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Theoretical increase of thyroid cancer induction from cervical spine multidetector computed tomography in pediatric trauma patients

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Abstract

BACKGROUND—The trend of increasing cervical spine multidirectional computed tomography (MDCT) imaging of pediatric trauma patients is characteristic of the overall dramatic increase in computed tomography utilization in the United States. The purpose of this study is to compare the amount of radiation a pediatric trauma patient absorbs to the thyroid from plain radiographs and MDCT of the cervical spine and to express risk by calculation of theoretical thyroid cancer induction.

METHODS—A retrospective evaluation of pediatric trauma patients admitted from October 1, 2004, to October 31, 2009, was performed at an academic, Level I trauma center. Inclusion criteria were Level I/II trauma patients, cervical spine imaging performed at our institution, and age <18 years. Absorbed thyroid radiation was calculated for patients receiving plain radiographs or MDCT. Thyroid cancer risk was calculated using the 2006 Biological Effects on Ionizing Radiation VII report.

RESULTS—Six hundred seventeen patients met inclusion criteria: 224 received cervical spine radiographs and 393 received cervical spine MDCT. The mean thyroid radiation absorbed from radiographs was 0.90 mGy for males and 0.96 mGy for females compared with 63.6 mGy (males) and 64.2 mGy (females) receiving MDCT ($p < 0.001$). The median excess relative risk of thyroid cancer induction from one cervical spine MDCT in males was 13.0% and females was 25.0%, compared with 0.24% (males) and 0.51% (females) for radiographs ($p < 0.001$).

CONCLUSIONS—The significant difference in radiation that MDCT delivers to the pediatric trauma patient when compared with plain radiographs should temper routine use of computed tomography in pediatric cervical spine clearance algorithms.

> Cervical spine injury occurs in 0.98% to 2.2% of pediatric blunt trauma patients^{1–5} necessitating a methodical screening process to avoid the potential devastation of a missed

DISCLOSURE

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injury.⁶ In patients who cannot be cleared on clinical grounds, screening with plain radiographs has historically been the recommended imaging modality. However, because of availability and improved sensitivity, computed tomography (CT) use for pediatric cervical spine screening has increased nearly fourfold.⁷ The comparatively higher doses of radiation from CT in pediatric trauma patients may be associated with untoward complications. $8-10$

Sixty-seven million CT scans are ordered annually as of 2006, accounting for 24% of the total amount of radiation the U.S. population is exposed to annually, while 11% was the contribution of all diagnostic medical imaging 20 years ago.¹¹ Increased exposure to radiation has raised concern toward its population health effects, namely carcinogenesis.^{12–16} Diagnostic radiation contains high-energy ionizing radiation, X-ray, which is sufficient to displace an electron from an atom or molecule, ionize, and destabilize that molecule. At a cellular level, ionizing radiation has the ability to alter DNA and create mutations, potentially inducing cancer.^{17,18}

One particularly radiosensitive tissue, the thyroid gland, $17,19$ is central to this discussion as it has become the fastest growing cancer in the United States.20 Although debate in the literature exists as to whether this is simply a function of improved detection methods, 21 the incidence of thyroid cancer has increased twofold since $1985²⁰$ and recent data suggest that increased diagnostic scrutiny is not the only reason.^{22,23} This does not directly implicate CT as a contributor to this dramatic increase in thyroid cancer, but an association should be considered given the susceptibility of the thyroid to radiation, particularly when factoring in its anatomic location with respect to cervical spine CT , 17,19,24

The pediatric population raises the most concern for harmful effects of diagnostic radiation from cervical spine CT because of the age-related sensitivity of the thyroid to radiation, placing youth at higher risk.^{18,25–28} Their intrinsic sensitivity to the effects of ionizing radiation plus the longer life expectancy during which risk may be expressed makes reducing exposure to children particularly important.²⁹

We hypothesize that CT examination of children to screen the cervical spine after trauma results in an important, measurable thyroid cancer risk greater than plain radiographs. The specific aims of this study are to compare the absorbed dose of radiation to the thyroid from plain radiographs with that from a cervical spine CT in pediatric trauma patients and to determine an estimated relative risk of induced thyroid cancer from the clearance process. This information will be useful to clinicians who must weigh the benefit of enhanced diagnostic imaging against the potential hazard of excess radiation exposure in a radiosensitive patient population.

