status Epilepticus in Adults (approved April 2023). Accessed online at: management-of-generalised-convulsive-status-epilepticus-in-adults.pdf (scot.nhs.uk) on 7/6/2023.
2. GGC Medicines- Adult Therapeutics Handbook. Published by NHS Greater Glasgow and Clyde, (content last updated September 2022). Accessed online at: <http://handbook.ggcmedicines.org.uk/guidelines/central-nervous-system/guideline-for-phenytoin-dose-calculations/> on 7/6/2023.
3. BNF 84. Accessed online at: <https://www.medicinescomplete.com/#/content/bnf/789756225> on 7/6/2023.
4. Summary of Product Characteristics for Phenytoin 100mg film coated tablets (last updated on 24-02-2023). Accessed online at: <https://www.medicines.org.uk/emc/product/4225> on 7/6/2023.
5. The NEWT guideline for administration of medication to patients with enteral feeding tubes or swallowing difficulties (last updated in August 2019). Accessed online at: [NEWT Guidelines - Drug Monographs - Phenytoin](http://newtguidelines.org.uk/guidelines/phenytoin) on 7/6/2023.

Written by: Claire Anderson

Approved by: Area Drugs and Therapeutics Committee

Updated: June 2023

Review date: June 2026