Pediatric computed tomography practice in Japanese university hospitals from 2008–2010: did it differ from German practice?

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ABSTRACT

Computed tomography (CT) is an essential tool in modern medicine and is frequently used to diagnose a wide range of conditions, particularly in industrial countries, such as Japan and Germany. However, markedly higher doses of ionizing radiation are delivered during CT imaging than during conventional X-ray examinations. To assess pediatric CT practice patterns, data from three university hospital databases (two in Japan and one in Germany) were analyzed. Anonymized data for patients aged 0 to 14 years who had undergone CT examinations between 2008 and 2010 were extracted. To assess CT practice, an interdisciplinary classification scheme for CT indications, which incorporated the most common examination types and radiosensitive tissues, was developed. The frequency of CT examinations was determined according to sex, age at examination, and indications. A total of 5182 CT examinations were performed in 2955 children. Overall, the frequency of CT examinations at the Japanese university hospitals did not differ significantly from that at the German hospital. However, differences were detected in the age distribution of the patients who underwent CT examinations (the proportion of patients <5 years of age was significantly higher in Japan than in Germany) and in the indications for CT. Substantial practice differences regarding the use of CT in pediatric health care were detected between the three hospitals. The results of this study point towards a need for approaches such as clinical guidelines to...
INTRODUCTION
Computed tomography (CT) is an essential tool in modern diagnostic radiology. Compared with conventional X-ray, ultrasound, and magnetic resonance imaging (MRI) examinations, CT scans deliver 3D images within seconds and are not affected by the physician’s skills level [1]. These attributes can be life-saving, especially in emergency situations, when rapid decision-making is essential. Consequently, the use of CT is steadily increasing in all industrialized countries [2]. However, considerably higher doses of ionizing radiation are involved than those used in conventional X-ray examinations. Ionizing radiation is a well-established risk factor for cancer [3]; the risk increases in a dose-dependent manner and is higher in younger patients [4]. Furthermore, children are assumed to be more radiosensitive than adults and at higher risk of developing cancer due to their long lifespan after the radiation exposure [5].

In pediatric patients, the doses delivered to exposed organs from repeated examinations can add up to 100–200 mGy [6–8], and this level is associated with an elevated risk of cancer [9]. In 2001, Brenner et al. estimated the CT-attributable cancer mortality rate of 600 000 children who were exposed to CT in the US in 2000. According to their findings, 500 children were expected to die from cancer attributable to CT examinations, i.e. they detected an attributable lifetime cancer mortality risk of 0.25% [6]. Several modeling studies have subsequently estimated the attributable cancer mortality and cancer incidence rates for various groups of pediatric and adolescent patients who had undergone CT [10]. Recent large epidemiological cohort studies that attempted to empirically confirm these results suggest that CT examinations might cause childhood cancers, such as leukemia and solid cancers [11–15]. However, this association has not yet been definitively confirmed. One major issue with these studies relates to the non-exclusion of CT examinations that have been performed because of a suspected malignancy, which might have introduced confounding bias. As a consequence, the indications that prompt the use of CT should be assessed when estimating cancer risk after CT exposure [16].

Recent research has also indicated that knowledge gaps regarding CT doses and the associated health risks exist among referring physicians [17, 18]. Furthermore, little is known about the indications under which CT examinations are most commonly performed and how such parameters vary among countries. This lack of information hampers efforts to assess CT practices and quantify unwanted exposure.

Japan and Germany both exhibit higher X-ray examination rates than other industrialized countries (the rates seen in the two countries are similar). However, the X-ray examination attributable cancer risk is estimated to be higher in Japan (the highest globally) than in Germany [19]. In addition, unlike other countries such as Germany, the USA and the UK, Japan does not have commonly agreed clinical guidelines for the use of ionizing radiation in diagnostic radiology, especially in the pediatric field [20].

We assessed pediatric CT practice patterns, including the associated indications, in Japan. In addition, German data were used to identify potential differences compared with Japanese CT practices, which could aid the development of best-practice guidelines for CT use in Japan.

MATERIALS AND METHODS
Study population and data collection
We extracted anonymized examination data from the Radiological Information Systems (RISs) at three university hospitals: two in Japan (Nagasaki University Hospital and Fukushima Medical University Hospital) and one in Germany (Mainz University Medical Center). All pediatric patients who had undergone at least one CT examination during the study period (1 July 2008 to 30 June 2010) and were between 0 and 14 years of age at the time of the first CT examination were included in this study. Only residents of the respective countries were eligible. Data regarding sex, date of birth, the date of the examination, and the indications under which the examinations were conducted were extracted for all eligible patients.