PATIENTS AND METHODS

Study Groups

Institutional Review Board (M-2009-1358) exemption was obtained. All patients admitted during the 5-year window October 1, 2004, to October 31, 2009, were identified from a prospective, Institutional Review Board-registered pediatric trauma database who met the inclusion criteria: age 17 years, Level I trauma (emergent activation of the trauma team

secondary to airway compromise, cardiopulmonary duress, gunshot wound to the neck, chest, or abdomen, or Glascow coma scale score <8) or Level II trauma (emergent trauma evaluation but no physiologic abnormalities), 30 and had cervical spine imaging by plain radiographs or CT at our institution.

Calculation of the Absorbed Dose to the Thyroid for Plain Radiographs

The calculation of absorbed thyroid radiation from plain radiographs is dependent on field of view (we assumed 35×20), the tube current and exposure time (mAs), maximum voltage (kVp), distance from source to patient (75 cm), and tissue-specific absorption value $(mrad/R).$ ³¹ The tissue-specific absorption value was calculated from the equation:

Tissue specific absorption $(mRad/R)=5+(kVp-60)/4.31$

Absorbed thyroid radiation was calculated for each radiographic view, e.g., anteroposterior, lateral, and open-mouth odontoid, and summed as follows:

> Absorbed thyroid dose $(mGy) = (tissue - specific$ absorption value $[\text{mrad/R}]$ \times $(\text{mAs}) \times (\text{X-ray exposure})$ output $[mR/mAs]) \times (10^{-5}$ for unit conversion)

where 1 -mrad = 0.01 mGy.³¹

The actual mAs and kVp for each radiograph were not recorded; therefore, we estimated these based on recommended imaging protocols that contain ranges of settings. The largest value in the range was chosen to avoid underestimating the dose and falsely expanding the difference between CT and radiographs. To confirm the validity of our calculations, a 1 month prospective survey of cervical spine radiograph image settings was performed. Data were collected on three patients, all of whom had their actual image settings fall within the range of recommended imaging protocols used to calculate the dose.

Calculation of the Absorbed Dose to the Thyroid for CT

The total thyroid radiation dose from CT was calculated via Monte Carlo technique using the ImPACT CT Patient Dosimetry Calculator (version 1.0, August 28, 2009). The Monte Carlo technique $32,33$ uses mathematical equations based on anthropomorphic phantoms to model photon transport and estimate the absorbed radiation doses. The calculator applies these models with data collected by the United Kingdom's National Radiation Protection Board.32 This allows the amount of radiation absorbed by each organ to be estimated specific to the manufacturer and model of CT scanner. The variables required include CT manufacturer and model, kV, tube current (mA), rotation time (s), spiral pitch, field of view (set to the skull base through T2), and collimation (mm) and were obtained from each patient's medical record. The absorbed thyroid dose was calculated for each patient receiving a CT.

Our institution used three CT scanners during the course of the study—GE LightSpeed VCT, GE LightSpeed Pro 16, and GE LightSpeed 16 (General Electric, Waukesha, WI). The protocols are given in Table 1 and differ based on three age groups.

Estimating Cancer Risk

The National Academies' Biological Effects of Ioniz-ing Radiation 7th Report was used to estimate the thyroid cancer risk from radiation expressed as excess relative risk (ERR) according to:

> ERR=0.53exp $[-0.83 \text{ (age}-30)] \times \text{Gy}$, for males ERR=1.05exp $[-0.083 \text{ (age}-30)] \times \text{Gy}$, for females

The absolute risk is calculated by identifying the rate in the exposed population:

