

139 - Treatment other possible options

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136 The Maudsley® Prescribing Guidelines in Psychiatry CHAPTER 1 Treatment – other possible options The large number of proposed treatments for TD undoubtedly reflects the somewhat limited effectiveness of standard remedies, at least before the introduction of valbenazine and deutetrabenazine.¹⁴ Table 1.30 lists some of these putative treatments, most of which have a low level of evidence.^{8,62} Treatment – additional agents Given that there is insufficient evidence to recommend dose reduction as a treatment for TD, and that switching or withdrawing antipsychotic medication is not always effective or advisable, add-on agents are often considered. Evidence--based, pharmacological treatment algorithms for TD have been published.^{41,56} A 2020 meta-analysis⁵⁷ concluded that vesicular monoamine transporter 2 (VMAT-2) inhibitors (such as deutetrabenazine and valbenazine), vitamin E, amantadine and vitamin B6 (pyridoxine) are probably effective treatments. VMAT-2 inhibitors are considered agents of first choice, given the body of evidence supporting their use^{58–60} and their additional antipsychotic action.⁶¹ Table 1.29 describes the most frequently prescribed add-on medications for TD. Table 1.29 First-choice agents prescribed for tardive dyskinesia (alphabetical order; no preference implied). Drug Comments Amantadine^{56,57,62–64} Rarely used and evidence for efficacy is limited. Dose is 100–300mg/day. Benzodiazepines^{37,38} Widely used for TD, but a Cochrane review considered that the limited evidence for efficacy was inconclusive.⁶⁵ Intermittent use may be necessary to avoid tolerance to effects. Most commonly used are clonazepam 1–4mg/day and diazepam 6–25mg/ day, with better supporting evidence for clonazepam.^{42,66} Deutetrabenazine^{8,56,58,60,67,68} Deutetrabenazine (a VMAT-2 inhibitor) is effective as a treatment for TD. Licensed for TD in the USA.⁶⁹ Better supporting evidence than for tetrabenazine. Longer half-life than tetrabenazine but still needs to be taken twice a day (a once daily slow-release tablet is in development).⁷⁰ Low incidence of psychiatric and neurological effects. Dose is 12–48mg/day. Ginkgo biloba^{56,71,72} Well tolerated. A Cochrane review concluded that while Ginkgo biloba could reduce TD symptoms, the available evidence did not justify its routine use as a treatment.⁷³ A meta-analysis of three Chinese RCTs showed a good effect with 240mg/day.⁷⁴ Pyridoxine⁷⁵ Supported by a Cochrane review⁷⁶ and a 2020 meta-analysis.⁵⁷ Dose is up to 400mg/day. Tetrabenazine^{77,78} The only licensed treatment for moderate to severe TD in UK. Depression, drowsiness, parkinsonism and akathisia may occur.^{66,79} Dose is 25–200mg/day. Reserpine (similar mode of action) also effective but rarely, if

ever, used. Valbenazine^{8,59,67,73,80} Evidence supports a favourable benefit-risk ratio for valbenazine (VMAT-2 inhibitor) as a treatment for TD. Licensed for TD in the USA.⁸¹ A dose of 80mg once daily is effective. It has a benign cardiovascular profile and a low incidence of depression and akathisia. Vitamin E^{57,82} Numerous studies but efficacy remains to be conclusively established. A Cochrane review suggested that there is evidence only for slowing the deterioration of TD,^{8,83} but a 2022 meta-analysis also suggested a treatment effect.⁸⁴ Dose is in the range 400–1600 IU/day.

Revision #1

Created 2026-01-04 20:12:46 UTC by Omar Ayman

Updated 2026-01-04 20:12:46 UTC by Omar Ayman