



# MINISTRY OF PUBLIC HEALTH & SANITATION

Division of Leprosy, Tuberculosis and Lung Disease (DLTLD)



## THE COMMON SIDE EFFECTS, LIKELY CAUSING AGENTS, AND MANAGEMENT STRATEGIES

| Side affect                                             | Suspected agent(s)                                               | Suggested management strategy                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      | Comments                                                                                                                                                                                                                                                                                                                                            |
|---------------------------------------------------------|------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Seizures                                                | Cs<br>H<br>FQ                                                    | <ul style="list-style-type: none"> <li>Suspend suspected agent</li> <li>Initiate anticonvulsant therapy (e.g. carbamazepine)</li> <li>Restart suspected agent or reinstate suspected agent at lower dose, <b>if essential</b> to the regimen</li> </ul>                                                                                                                                                                                                                                                                            | <ul style="list-style-type: none"> <li>Continue anti depressants to until treatment is completed</li> </ul>                                                                                                                                                                                                                                         |
| Peripheral neuropathy                                   | CS<br>H<br>FQ<br>KM<br>AMK<br>CM<br>E<br>Ethio                   | <ul style="list-style-type: none"> <li>Increase pyridoxine to maximum daily dose (<b>200 mg per day</b>).</li> <li><b>Change</b> parenteral to CM if patient has documented susceptibility to CM.</li> <li>Initiate tricyclic antidepressants e.g. amitriptyline. NSAIDS</li> <li>Lower dose of suspected agent, if this can be done without compromising regimen.</li> </ul>                                                                                                                                                      | <ul style="list-style-type: none"> <li>Patients with co-morbid disease (e.g., diabetes, HIV, alcoholism) may be more likely to develop peripheral neuropathy.</li> <li>Neuropathy is irreversible; some patients may experience improvement when offending agents are suspended.</li> </ul>                                                         |
| Hearing loss                                            | S<br>KM<br>AMK<br>CM                                             | <ul style="list-style-type: none"> <li>Change parenteral to CM if patient has documented susceptibility to CM</li> <li>Injectable should never be stopped before conversion of patients</li> </ul>                                                                                                                                                                                                                                                                                                                                 | <ul style="list-style-type: none"> <li>Audiometry at the initiation of MDR TB therapy.</li> <li>Hearing loss is <b>generally not</b> reversible.</li> </ul>                                                                                                                                                                                         |
| Psychotic symptoms                                      | Cs,<br>H,<br>FQ,<br>Ethio                                        | <ul style="list-style-type: none"> <li>Usually caused by Cs. Withhold suspected agents till symptoms are brought under control.</li> <li>Initiate anti-psychotic drugs e.g. Haloperidol</li> <li>Start Cycloserine at 250mg per day, observe for 5 days. If stable increase to 250mg BD for 5 days. Increase the dose again to 750mg per day. If patient can't tolerate, reduce to where the pat can tolerate. <b>NB. Cycloserine is given in divided doses.</b></li> <li>In case of severe psychosis, replace with PAS</li> </ul> | <ul style="list-style-type: none"> <li>Continue anti-psychotic treatment throughout MDR TB therapy.</li> <li>Prior history of psychiatric disease is <b>not a contraindication Second line drugs</b></li> <li>Psychotic symptoms <b>are generally reversible</b> upon completion of MDR TB treatment or cessation of the offending agent</li> </ul> |
| Depression                                              | Socio-economic circum-stances, chronic disease, CS, FQ, H, Ethio | <ul style="list-style-type: none"> <li>Improve socioeconomic conditions.</li> <li>Group or individual counseling.</li> <li>Initiate antidepressant drugs.</li> <li>Lower dose of suspected agent, if this can be done without compromising the regimen.</li> </ul>                                                                                                                                                                                                                                                                 | <ul style="list-style-type: none"> <li>Socioeconomic conditions &amp; chronic illness are contributing factors to depression.</li> <li>Depressive symptoms may fluctuate during therapy</li> <li>History of prior depression is not a contraindication to the use of the Second line drugs</li> </ul>                                               |
| Hypo-thyroids                                           | PAS<br>Pto/Ethio especially when given in combination            | <ul style="list-style-type: none"> <li>Initiate thyroxine therapy.</li> <li>Thyroxine should be given for till one month after completion of treatment</li> <li>Follow TSH and adjust thyroxine periodically</li> </ul>                                                                                                                                                                                                                                                                                                            | <ul style="list-style-type: none"> <li><b>Completely reversible</b> upon discontinuation of PAS or Ethio.</li> </ul>                                                                                                                                                                                                                                |