Absolute risk=ERR \times (thyroid cancer risk in the unexposed $[R_e])+R_e$

where $R_{\rm e}$ was 5.2/100,000 for males and 15.2/100,000 for females.²⁰

Statistical Methodology

The male and female groups were divided by age 0 to 6 years, 7 years to 11 years, and 12 years to 17 years. This allowed associations based on CT protocols (the 0- to 6-year-old group had different imaging protocols) and prestudy, arbitrary division of the older ages was done to allow comparison. Each group had radiation doses calculated and averaged using analysis of variance to look at differences between groups and pairwise two-tailed *t* test to calculate *p* values. The calculated age and radiation dose value for each group was used to estimate the gender-specific cancer risk. ERR data are not normally distributed, and so the nonparametric Kruskal-Wallis test was used to look at overall difference in ERR between age groups by sex. A Wilcoxon rank-sum test examined the pairwise differences between age groups. Bonferroni correction was used and $p < 0.05$ was considered statistically significant.

RESULTS

Demographics

Six hundred seventeen patients met the inclusion criteria: 224 had plain cervical radiographs and 393 had cervical CT (Table 2).

Plain Radiograph Radiation

The mean radiation absorbed to the thyroid from plain radiographs in males was 0.90 mGy and 0.96 mGy for females, which was not significantly different. Similarly, when analyzed by age there was no difference between males and females. However, increased age was associated with significantly greater thyroid radiation absorption for both genders in the oldest age group (12–17 years) compared with the two younger groups (Table 2).

CT Radiation

The mean radiation dose from CT absorbed by the thyroid in males was 63.6 mGy and 64.2 mGy for females, this difference was not statistically significant. Age significantly impacted the dose as younger patients received, on average, less radiation to the thyroid: young males (0–6 years) = 51.7 mGy, young females (0–6 years) = 55.5 mGy; males (7–11 years) = 51.3 mGy, females $(7-11 \text{ years}) = 56.2 \text{ mGy}$; and adolescent males $(12-17 \text{ years}) = 70.2 \text{ mGy}$, adolescent females $(12-17 \text{ years}) = 68.3 \text{ mGy}$. The radiation dose received by the younger two age groups was significantly lower than the adolescent age groups but not significantly different between the youngest two age groups, males and females alike (Table 2).

Excess Relative Risk

The median lifetime ERR for a plain radiograph cervical spine series in males was 0.24% and females was 0.51% (Table 3; Fig. 1, A and B).

The median ERR for one cervical spine CT in males was 13.0% (range, 10–66%) and females was 25.0% (range, $8.0-116.0\%$), $p < 0.001$ when compared with plain radiographs (Table 3). Young (0–6 years) males and females have the highest ERR for one CT scan, 21.5% and 45.6%, respectively, compared with the older two groups. Males 7 to 11 years had a 13.5% ERR and females 7 to 11 years a 29.7% ERR, whereas the adolescent male group (12–17 years) had a 12.0% ERR and female adolescent group (12–17 years) a 22.7% ERR. Furthermore, younger age resulted in significantly higher ERR when compared against the older age groups (Fig. 2, A and B).

Absolute Risk

The prevalence of thyroid cancer in the general population is 5.2/100,000 males and 15.2/100,000 females.20 The risk after plain radiographs increased from 5.20 to 5.21/ $100,000$ (0.0052% \rightarrow 0.00521%) for males and from 15.2 to 15.3/100,000 (0.0152%) \rightarrow 0.0153%) for females. For CT, the estimated absolute risk increased from 5.2 to 5.87/100,000 (0.0052% \rightarrow 0.00587%) for males and from 15.2 to 19.0/ 100,000 (0.0152%) \rightarrow 0.019%) for females (Table 4).

Comparison Between Plain Radiograph and CT ERR

The overall median ERR for CT compared with plain radiographs was significantly higher, 0.24% versus 13.0% for males and 0.51% versus 25.0% for females, for inducing thyroid cancer $(p < 0.001)$. This was true across both genders and all age groups (Table 3).

