

Checklist for Phenytoin Pharmacokinetic Dosing and Monitoring¹

Pre-Dosing Background Information Gathering (from chart +/- in discussion with preceptor)

- ✓ Confirm indication (i.e. type of seizure); confirm when last seizure occurred. If seizures pre-existing, confirm frequency of seizures in the past
- ✓ Review for potential drug interactions with current prescription (Rx) meds, non-Rx meds, natural health products (NHPs) and alternative medicines
- ✓ Confirm allergies status
- ✓ Confirm pregnancy & lactation status (If Applicable)
- ✓ Confirm that patient has no contraindications: i.e. allergic to hydantoin products
- ✓ Review patient's previous phenytoin levels, if any, and if doses were altered as a result (check both outpatient and inpatient records)
- ✓ Review the pharmacokinetics of phenytoin via class notes or alternative sources. If available, ask the preceptor for a local phenytoin dosing and monitoring guideline.*

Initial Phenytoin Dosing

- ✓ Establish patient weight, serum creatinine, eGFR, BUN, and albumin
- ✓ Identify target pre-phenytoin level
- ✓ Identify if a phenytoin loading dose is needed
- ✓ Calculate the initial loading dose (if needed), maintenance dose, route and frequency according to patient weight, presence/absence of clinical symptoms, kidney function and ability to adhere to therapy

Therapeutic Drug Monitoring

- ✓ Identify if a phenytoin serum level should be ordered. Here are examples of when levels should be ordered:
 - Recent medication initiation. Typically, a level is taken 2-4 hours post IV or 24 hours post PO load. Generally, a trough level (within 30 to 60 minutes prior to next dosage administration) is then drawn within a few days after phenytoin maintenance dose gets started and will be repeated after 1-2 weeks until there is a steady level of drug in the body (7-21 days). Levels can then be monitored every 3 months to 1 year depending on stability of condition being treated and patient.
 - Recent medication titration (repeat level ~5-7 days following any dose change).
 - Recent change in route of administration (PO, NG, IV)
 - Acute seizure episode
 - Query adherence to medication
 - Query absorption of medication
 - Addition of an interacting medication
 - Presence of dose related side effects or if toxicity presents:
 - Nausea, stomach discomfort
 - Confusion or slurred speech, drowsiness
 - Chest pain or tightness or irregular/pounding heart beat
 - Uncontrolled eye or muscle twitching
 - Light headedness or fainting, dizziness
 - Change in renal function
 - Patient is obese (>125% IBW), pregnant, pediatric or hypermetabolic (e.g. burn patient, cystic fibrosis)
 - Significant changes in hepatic function
- ✓ If a level is necessary, the following should be considered:
 - Is the drug at steady state? (e.g. 7-21 days, note: time to steady state is variable and can range from 3-50 days)
 - Were there any missed doses?
 - Were the previous doses given at the appropriate time? If NG, were feeds held before and after administration?
 - Are there any drug interactions?
 - Is an albumin level available to calculate the corrected level?
 - Is a free phenytoin level required? (e.g. in presence of uremia, dialysis, severe hypoalbuminemia, seizures/toxicity with target concentrations)

Drug Levels Interpretation

- ✓ Interpretation and documentation of the drug level should encompass the following:
 - ID
 - Indication
 - Phenytoin dosing, route of administration, and frequency
 - Date of last dose given/change (if applicable)
 - Subjective
 - Signs and symptoms relevant to phenytoin related adverse effects or lack of efficacy
 - Objective
 - Relevant vitals and laboratory parameters for resulted and pending results (if available)
 - Phenytoin level (include time/date for when the level was drawn)
 - Corrected or free phenytoin level (if applicable)
 - Intended target pre-phenytoin (corrected/free) phenytoin level based on clinical indication
 - Assessment
 - Review the following when interpreting your level and describe how these can affect your interpretation:
 - Missed doses?
 - Previous doses given at the appropriate time?
 - True trough level (within 30minutes to 60 minutes prior to next dosage administration)?
 - Corrected level given other laboratory parameters?
 - Were there interactions with other medications that can affect the interpretation of the level?
 - Is the level subtherapeutic, therapeutic and suprathreshold?
 - Is the level representative of steady state? And how would this affect your interpretation of the drug level?
 - How does this level compare to previous levels the patient has had as an out-patient or inpatient? Is the patient experiencing the same or different level of seizure control now as compared to when those levels were taken?

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- Is dose adjustment or discontinuation needed?
 - What new regimen would you recommend?
 - Is a mini-load needed?
 - Is titration schedule needed?
- Plan
 - Suggestion for subsequent phenytoin dose, route and frequency
 - Suggestion for subsequent monitoring (i.e. repeat drug level or other lab work if necessary)

Subsequent Laboratory Monitoring

- ✓ Repeat level ~5-7 days following any dose change, or if patient seizes, or if toxicity presents, if drug interactions are suspected, or if any significant changes in hepatic function occur.
- ✓ Once desired therapeutic range is reached and there is a steady state of drug level in the body levels can be monitored every 3 months to 1 year as needed.
- ✓ Repeat CBC, LFTs, INR, albumin, serum creatinine as necessary

* If not available at their site, here is a link to a phenytoin empiric dosing guideline: <http://www.vhpharmsci.com/vhformulary/tools/Phenytoin-Kinetic-Monitoring.htm>