

#### Distribution of Expenses Associated with THC Residency Training, Academic Year 2013–2014.

Faculty compensation includes salary plus benefits associated with a full-time-equivalent position dedicated to residency administration, precepting outpatient and inpatient service, allocation of clinic-administration time on the basis of volume of residents' patient service, and precepting contracts paid by the residency. Resident compensation includes salary and benefits for residents. Clinical service administration and operational costs include inpatient administration and outpatient operational and administration costs allocated to residents' patient visits. Other educational costs include residency personnel, educational supplies, information technology, occupancy and other residency administration items, and residency overhead. In-kind costs include all items in other educational costs supported by hospital precepting contracts and community partners. Numbers do not add to 100 because of rounding.

The costing data make it clear that residents contribute to THC's finances, as well as to the care of patients — a reality that needs to be taken into account in establishing the true cost of residency training to a sponsoring institution.

As community-based practices that rely largely on ambulatory care reimbursement for financial viability, most THC-sponsoring organizations cannot support residency programs without specific, adequate, and stable funding. Whereas teaching hospitals need residents to assist with clinical coverage for acute and inpatient care, community-based ambulatory care practices don't have analogous coverage demands. For THCs, initiating residency programs is an elective decision that can't be considered if it jeopardizes the practice. The decision by the secretary of health and human services to provide \$150,000 per resident per year recognized this reality and established a funding level that, absent hard data, met the best estimates available at the time. Our findings suggest that \$150,000 per resident per year is reasonably reflective of the true

current cost of a resident to a community-based sponsor.

The views expressed in this article are those of the authors and do not necessarily reflect those of the Health Resources and Services Administration, Department of Health and Human Services, or the U.S. government.

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## Worldwide Thyroid-Cancer Epidemic? The Increasing Impact of Overdiagnosis

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Several reports have described dramatic increases over recent decades in the incidence of thyroid cancer,<sup>1</sup> predominantly small papillary carcinomas,<sup>2</sup> even as thyroid-cancer-related mortality rates have not changed substantially.<sup>3</sup> The largest increase has been observed in South Korea:

the incidence among people 15 to 79 years of age (standardized to the world population) increased from 12.2 cases per 100,000 persons in 1993–1997 to 59.9 cases per 100,000 persons in 2003–2007,<sup>1</sup> making thyroid cancer the most commonly diagnosed cancer among women in that country.

The introduction of new diagnostic techniques (ultrasonography, computed tomography, and magnetic resonance imaging), combined with increased medical surveillance and access to health care services, can lead to massive increases in detection of small papillary lesions caused by the

large reservoir of asymptomatic, nonlethal disease known to exist in the thyroid gland.<sup>1,2</sup> Sudden changes in thyroid-cancer incidence have also accompanied large-scale thyroid-gland surveillance in purportedly high-risk populations. In Japan's Fukushima Prefecture, thyroid-cancer incidence among screened children and adolescents was approximately 30 times as high as the national average only a few months after intensive screening programs for these age groups began in response to the 2011 nuclear accident.<sup>4</sup>

Studies comparing thyroid-cancer incidence during different periods and across countries can help in estimating overdiagnosis — that is, diagnosis of thyroid tumors that would not, if left alone, result in symptoms or death. Here, we provide a measure of thyroid-cancer overdiagnosis over the past two decades in selected high-income countries, based on recently developed methods and high-quality cancer-registry data (Cancer Incidence in Five Continents, International Agency for Research on Cancer, available at <http://ci5.iarc.fr/CI5I-X>). Using as reference populations those in countries with long-standing cancer registries, we deduced the shape of age-specific trends in thyroid-cancer incidence in the 1960s, before ultrasonography was introduced in the late 1970s.<sup>1</sup> The “historical” age curves were remarkably similar across populations, showing exponential growth of rates with age, a finding consistent with the multistage model of carcinogenesis described by Armitage and Doll and with the behavior of most epithelial cancers. The historical age curve was used to estimate the number of cases that would

have been expected if thyroid cancer had continued to be detected primarily through palpation.

The shape of the age-specific curves has changed dramatically since the 1980s, albeit to different extents in various countries (see graphs). Incidence has progressively increased among middle-aged women (and men; see Fig. S1 in the Supplementary Appendix, available at [NEJM.org](http://NEJM.org)) but has varied to a much lesser extent at older ages, thus altering age curves over the years roughly from an exponential-growth shape to an inverted-U shape.