DISCUSSION

A 5-year retrospective review of our institution's pediatric trauma database was performed to determine the relative difference in radiation dose to the thyroid between CT and plain radiographs of the cervical spine. Our protocol was to obtain CT in patients who had symptoms, were at higher risk for injury, or who were otherwise being evaluated with CT. Clearance of the cervical spine using CT at our institution resulted in a 60-fold increase in radiation absorbed to the thyroid when compared with plain radiographs, 64 mGy to 0.9

mGy, $p < 0.001$. This amount of radiation to the thyroid from a CT cervical spine is approximately equivalent to the absorbed dose to the lungs from 80 chest radiographs.³⁴

The true clinical value, however, lies in gauging the potential for the CT radiation dose to be carcinogenic. On the basis of the linear nonthreshold theory (LNT), the National Research Council described a formula for calculating thyroid cancer risk.18 A single CT cervical spine in our pediatric trauma population increases the median lifetime risk 13.0% in males and 25.0% in females compared with 0.24% in males and 0.51% in females for plain radiographs. Further subdividing the patients by age demonstrates the significant influence youth has on increasing risk despite overall reductions in absorbed radiation from CT. The youngest patient group (0–6 years) received the lowest amount of radiation to the thyroid, yet had the highest median ERR. This supports the age-related sensitivity of the thyroid to radiation and provides evidence for heightened scrutiny when ordering cervical CT in the younger patient populations.

The observation that significantly lower doses were absorbed by younger patients is intuitive but also reflected differences in protocols between age groups at our institution. To decrease total radiation, CT protocols were adjusted in children younger than 6 years. Compared with older children, this demonstrated that imaging protocols that consider radiation dose can have a significant impact on lowering the radiation doses. The quality and diagnostic sensitivity of these studies was not assessed in this study, but clinically there was no issue with inferior image quality in the population that received less radiation. Further efforts to diminish radiation dose during CT are needed.

The impact of age on thyroid cancer induction is secondary to its innate sensitivity to radiation and the time-dependency of thyroid cancer expression.^{17,19,24} Therefore, the youngest patients have the most radiosensitive thyroids and the greatest time with which to express induction of thyroid cancer. The radiation dose calculator, ImPACT CT Patient Dosimetry Calculator (version 1.0, August 28, 2009), used in this study is based on adult phantoms and potentially underestimates the dose absorbed to pediatric patients.35 After performing another dosimetry study involving pediatric phantoms, Khursheed et al.³⁵ concluded that effective doses for pediatric patients calculated on this dosimetry calculator ranged from 1.1 (15 years) to 2.2 (1 year) times higher than that of an adult. To avoid inaccurately inflating the calculated absorbed doses, this age-dependent factor was omitted from our calculations with the acknowledgment that the doses arrived at are potentially underestimated.

Other limitations of the study potentially exist in both the calculation of absorbed doses to the thyroid and the determination of the risk for developing cancer. The absorbed doses from plain radiographs were estimates based on imaging protocols and not actual dose data for each patient. Rybicki et al.10 has published absorbed doses to the thyroid in adults from plain radiographs—lateral radiograph 0.09 mGy, anteroposterior 0.46 mGy, and open-mouth odontoid 1.04 for a total of 1.59 mGy. The mean radiation in the oldest patient group (that most closely resembling adults) from our data set was 1.55 mGy. Furthermore, the difference between plain radiographs and CT doses was significantly large such that any potential miscalculation would be too minimal to affect the significance. Arbitrary grouping

of patients could misrepresent the true effect of age on thyroid dose and cancer induction. In addition, different-sized children within each age group creates a variability in the radiation dose calculated that may cloud comparison by age alone and necessitated data analysis using median values instead of means. Finally, three different CT scanners were used to perform the cervical spine imaging possibly introducing an uncontrolled variable.

Determining the risk for developing thyroid cancer from cervical spine CT using the linear nonthreshold theory is not uniformly accepted. Simply, this theory states that no threshold exists for radiation's consequence and the effect is dose-dependent in a linear fashion and additive across time. Opponents of LNT refer to studies illustrating a potential protective effect of low dose radiation, so-called radiation hermesis.^{36–40} The science to support or negate the effect of low-dose (<100 mGy) radiation on cancer induction is weak at best and conclusions made based off the linear LNT should be regarded with caution.⁴¹ The application of LNT in this study is solely to illustrate the difference between plain radiography and CT radiation doses. It would be inaccurate to conclude that CT cervical spine causes thyroid cancer—rather, in a highly susceptible organ and sensitive patient population, the significant difference between plain radiographs and CT for potentially inducing cancer should be known so the physician can make the most educated decision to diagnose the pediatric trauma patient.