Data management and analysis
We calculated the mean number of CT examinations for the whole study population and according to sex, age at the time of the examination (hereafter referred to as ‘age at examination’), indication, and the hospital at which the scan was carried out. The Chi-squared test for independent samples was used to assess the differences in the mean of the examination number, number of each sex, or age at examination between the two countries. An interdisciplinary classification scheme combining the most common examination types and radiosensitive tissues was jointly developed by a small team led by senior physicians from each country. The classification scheme was piloted on a randomly drawn subset of the original data. The categories included tumor-related examinations (‘tumor diagnoses and therapy’ or ‘tumor exclusion’), trauma, congenital/other typical childhood diseases, and general examination regions (‘cranium’, ‘spine’, ‘thorax’, and ‘abdomen’), as well as ‘other’ examinations (Fig. 1). We separated out the cases involving tumor- or tumor-related or trauma indications because CT examinations are usually essential in such cases. As our aim was to develop best practice guidelines based on an international comparison, it was considered that separating cases in which CT examinations were essential from other cases might provide some insight into the number of ‘non-essential’ (i.e. potentially unnecessary) examinations. Statistical analysis was performed using SPSS Statistics 22.0 (IBM Japan, Tokyo, Japan).

Ethics statement
This study was approved by the Ethics Committees of Nagasaki University Hospital, Fukushima Medical University and the

KEYWORDS: computed tomography, ionizing radiation, children, practice pattern, international comparison

reduce unwarranted medical radiation exposures, particularly abdominal and head CT, in the Japanese health system.
Medical Chamber of Mainz, and from the Data Protection Officer of Mainz University Medical Center before the start of the study. The study was conducted in accordance with the guidelines outlined in the Declaration of Helsinki. All of the data used in this study were analyzed anonymously and securely protected under the guidance of the Data Protection Officer of each hospital.

RESULTS
The three university hospitals differ considerably in size. Compared with the two Japanese hospitals, Mainz University Medical Center has twice as many beds and 2-fold and 5-fold higher numbers of physicians and annual inpatients, respectively. However, the number of out-patients was highest at Fukushima Medical University Hospital, while similar numbers of outpatients visited the other two

<table>
<thead>
<tr>
<th>Hospital</th>
<th>Nagasaki University Hospital</th>
<th>Fukushima University Hospital</th>
<th>Mainz University Medical Center</th>
</tr>
</thead>
<tbody>
<tr>
<td>Number of Physicians</td>
<td>436</td>
<td>488</td>
<td>960</td>
</tr>
<tr>
<td>Annual in-hospital patients</td>
<td>13 586</td>
<td>12 995</td>
<td>68 661</td>
</tr>
<tr>
<td>Annual out-hospital patients</td>
<td>280 162</td>
<td>373 393</td>
<td>241 429</td>
</tr>
<tr>
<td>Number of beds</td>
<td>861</td>
<td>778</td>
<td>1640</td>
</tr>
</tbody>
</table>

Examining and patients

<table>
<thead>
<tr>
<th>Category</th>
<th>Nagasaki University Hospital</th>
<th>Fukushima University Hospital</th>
<th>Mainz University Medical Center</th>
</tr>
</thead>
<tbody>
<tr>
<td>All CT examinations</td>
<td>40 746</td>
<td>40 609</td>
<td>46 489</td>
</tr>
<tr>
<td>Pediatric CT examinations (% from all)</td>
<td>1962 (4.8%)</td>
<td>2220 (5.5%)</td>
<td>1000 (2.2%)</td>
</tr>
<tr>
<td>Pediatric patients examined with CT</td>
<td>1203</td>
<td>1138</td>
<td>614</td>
</tr>
<tr>
<td>Mean age at CT examination (SD)</td>
<td>5.7 ± 4.7</td>
<td>5.0 ± 4.8</td>
<td>7.7 ± 4.2</td>
</tr>
</tbody>
</table>

Fig. 1. Classification scheme for CT indications.

