# Eye cancer in a young male with a vaping history

#### Carol L Shields, Min Kim, Sara E Lally, Patricia Chévez-Barrios<sup>1</sup>, Jerry A Shields

We report a new observation of conjunctival intraepithelial neoplasia (CIN) in a young man following years of electronic cigarette use. A 22-year-old man with a 5-year electronic cigarette use (vaping) developed painless unilateral blurred vision in the right eye from mild superficial corneal opacification, unresponsive to topical antiviral therapy. Corneal scraping documented no infectious etiology. The abnormality persisted for 1 year and superficial keratectomy revealed high-grade CIN with enlarged pleomorphic and dyskeratotic cells. Interferon-alpha-2b was instituted. In this case, chronic exposure to electronic cigarette vapors (vaping) could have been associated with CIN in this young patient.

Key words: Cancer, conjunctiva, electronic cigarettes, eye, squamous cell carcinoma, vaping

Electronic cigarettes were introduced in the United States in 2006 and touted as a safer alternative to standard cigarettes. This product has become popular, especially among young adults. Electronic cigarettes produce vapor rather than smoke, hence the activity is termed vaping rather than smoking.

Within the past year, vaping-related serious lung disease (acute respiratory distress) has surfaced, often requiring intensive care monitoring on a ventilator.<sup>[1]</sup> The first vaping-related death was reported in August 2019 and there have been over 1000 vaping-related illnesses, most often in young males ≤35 years of age.

Standard cigarette use is well-documented to impact risk for head and neck cancer, including conjunctival malignancies.<sup>[2]</sup> There is limited literature on cancer related to electronic cigarette use.<sup>[3,4]</sup> A systematic review of articles addressing electronic cigarettes and head/neck cancers found that vaping exposure could lead to DNA damage with increased

Access this article online	
Quick Response Code:	Website:
	www.ijo.in
	<b>DOI:</b> 10.4103/ijo.IJO_27_20

Ocular Oncology Service, Wills Eye Hospital, Thomas Jefferson University, Philadelphia, PA, <sup>1</sup>Department of Pathology and Genomic Medicine, Houston Methodist Hospital, Houston TX, USA

Correspondence to: Dr. Carol L Shields, Ocular Oncology Service, 840 Walnut Street, Suite 1440, Philadelphia, PA - 19107, USA. E-mail: carolshields@gmail.com

Received: 21-Jan-2020Revision: 18-Mar-2020Accepted: 28-Mar-2020Published: 24-Jul-2020

oxidative stress, especially in flavored products.<sup>[3]</sup> Electronic cigarettes aerosols can promote DNA damage and impair DNA repair with potential for carcinogenesis.<sup>[4,5]</sup> A PubMed search for < electronic cigarette eye>, <electronic cigarette conjunctiva>, and < electronic cigarette eye cancer > yielded corneal chemical toxicities with tear film abnormalities, but no case of malignancy. Herein, we describe vaping-related conjunctival squamous neoplasia in a young male.

# **Case Report**

A male in his early 20s noted painless blurred vision and was found to have possible viral keratitis, which did not respond to oral antiviral therapy. One year later, corneal scraping revealed atypical epithelial cells and polymerase chain reaction (PCR) documented the lack of infectious etiology. Subsequently, 1 year later, superficial keratectomy demonstrated full-thickness atypia with mitotic activity, enlarged pleomorphic and dyskeratotic cells, and thickened basement membrane, suggesting conjunctival intraepithelial neoplasia (CIN) with high-grade features. [Fig. 1] Immunohistochemistry surrogate marker for HPV-driven lesions (p16) was negative. Interferon-alpha-2b (1 million international units/cc) was instituted and the patient was referred to us.

On examination, visual acuity was 20/30 in the right eye (OD) and 20/25 in the left eye (OS). There was limbal pannus with diffuse epithelial thickening and opacification OD involving 95% of the corneal surface, sparing only a superonasal sector. [Fig. 2] A subtle gelatinous limbal mass was noted superotemporally, measuring 8 mm in diameter. Anterior segment optical coherence tomography confirmed epithelial thickening of 110–180  $\mu$  (normal 52.2–54.6  $\mu$ ). [Fig. 2] The left eye was normal.

The patient denied immune suppression, human immunodeficiency virus (HIV), organ transplantation, and xeroderma pigmentosum. He admitted to vaping flavored (mint) electronic cigarettes for 5 continuous years (5 vapes per day or more). He denied cigarette smoking. Topical interferon was continued for 3 additional months.

## Discussion

In an analysis of 5002 conjunctival tumors, squamous neoplasia, (CIN and squamous cell carcinoma (SCC)), represented 715 (14%) cases.<sup>[6]</sup> This malignancy most often occurred in older adults (>60 years) (n = 494/715, 69%) or mid-adults (41–60 years) (n = 181/715, 25%) and rarely occurred in young adults (21–40 years) (n = 38/715, 5%) or children ( $\leq$ 20 years) (n = 2/715, <1%),<sup>[6]</sup> typically those

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**Cite this article as:** Shields CL, Kim M, Lally SE, Chévez-Barrios P, Shields JA. Eye cancer in a young male with a vaping history. Indian J Ophthalmol 2020;68:1699-701.



**Figure 1:** Intraepithelial corneal neoplasia. (a) Histopathology demonstrates corneal full-thickness atypia, loss of cellular polarity, and dyskeratosis (eosinophilic cells). (Hematoxylin-eosin, original magnification 20×). (b) Higher magnification shows pleomorphic cells and frequent atypical mitoses (arrows) in the superficial layers of the epithelium. (Hematoxylin-eosin, original magnification 40×). (c) PAS stain highlights the abnormally thickened basement membrane and shows the atypical mitosis (arrow). (Periodic Acid Schiff, original magnification 40×). (d) Full-thickness epithelial proliferation is demonstrated by Ki67 nuclear cell proliferation marker (brown nuclei). (Mib-1 antibody, DAB chromogen, immunohistochemistry original magnification 40×)



**Figure 2:** (a) Corneal epithelial opacification from intraepithelial neoplasia (arrows) is seen on the entire corneal surface, sparing a superonasal sector. (b) On the anterior segment, optical coherence tomography, the corneal epithelial thickening (arrows) demonstrated smooth surface, increased optical reflectivity, and thickness of 110–80  $\mu$ 

with immune suppression, HIV, organ transplantation, or xeroderma pigmentosum, not found in this case.

#### Conclusion

To our knowledge, the squamous neoplasia in this young male could be related to the 5-year chronic vapor exposure from electronic cigarettes. Although more data is needed to prove this correlation, we understand that e-cigarettes contain variable amounts of potential carcinogens that could result in malignancy, as seen in this young patient. All patients with squamous neoplasia should be examined regarding the use of standard or electronic cigarettes.

Financial support and sponsorship

### Nil.

**Conflicts of interest** 

There are no conflicts of interest.

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