There are no national guidelines to direct clearance of the cervical spine in pediatric trauma patients and current algorithms lack support from evidence-based research.42 The Eastern Association for the Surgery of Trauma, American College of Radiology (ACR), and the American Academy of Orthopedic Surgery (AAOS) have all authored imaging guidelines for the clearance of the cervical spine, with the AAOS and ACR specifically addressing the pediatric trauma patient.43–45 The National Emergency X-Radiography Utilization Study (NEXUS) has provided practitioners with information to guide the decision to obtain imaging based on clinical findings—no midline cervical tenderness, no focal neurologic deficits, no intoxication or indication of brain injury, no painful distracting injuries, and normal alertness.^{1,46} However, once the decision to image a patient has been determined, the proper imaging study to use remains controversial. In the journal for the AAOS, Eubanks et al.⁴³ state that plain radiographs are the proper imaging study in the conscious patient at risk (those that meet NEXUS criteria) with imaging by CT of the upper cervical spine if proper C1–2 visualization cannot be achieved or if head CT is indicated. The ACR agrees with applying the NEXUS criteria to older children but does not validate the criteria for younger children (<9 years). Radiographs were deemed appropriate by the ACR committee when NEXUS criteria were met while CT was not an appropriate imaging option.43 Eubanks et al.43 approach the unconscious patient differently as cervical spine CT is obtained with the head CT because of the altered mental status. Then, if unconsciousness is expected to last longer than 24 hours, a magnetic resonance imaging of the cervical spine is performed to definitively clear the cervical spine.⁴³ A recent review of the literature by Hutchings and Willett⁴² agreed with the approach outlined by Eubanks et al; however, no firm statement was able to be made for the imaging choice in the unconscious/intubated patient.

The significant difference in radiation that CT delivers to the pediatric trauma patient when compared with plain radiographs should temper excessive use. Thus, CT is not

recommended as the initial study in conscious patients that meet NEXUS criteria for imaging. In the unconscious patient already obtaining a CT head, we would urge practitioners to consider using plain radiographs as the initial imaging modality followed by magnetic resonance imaging if advanced imaging proved necessary—either from inadequate films or an anticipated prolonged unconscious state. The concern for missed injury is real, but the possible impact of the radiation dose on thyroid cancer induction should factor into the diagnostic algorithm. The significant difference in thyroid dose between CT and plain radiographs, along with the theoretical increase in thyroid cancer estimated relative risk, supports a more conservative cervical spine clearance protocol in the pediatric trauma patient. These results will hopefully facilitate a sensible approach to the cervical spine clearance process and more accurately allow clinicians to measure the risk of advanced imaging in the pediatric trauma patient.

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(*A*, *B*) Excess relative risk box plot for males and females receiving cervical spine radiographs. *White boxes* indicate 1 SD from the median value (single line within the box), while the bars indicate the range of values.

(*A*, *B*) Excess relative risk box plot for males and females receiving cervical spine computed tomography. *White boxes* indicate 1 SD from the median value (single line within the box), while the bars indicate the range of values.

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TABLE 2

Absorbed Radiation to the Thyroid in mGy for Males and Females From Plain Radiographs and CT Cervical Spine Absorbed Radiation to the Thyroid in mGy for Males and Females From Plain Radiographs and CT Cervical Spine

Statistically significant difference in thyroid absorbed radiation between the 12- and 17-yr age group and the younger two age groups.

Excess Relative Risk of Thyroid Cancer Induction for CT Cervical Spine Versus Plain Radiographs Excess Relative Risk of Thyroid Cancer Induction for CT Cervical Spine Versus Plain Radiographs

Median values reported with the range in parentheses; *p* value indicates significance for age- and gender-matched groups comparing CT and plain radiographs.

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Estimated Absolute Risk of Thyroid Cancer Induction From Plain Radiographs and CT Cervical Spine Estimated Absolute Risk of Thyroid Cancer Induction From Plain Radiographs and CT Cervical Spine

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