# Letter to the Editor: Reversal in Thyroid Cancer Incidence Trends in the United States, 2000–2017

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## **Dear Editor:**

THE INCIDENCE OF THYROID CANCER in the United States L increased exponentially between 1974 and 2013, at a rate of 3.6% per year (1). Rates of thyroid cancer diagnosis have been found to be highly sensitive to clinical behavior, such as how widely diagnostic imaging is used. This was most clearly demonstrated by the South Korean experience, where screening asymptomatic persons with thyroid ultrasound led to a nearly 15-fold surge in thyroid cancer incidence for an 18-year period (1993–2011), and subsequently in 2014, a rapid decrease of 30% after public education on overdiagnosis and overtreatment (2). In the United States, the incidence of thyroid cancer plateaued for the first time in 2014-2016, possibly reflecting changes in diagnostic intensity (3). In 2009, the American Thyroid Association (ATA) clinical practice guidelines first recommended against biopsy of most subcentimeter or small, radiographically low-risk thyroid nodules (4). Subsequently, in 2015, ATA published revised guidelines further refining this recommendation (5). In this study, we examined if thyroid cancer incidence in the United States has changed in recent years.

The Weill Cornell Medicine Institutional Review Board deemed this study exempt from review. Annual thyroid cancer incidence (all histological types, stratified by size) between 2000 and 2017 was obtained from the Surveillance, Epidemiology, and End Results (SEERs) 18 registry using SEER\*Stat (National Cancer Institute). SEER-18 captures cancers diagnosed in ~27% of the U.S. population. Segmented log-linear regression was used to identify breakpoints in incidence trends and mean annual percentage change (APC) over time segments. *p*-Values represent two-tailed probabilities of the *t*-distribution of APC, including 0.

Thyroid cancer incidence (Fig. 1) increased from 7.5 per 100,000 (age-adjusted to the 2000 U.S. population) in 2000 to 13.5 per 100,000 in 2009 (APC: 7.1%, 95% confidence interval [CI 6.8–7.5%], p < 0.001). The rate of increase slowed from 2009 to 2015 (APC: 2.7% [CI 2.0–3.4%], p < 0.001), reaching an incidence of 14.9 per 100,000, and for the first time, decreased at a statistically significant rate from 2015 to

2017 (APC: -4.6% [CI -7.5% to -1.6%], p = 0.007), reaching an incidence of 13.7 per 100,000. After stratifying by size, a statistically significant decrease was seen in cancers  $\leq 1$  cm from 2013 to 2017 (APC: -3.3 [CI -5.8% to -0.8%], p = 0.016), but not in cancers >1 cm from 2015 to 2017 (APC: -2.0 [CI -6.0% to 2.2%], p = 0.312).

After three decades of exponential growth, for the first time, thyroid cancer incidence in the United States has begun to decrease at a statistically significant rate. We caution that observational data alone cannot rule out changes in environmental or other risk factors as causes for these changes. However, the temporal relationship with recent clinical management guidelines (4,5), and the similar reversal of trends in South Korea (2), suggest that changing clinical practices are likely to have led to the recent decline in thyroid cancer diagnoses. A similar reversal was observed with prostate cancer incidence in the 1990s, after adoption and subsequent decrease of widespread prostate-specific antigen screening (6). The reclassification of some cancers as noninvasive follicular thyroid neoplasm with papillary-like nuclear features (NIFTP) in 2016 may also have contributed to some of the decrease in incidence in the final year of the study (7). In addition, the inclusion of active surveillance of small cancers as an acceptable alternative to surgery in the 2015 ATA guidelines may also have begun to affect some biopsy decisions for small nodules, although the impact of this is not yet known.

The contemporary trends in thyroid cancer incidence parallel the introduction of radiographic features and size as risk-stratification criteria. Future data will help establish if these declining trends remain durable as clinical practice continues to evolve.

#### **Author Disclosure Statement**

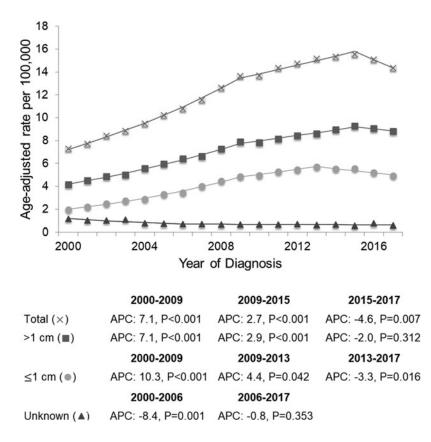
The Morris laboratory receives research funding from Illumina, Inc. and AstraZeneca for research unrelated to this study.

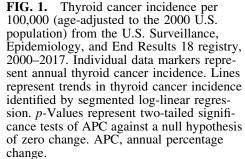
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