We attribute the excess cases beyond the number predicted by the multistage model to diagnoses of asymptomatic disease subsequent to improved diagnostic technology and increased surveillance, predominantly in young or middle-aged populations.<sup>1</sup>

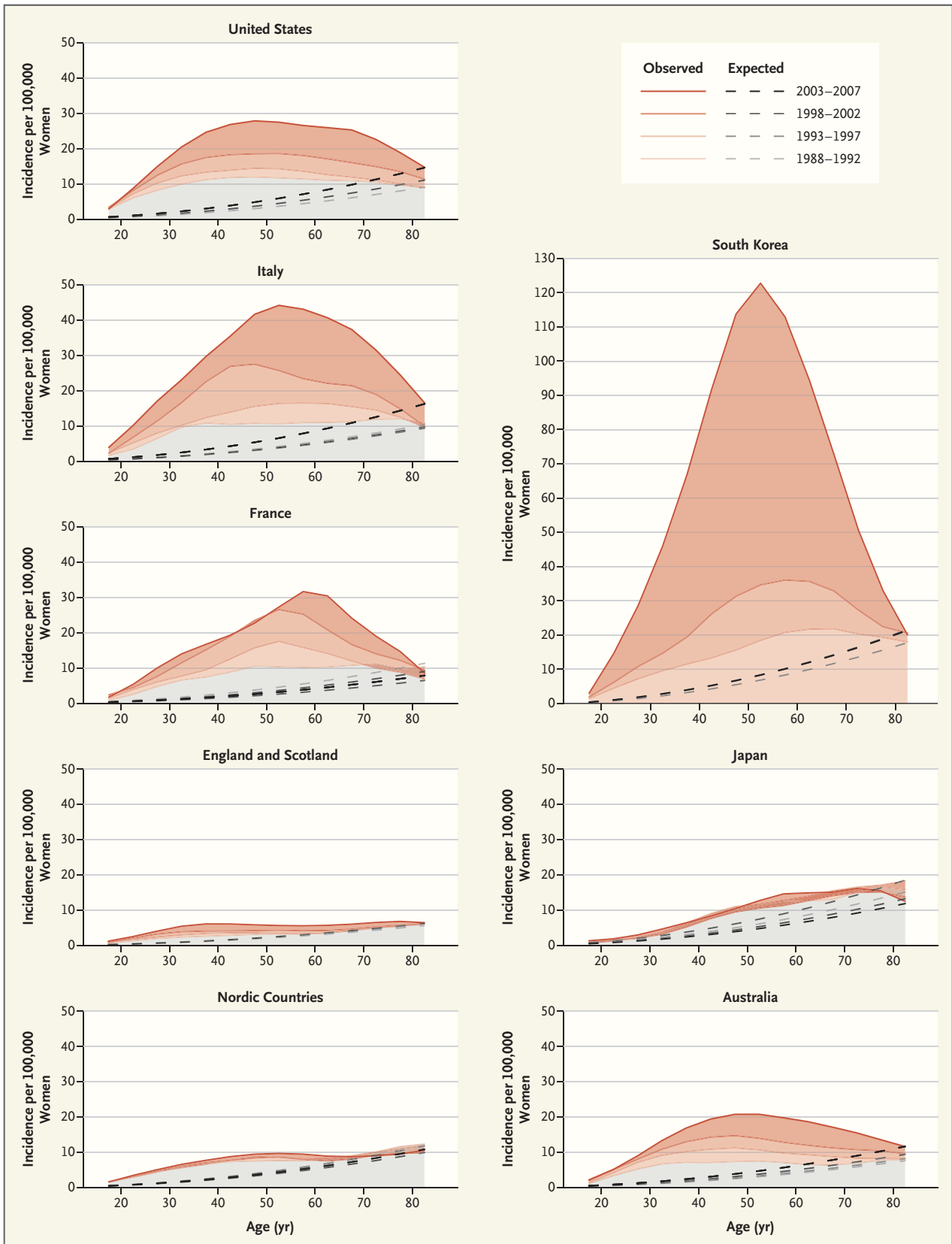
Large changes in observed age curves for women emerged in the countries most affected by the thyroid-cancer “epidemic,” including the United States, Italy, and France. A striking increase was observed among South Korean women 50 to 59 years of age, from about 35 cases per 100,000 women in 1998–2002 to more than 120 cases per 100,000 women in 2003–2007. Indeed, since 1999, approximately 13% of South Korean adults had voluntarily undergone thyroid ultrasonography within the framework of organized screening programs for five other types of cancer, and the highest rate of participation (26%) was among women 50 to 59 years of age.<sup>1</sup> Conversely, increases in incidence in the United States, Australia, and Italy began in the 1980s, particularly among women younger than 45 years, around the time of the earliest introduction of

ultrasonography in gynecologic and obstetric clinics, which favored opportunistic examinations of the thyroid gland in women of reproductive age.