Table 1. Hospital and study population characteristics for Nagasaki University Hospital, Fukushima Medical University Hospital, and Mainz University Medical Center (2008–2010)
hospitals (Table 1). At Mainz, CT procedures were managed by specialized pediatric radiologists in the Radiology Department, whereas at Nagasaki and Fukushima CT examinations were conducted by general radiologists who did not specialize in pediatric radiology.

In total, 5182 CT examinations were performed on 2955 patients between 1 July 2008 and 30 June 2010 at the three hospitals (Nagasaki: 1962 CT examinations, 1203 children; Fukushima: 2220 CT examinations, 1138 children; Mainz: 1000 CT examinations, 614 children) (Table 1). The sex distribution of the patients that underwent CT examinations was similar at each hospital, with males undergoing CT scans slightly more often than females. The mean age at examination for the Japanese hospitals (5.5 years) was significantly lower than that for the German hospital (7.7 years) (P < 0.001; Table 2).

At the Japanese hospitals, almost 20% of all pediatric CT examinations involved children younger than 12 months of age. In contrast, at the Mainz University Medical Center significantly fewer CT examinations were performed on patients below 12 months (2.2%). The proportion of pediatric CT scans that involved patients of ≤5 years old was significantly higher at the Japanese hospitals than at the German hospital (49.1% vs 28.8%, P < 0.001; Table 2). On the other hand, the proportions of patients aged 5–9 years and 10–14 years who underwent CT examinations were significantly lower at the Japanese hospitals than at the German hospital (P < 0.001, respectively; Table 2).

Of the 5182 extracted CT examinations, 5115 were classified according to the classification scheme developed for this study (Fig. 1). No unambiguous indications could be extracted from the RIS at Mainz University Medical Center for the remaining 67 examinations, and so these cases had to be excluded from the analysis. Among the 4182 examinations performed at the Japanese hospitals, cranial indications were the most common indications for CT (42% of all CT scans), followed by tumor-related (18%), trauma-related (16%), and abdominal (11%) indications (Table 3). In comparison, trauma-related was the most common indication for CT at the German hospital (n = 933) (34%), followed by cranial and tumor-related indications, which accounted for 27% and 23% of all CT examinations, respectively. The proportions of patients who underwent CT examinations of the cranium, thorax and abdomen were significantly higher at the Japanese hospitals than at the German hospital (P < 0.001, P < 0.01, P < 0.001, respectively; Table 3). On the other hand, the proportion of patients who underwent CT examinations under tumor-related, trauma-related, or spine indications were significantly higher at the German hospital than at the Japanese hospitals (P < 0.01, P < 0.001, P < 0.05, respectively; Table 3). Overall, the three most common indications (cranial, trauma-related, and tumor-related) accounted for ~80% of all examinations in both countries. Additional analyses showed that at the Japanese hospitals the majority of CT examinations performed on patients below 12 months of age were cranial scans (61%, n = 553), while at Mainz, the majority of these examinations were tumor-related (45%, n = 10).

The mean number of CT examinations per patient was 1.79 (standard deviation (SD): 2.03, range: 1–27) at the Japanese hospitals, and 1.63 (SD: 1.39, range: 1–12) at the German hospital. The frequency distributions for the number of scans performed in each individual did not differ significantly between the two countries (1 scan: P = 0.14, 2–6 scans: P = 0.38, ≥7 scans: P = 0.10, respectively; Table 4). Approximately one-third of the patients at each university hospital underwent more than one CT examination (30% at the Japanese hospitals, 26% at the German hospital). Of these, the majority had two to three CT scans (Fig. 2).

**DISCUSSION AND CONCLUSION**

To the best of our knowledge, this is the first study to assess the CT practice pattern in Japan and to compare detailed information regarding the number of CT examinations performed and the corresponding indications with those from another industrialized country that exhibits similar levels of X-ray use. In the present study, the overall frequency of pediatric CT examinations did not differ significantly between the examined Japanese and German university hospitals. However, substantial differences, in the age distribution of the examined patients and the most common CT indications, particular for abdominal and cranial CT scans, were seen between the Japanese and German hospitals.

The differences observed in the present study might reflect variations in the availability and use of CT scanners between the two countries, as Japan has a 5.7-fold higher number of CT scanners per
more radiosensitive than adults and have longer to live and so are young children examined in Japan, as children are considered to be particularly relevant when considering the large proportion of very population due to less frequent usage of CT in Germany. A previous study that indicated that both Japan and Germany CT-attributable cancer risk than the Germany population, probably same study found that the Japanese population has a 2-fold higher frequency of pediatric CT examinations by indication.

