## Accepted Manuscript

Title: Two-Fold Excess of Fluoride in the Drinking Water Has No Obvious Health Effects Other Than Dental Fluorosis

Authors: Batın İlgit Sezgin, Şirin Güner Onur, Ali Menteş, Alev Eda Okutan, Eda Haznedaroğlu, Alexandre Rezende Vieira

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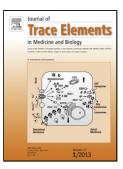
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## ACCEPTED MANUSCRIPT

Two-Fold Excess of Fluoride in the Drinking Water Has No Obvious Health Effects Other Than Dental Fluorosis

Short title: Excess of Fluoride in the Water Has No Long-Term Health Effects

Batın İlgit Sezgin, D.D.S.<sup>1</sup> Şirin Güner Onur, D.D.S., Ph.D<sup>2</sup> Ali Menteş, D.D.S., Ph.D<sup>1</sup> Alev Eda Okutan, D.D.S.<sup>1</sup> Eda Haznedaroğlu, D.D.S., Ph.D<sup>1</sup> Alexandre Rezende Vieira, D.D.S., M.S., Ph.D<sup>3</sup>

batinilgit@gmail.com sirin\_guner@yahoo.com alimentes@gmail.com alevedaokutan@gmail.com ehaznedaroglu@marmara.edu.tr alexandre\_vieira@pitt.edu

Corresponding author: Ali Menteş Email: alimentes@gmail.com

Address: Marmara Üniversitesi Başıbüyük Sağlık Yerleşkesi, Dişhekimliği Fakültesi, Başıbüyük Yolu 9/3

34854 Başıbüyük / Maltepe / İstanbul / Türkiye

Phone: +905323441995

#### Abstract

Background: There is concern that fluorides in the drinking water is hazardous to health.

Methods: We conducted an observational study in the village of Hanliyenice (population 280), Turkey, which has 2.5 times higher than optimal levels of fluoride in the drinking water and evaluated all children 7 to 13 years of age (N=30). We collected information on dental decay, fluorosis, daily water consumption and diet, child history and her family history of cancer, cardiovascular risks/diseases, and asthma, and obtained a blood sample for extraction of genomic DNA. We genotyped ten single nucleotide polymorphisms in aquaporins.

Results. As expected, a high number of children were dental caries free (19 out of 30) and had fluorosis (25 out of 30). Family history of cancer, cardiovascular events, and asthma was not different from the expected figures based on Turkey. One variant just upstream of AQP5 was associated with being fluorosis free. (G allele of AQP5 rs296763, p=6.0E<sup>-6</sup>).

Conclusions: Exposure to levels of fluoride twice as high than the optimum in the drinking water increases the prevalence of fluorosis, dramatically decreases dental caries, and does not increase the risk of cancer, cardiovascular events, and asthma.

Key Words: Fluorosis; Cardiovascular Disease; Cancer; Asthma; Dental Caries; Aquaporin

<sup>&</sup>lt;sup>1</sup>Department of Pediatric Dentistry, School of Dentistry, Marmara University, Istanbul, Turkey

<sup>&</sup>lt;sup>2</sup>Department of Pediatric Dentistry, School of Dentistry, Trakya University, Edirne, Turkey

<sup>&</sup>lt;sup>3</sup>Department of Oral Biology, School of Dental Medicine, University of Pittsburgh, Pittsburgh, USA

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