The estimated number of thyroid-cancer cases attributable to increased thyroid-gland surveillance is also striking and has increased markedly over time. If there had been no changes in the shape of the age curve (estimated on the basis of data from the pre-ultrasonography period), then approximately 228,000 cases diagnosed in women in the United States between 1988 and 2007 would be considered overdiagnoses (see Fig. S2 in the Supplementary Appendix). Corresponding numbers are 65,000 in Italy, 46,000 in France, and 36,000 in Japan. Among South Korean women, overdiagnosis accounted for approximately 77,000 extra cases of thyroid cancer between 1993 and 2007. The number of overdiagnoses was smaller but still substantial in Australia (10,000), England and Scotland (7000), and the Nordic countries (Denmark, Finland, Norway, and Sweden; 6000).

Overdiagnosis increased in many countries between 1988 and 2007 (see graphs and the Supplementary Appendix). We don't know whether this trend has continued, since post-2007 data were not available. However, if we take the most recent available period, 2003–2007, as typical of current practice, we estimate that overdiagnosis in women accounts for 90% of thyroid-cancer cases in South Korea; 70 to 80% in the United States, Italy, France, and Australia; and 50% in Japan, the Nordic countries, and England and Scotland.

A similar pattern of age curves was observed for men, al-



**Facing page: Observed versus Expected Changes in Age-Specific Incidence of Thyroid Cancer per 100,000 Women, 1988–2007.**

The observed rates were derived from Cancer Incidence in Five Continents, International Agency for Research on Cancer (available at <http://ci5.iarc.fr/CIS1-X>). The expected rates were based on the observation that before the introduction of ultrasonography and other novel diagnostic techniques, thyroid-cancer incidence increased exponentially with age in all countries with available long-term data, in keeping with the multistage model of carcinogenesis described by Armitage and Doll (rate proportional to age<sup>k</sup>, where the exponent *k* is to be estimated from incidence data). For each 5-year period, the expected age-specific rates were obtained by hypothesizing that the disease would have retained the historical age curve described by the multistage model. Since thyroid-cancer incidence varied only minimally across periods among people 80 to 84 years of age, we added a constraint that sets as equal the expected and observed incidence rates for this age group. We hypothesized that the progressive departure of the observed rates from the multistage model was attributable to the increased detection of asymptomatic, nonlethal disease — that is, overdiagnosis.

though it was less pronounced than the pattern for women and the peaks occurred at older ages. The estimated number of overdiagnosed cases is much lower than that among women. The proportion of cases of thyroid cancer in men that we estimate to be overdiagnoses in 2003–2007 is approximately 70% in France, Italy, and South Korea; 45% in the United States and Australia; and less than 25% in all other countries we examined (see Fig. S1 and S2 in the Supplementary Appendix).

Overall, we estimate that more than 470,000 women and 90,000 men may have been overdiagnosed with thyroid cancer over two decades in these 12 countries, with steady incremental increases over time and little evidence of stabilization. Therefore, there does not appear to be a limit to the identification of asymptomatic thyroid cancer at present, even though the related

mortality rates have remained stable or declined slightly.<sup>3</sup>

There is no evidence of new risk factors or increased exposure to known contributors to thyroid cancer<sup>1,2</sup> that might explain these steep upward trends or the differences among otherwise similar or geographically close high-income countries. Changes in patterns of somatic mutations in thyroid cancer over time have been reported<sup>2</sup> and may reflect the increasing proportion of papillary carcinomas after the introduction of ultrasonography. In addition, possible changes in exposure to risk factors such as diagnostic radiation, overweight, or diabetes may themselves be correlates of increased medical surveillance.<sup>2</sup> Changes in access to health care, physicians' practices, and the extent of intentional inspection of the thyroid gland or incidental findings are therefore the most likely explanations of our findings.

It's important to bear in mind that the vast majority of patients who received a diagnosis of thyroid cancer in the countries we studied underwent total thyroidectomy, and a high proportion also received other harmful treatments (neck lymph-node dissection and radiotherapy) — practices recently discouraged in the guidelines of the American Thyroid Association. Furthermore, studies from Japan have shown that immediate surgery and watchful waiting are equally effective in averting deaths from thyroid cancer: only a small minority (3.5%) of the 1235 patients with papillary microcarcinomas who were followed for an average of 75 months had clinical progression of disease, and none died.<sup>5</sup> The examples of South Ko-

rea, the United States, Italy, and France suggest that other countries should exercise caution against systematic screening for thyroid cancer and overtreatment of small nodules (<1 cm) and suggest that watchful-waiting approaches should be considered a research priority and a preferable option for patients with low-risk papillary thyroid cancers.

Finally, the enormous increase in thyroid-cancer incidence in South Korea subsequent to opportunistic ultrasonography-based screening sends a strong warning about data interpretation in the context of large-scale screening of the thyroid gland after radiation exposure from exceptional events like the nuclear accident in Fukushima.<sup>4</sup>

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