Table 3. Proportion of pediatric CT examinations by indication

<table>
<thead>
<tr>
<th>Indication category</th>
<th>Japan (two hospitals, n = 4182)</th>
<th>Germany (n = 933)</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Tumor</td>
<td>749 (17.9%)</td>
<td>216 (23.2%)</td>
<td>&lt;0.01**</td>
</tr>
<tr>
<td>Tumor exclusion</td>
<td>45 (1.1%)</td>
<td>26 (2.8%)</td>
<td>&lt;0.01**</td>
</tr>
<tr>
<td>Trauma</td>
<td>674 (16.1%)</td>
<td>315 (33.8%)</td>
<td>&lt;0.001***</td>
</tr>
<tr>
<td>Cranium</td>
<td>1754 (41.9%)</td>
<td>255 (27.3%)</td>
<td>&lt;0.001***</td>
</tr>
<tr>
<td>Spine</td>
<td>102 (2.4%)</td>
<td>38 (4.1%)</td>
<td>&lt;0.05*</td>
</tr>
<tr>
<td>Thorax</td>
<td>366 (8.8%)</td>
<td>59 (6.3%)</td>
<td>&lt;0.01**</td>
</tr>
<tr>
<td>Abdomen</td>
<td>444 (10.6%)</td>
<td>16 (1.7%)</td>
<td>&lt;0.001***</td>
</tr>
<tr>
<td>Childhood disease</td>
<td>36 (0.9%)</td>
<td>6 (0.6%)</td>
<td>0.56</td>
</tr>
<tr>
<td>Others</td>
<td>12 (0.3%)</td>
<td>2 (0.2%)</td>
<td>0.99</td>
</tr>
</tbody>
</table>

*P < 0.05, **P < 0.01, ***P < 0.001, *67 CTs could not be classified.

Table 4. Proportion of pediatric CT examinations by frequency

<table>
<thead>
<tr>
<th>Frequency</th>
<th>Japan (two hospitals)</th>
<th>Germany (n = 933)</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Pediatric patients</td>
<td>2341 (70.6%)</td>
<td>600 (73.7%)</td>
<td>0.14</td>
</tr>
<tr>
<td>1 scan</td>
<td>1652 (50.1%)</td>
<td>442 (54.0%)</td>
<td>0.94</td>
</tr>
<tr>
<td>2–6 scans</td>
<td>616 (26.3%)</td>
<td>147 (25.4%)</td>
<td>0.38</td>
</tr>
<tr>
<td>≥7 scans</td>
<td>73 (3.1%)</td>
<td>11 (1.8%)</td>
<td>0.1</td>
</tr>
</tbody>
</table>

The data for 14 people from Germany had to be excluded from the analysis.

In the Japanese hospitals, the proportion of CT examinations involving patients aged <12 months was not only considerably higher than the equivalent values for all other age groups at the Japanese hospitals, but was also higher than those for all age groups at the Mainz University Medical Center. In contrast, the lower frequency of CT examinations involving very young patients recorded at the Mainz University Medical Center might indicate that the clinicians at that institution are acutely aware of the pediatric radiation risk associated with CT examinations. However, recent studies have reported mixed results regarding physicians’ knowledge of the CT-related risks of pediatric and adult patients [17, 18].

In the current study, we found large differences in the frequencies of various indications for CT between hospitals in Japan and Germany. In Germany, the Radiation Protection Commission together with clinical experts has developed guidelines for diagnostic radiological procedures [20]. Consequently, the low overall number of CT examinations per-patient is consistent with the results of these guidelines.

In Japan, at present there are no standardized clinical guidelines for pediatric CT. In 2013, guidelines for medical imaging were published in Japan [23], but merely 6 out of the 476 pages were devoted to pediatric imaging. In addition, it appears that these guidelines have not yet found wide acceptance among Japanese referring physicians. The large variations in CT examination patterns between the examined hospitals in Japan and Germany might be associated with the lack of commonly accepted diagnostic guidelines in Japan, but this has not been empirically verified. It has been reported that the use of CT examinations varies significantly between individual doctors, even under similar conditions [24, 25].