| Nausea and vomiting                                     | PAS<br>Pto<br>H<br>E<br>Z<br>CFZ                                 | <ul style="list-style-type: none"> <li>Rehydrate</li> <li>Initiate anti-emetic therapy.</li> <li>Take medication after meals</li> <li>Monitor electrolytes especially potassium and replace</li> </ul>                                                                                                                                                                                                                                                                                                                             | <ul style="list-style-type: none"> <li>Nausea and vomiting are common in early weeks of therapy and usually <b>abate</b> with time supportive therapy.</li> <li>Reversible upon discontinuation of suspected agent.</li> </ul>                                                                                                                      |
| Gastritis                                               | PAS<br>Ethio<br>H<br>E<br>Z<br>CFZ                               | <ul style="list-style-type: none"> <li>Antacids (e.g., calcium carbonate, H2-blockers, proton-pump inhibitors).</li> <li>Dosing of antacids should be taken two hours before or after anti-TB medications.</li> </ul>                                                                                                                                                                                                                                                                                                              | <ul style="list-style-type: none"> <li>Severe gastritis, as manifested by hematemesis, melena, or hematechezia, is rare.</li> <li>Reversible upon discontinuation of suspected agent(s).</li> </ul>                                                                                                                                                 |
| Hepatitis                                               | Z<br>R<br>H<br>Ethio<br>PAS<br>E<br>FQ                           | <ul style="list-style-type: none"> <li>Stop all therapy pending resolution of hepatitis. <i>(If the LFT results shows a &gt;5 times more than the reference range)</i></li> <li>Rule out other potential causes of hepatitis.</li> <li>Re-introduce remaining drugs, one at a time with the <b>LEAST</b> suspected hepatotoxic agents first, while monitoring liver function <i>(see SOPs – Hepatic regimen)</i></li> </ul>                                                                                                        |                                                                                                                                                                                                                                                                                                                                                     |
| Renal failure                                           | S<br>KM<br>AMK<br>CM                                             | <ul style="list-style-type: none"> <li>CM if an aminoglycoside had been the prior parenteral in regimen.</li> <li>Use intermittent dosing while monitoring the creatinine clearance</li> <li>Adjust all TB medications according to the creatinine clearance</li> </ul>                                                                                                                                                                                                                                                            | <ul style="list-style-type: none"> <li>History of diabetes or renal disease is not a contraindication to the use of the agents listed here, although patients with these co-morbidities may be at increased risk for developing renal failure.</li> </ul>                                                                                           |
| Electrolyte disturbance (Hypomagnesaemia & Hypokalemia) | CM<br>KM<br>AMK<br>S                                             | <ul style="list-style-type: none"> <li>Check potassium.</li> <li>If potassium is low, also check magnesium (and calcium if hypocalcemia is suspected).</li> <li>Replace electrolytes as needed as per guideline</li> </ul>                                                                                                                                                                                                                                                                                                         | <ul style="list-style-type: none"> <li><b>If severe hypokalemia is present, consider hospitalization. See SOPS</b></li> </ul>                                                                                                                                                                                                                       |
| Optic neuritis                                          | E                                                                | <ul style="list-style-type: none"> <li>Stop E.</li> <li>Refer patient to an ophthalmologist</li> </ul>                                                                                                                                                                                                                                                                                                                                                                                                                             | <ul style="list-style-type: none"> <li>Usually reverses with cessation of E.</li> </ul>                                                                                                                                                                                                                                                             |
| Arthralgias                                             | Z<br>FQ                                                          | <ul style="list-style-type: none"> <li>NSAIDS.</li> <li>Initiate exercise regimen.</li> <li>Symptoms of arthralgia generally diminish over time, even without intervention</li> </ul>                                                                                                                                                                                                                                                                                                                                              | <ul style="list-style-type: none"> <li>Uric acid levels may be elevated in patients on pyrazinamide. Allopurinol <b>appears not to</b> remediate uric acid levels.</li> </ul>                                                                                                                                                                       |

NB: For further information, refer to the guidelines and standard operating procedures.

