



ELSEVIER

Contents lists available at ScienceDirect

Current Therapeutic Research

journal homepage: www.elsevier.com/locate/cuthre

Mebendazole Is a Potential Alternative in the Treatment of *Giardia duodenalis* Infection



To the Editor:

Parasites are widely distributed throughout the world and represent an important cause of morbidity and mortality, mainly in tropical and subtropical regions.¹ Most of those parasites exist and persist for social and economic reasons that enable pathogens to take advantage of changes in behavioral and physical environments.

Giardia duodenalis (also known as *Giardia lamblia* and *Giardia intestinalis*) is the most common intestinal pathogenic protozoan infection reported in humans, and the disease it causes, giardiasis, is now included in the neglected diseases initiative. Approximately 280 million people are infected each year with this parasite, but this estimation could be low considering that infection rates are higher in endemic countries where diagnostic facilities and reporting systems are unavailable or not functional.

The lack of any useful parasite vaccine means that prevention of this and many other parasitic diseases continues to be based on ecologic measures aimed at interrupting the biological cycle of the parasite combined with the use of antiparasitic drugs.

Mebendazole (MBZ) has been used worldwide because of its relatively poor absorption from the intestine, low level of adverse events, and broad spectrum of action against soil-transmitted helminths, even in single doses. The low cost, effectiveness, lack of action on intestinal microbiota, and the safety of this drug further enhances its therapeutic appeal. MBZ has been evaluated for its potential use against protozoan and helminth infections other than the common soil-transmitted helminths, providing some evidence that encourages scientists to use it in certain situations, such as for cases of treatment failure or resistance.²

Some in vitro studies have demonstrated MBZ activity against *G duodenalis* infection. The authors of these studies showed that this drug affects the growth of the protozoa, inducing trophozoite detachment and distortion of both morphology and general structure.^{7–10}

Based on these in vitro studies, our group carried out 4 clinical trials in Cuba—3 in children and 1 in adults—evaluating the efficacy and safety profile of MBZ in the treatment of *G duodenalis*. These studies offered clear evidence of the effectiveness and tolerability of MBZ to treat *G duodenalis*.^{3–6}

Three days' treatment with MBZ was similar in efficacy to the first-line drugs used for the treatment of giardiasis, both in children and in adult patients.

After 8 years of extensive clinical use, we have found MBZ (200 mg 3 times daily for 3 days)³ to be excellent in treating children aged 5 to 15 years mono infected with *G duodenalis*. From the beginning of 2008 to the end of 2015, of 522 children attending the

Centre of Hygiene, Epidemiology, and Microbiology seeking treatment for this intestinal protozoa, 450 (86.2%) were healed. The efficacy of the chemotherapy was assessed by microscopic examination (as direct wet mounts and after formol–ether concentration) of fecal samples collected soon (ie, 3, 5, and 7 days) after treatment completion, to avoid the bias that would be introduced by reinfection. A child was only considered cured if no *Giardia* trophozoites or cysts could be found in any of the 3 posttreatment fecal specimens.

The drug was well tolerated. Only mild, transient, and self-limited side effects were reported and these did not require discontinuation of treatment. Abdominal pain (29 out of 522 patients; 5.6%), nausea (15 out of 522 patients; 2.9%), and vomiting (12 out of 522 patients; 2.3%) were the only side effects reported.

Additionally, from 2010 to the end of 2015, 423 adults mono infected by *G duodenalis* were also treated successfully using MBZ (200 mg 3 times daily for 3 days).⁶ Similar to the children group, MBZ was an exceptional option. Of 423 patients treated, 392 (92.7%) were cured. In this group the only adverse effect reported was abdominal pain in 26 out of 423 patients (6.2%).

The effective management of *G duodenalis* infection has been considered problematic, especially in tropical and subtropical areas. Based on our clinical trials as well as anecdotal experience, MBZ could be an excellent alternative in the treatment of *G duodenalis* infections both in children and in adults.

Acknowledgment

The author thanks all patients who generously participated in the studies referred to in this letter.

Roberto Cañete, MD, MS, PhD*

Council of Scientific Societies of Health Director

Full Professor and Senior Researcher

University of Medical Sciences and Centre of Hygiene

Epidemiology, and Microbiology, Matanzas, Cuba

E-mail address: roberto.villafranca@infomed.sld.cu

* Address correspondence to: Roberto Cañete Villafranca, Council of Scientific Societies of Health and Parasitology, University of Medical Sciences and Centre of Hygiene, Epidemiology, and Microbiology, Calle Milanés Esquina a Buena Vista, Matanzas, Matanzas 40100 Cuba.

References

- [1] Cañete R, Díaz MM, Avalos García R, et al. Intestinal parasites in children from a day care centre in Matanzas City, Cuba. *PLoS One*. 2012;7:e51394.
- [2] Cañete R, Escobedo AA, Almirall P, et al. Mebendazole in parasitic infections other than those caused by soil-transmitted helminths. *Trans R Soc Trop Med Hyg*. 2009;103:437–442.
- [3] Escobedo AA, Cañete R, González ME, et al. A randomized trial comparing mebendazole and secnidazole for the treatment of giardiasis. *Ann Trop Med Parasitol*. 2003;97:499–504.
- [4] Cañete R, Escobedo AA, González ME, et al. A randomised, controlled, open-label trial of a single day of mebendazole versus a single dose of tinidazole in the treatment of giardiasis in children. *Curr Med Res Opin*. 2006;22:2131–2136.
- [5] Cañete R, Escobedo AA, González ME, Almirall P. Randomized clinical study of five days therapy with mebendazole compared to quinacrine in the treatment of symptomatic giardiasis in children. *World J Gastroenterol*. 2006;12:6366–6370.
- [6] Almirall P, Escobedo AA, Ayala I, et al. Mebendazole compared with secnidazole in the treatment of adult giardiasis: a randomised, no-inferiority, open clinical trial. *J Parasitol Res*. 2011;2011:636857.
- [7] Katiyar SK, Gordon VR, McLaughlin GL, et al. Antiprotozoal activities of benzimidazoles and correlations with beta-tubulin sequence. *Antimicrob Agents Chemother*. 1994;38:2086–2090.
- [8] Chávez B, Cedillo-Rivera R, Martínez-Palomo A. *Giardia lamblia*: ultrastructural study of the in-vitro effect of benzimidazoles. *J Parasitol*. 1992;39:510–515.
- [9] Morgan UM, Reynoldson JA, Thompson RC. Activities of several benzimidazoles and tubulin inhibitors against *Giardia* spp. in vitro. *Antimicrob Agents Chemother*. 1993;37:328–331.
- [10] Cedillo-Rivera R, Munoz O. In vitro susceptibility of *Giardia lamblia* to albendazole, mebendazole and other chemotherapeutic agents. *J Med Microbiol*. 1992;37:221–224.