Thus, standardized clinical guidelines might be useful for promoting the consistent use of CT examinations. Furthermore, in Japanese university hospitals CT examinations are conducted by general radiologists rather than by specialized pediatric radiologists (in Germany, specialized pediatric radiologists generally perform pediatric CT, when available). This might partly explain the differences between the two countries.

We assumed that differences in common medical practice played the most important role in the observed differences in CT practice, since the Japanese and German health systems (including the reimbursement schemes for radiological examinations employed as part of the health insurance systems in operation in each country) are quite similar [26]. Therefore, the standardization and appropriate use of CT practice is essential. Some professional societies, such as the American College of Radiology (ACR) and the US Society for Pediatric Radiology, have produced recommendations and guidelines for pediatric CT use [27, 28]. Moreover, the ACR recently developed a CT dose index registry in the USA as a quality control and safety instrument, and the European Society of Radiology recently launched the ‘Eurosafe Imaging’ campaign, which aims to educate clinicians about the appropriate usage of medical radiation [29]. Unlike in the USA and the EU, no nationwide framework or campaign about the appropriate use of medical radiation in pediatrics exists in Japan.

In addition to clinical guidelines, clinical ordering aids and the exchange of CT images via electronic imaging communication.
platforms (e.g. the 'Image Share project'; [www.rsna.org/Image_Share.aspx](http://www.rsna.org/Image_Share.aspx)) can also help to reduce unnecessary diagnostic imaging. The use of non-ionizing imaging techniques, such as ultrasound and MRI, instead of ionizing imaging is another approach that can be used to further reduce ionizing radiation exposure. In cases of suspected appendicitis, the ACR recommends that non-contrast-enhanced ultrasound or MRI examinations should be performed instead of abdominal CT in pediatric patients [26]. Moreover, innovations in medical technology have resulted in the development of pre- and post-processing methods that make it possible to reduce the radiation dose delivered during CT without a deterioration of image quality [30–32]. In addition to such measures, education regarding radiation and the associated risks might contribute to optimizing the use of radiological examinations.

Our study had several limitations. We could not identify the reason why the frequencies of CT examinations involving very young patients and cranial/abdominal CT are so high in Japan. The radiological report data for the Japanese hospitals indicated that many of the CT examinations were performed for hydrocephalus or suspected appendicitis. However, these observations were incomplete, and no radiological report data were available for the German hospital. In addition, the study data related to the years 2008–2010 and, hence, were not up to date. However, we assume that any major changes in pediatric CT practice would have been influenced by the initial report about CT risk prediction by Brenner et al., which was published in 2001 [6], and thus, would have occurred before our study period. In addition, a 2004 study obtained similar findings regarding the number of CT examinations and the age distributions of patients who had undergone CT scans to those we obtained for the Nagasaki University Hospital [22], confirming our observations. Furthermore, the current study only analyzed data from three university hospitals. Thus, the results are unlikely to accurately reflect the general practice patterns in the respective countries and hospitals. We consider that the basic characteristics of the three examined hospitals were similar because they were all general university hospitals. However, it is possible that differences between the characteristics of the examined hospitals affected the results of the present study. Finally, we consider that dose estimation is also important for risk assessments of pediatric CT examinations; however, the data available from the RIS did not allow us to calculate organ doses. Another limitation of this study was that we only had data about CT examinations and were not able to acquire data about the other diagnostic imaging techniques employed at the target hospitals. The increased use of MRI might explain the differences in the use of CT for tumor diagnostics between the hospitals. However, analyzing all of the indications for diagnostic imaging at the target hospitals was beyond the scope of this study.

In conclusion, variations in CT examination practices were detected between hospitals in Japan and Germany, with large differences being observed in the age distribution of the patients who underwent CT and in the most common indications for CT examinations among pediatric patients. To reduce unwarranted medical radiation exposure in pediatric patients, simple and useful clinical guidelines that have been designed for use by referring physicians should be developed. Risk communication and educational materials for parents that aim to reduce the number of unnecessary CT examinations performed should also be produced. In addition, the active implementation of technical dose-reduction approaches should be promoted in cases in which CT cannot be replaced by another diagnostic modality.

**CONFLICT OF INTEREST**
There are no conflicts of interest for any of the authors of this manuscript.

**ETHICS STATEMENT**
Approvals for the study were obtained before the beginning of the study from the Ethics Committee of the Nagasaki University.
REFERENCES


