Differences in upper gastrointestinal neoplasm detection rates based on inspection time and esophagogastroduodenoscopy training

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ABSTRACT

Background and study aims Esophagogastroduodenoscopy (EGD) has utility in early detection of upper gastrointestinal (UGI) neoplasms. However, previous studies report shorter inspection times and inexperienced endoscopists contribute to overlooking gastric neoplasms. We investigated neoplasm detection rates according to inspection time and extent of EGD training.

Patients and methods In this retrospective observational study, we reviewed routine EGDs for 3,925 consecutive cases between October 2014 and March 2015. We divided the endoscopists into three groups based on median inspection time during EGD without undergoing biopsy. Using cut-off median inspection times of 7 and 10 minutes, three, five, and eight endoscopists were classified into the fast, moderate, and slow groups, respectively. We compared detection rates according to inspection time and the extent of EGD training.

Results The median inspection time among all endoscopists was 9.3 minutes (range, 6.6–12.0 min). The detection rate for UGI neoplasms was as follows: fast group, 3.6%; moderate group, 3.3%; and slow group, 3.1% (P=0.807). The median inspection time was significantly shorter among the intensive training ≥1-year group than among the < 1-year group (< 1-year: median 6.3 min; range 8.2–13.9 min, ≥ 1-year: median 8.9 min; range 6.4–11.4 min, P<0.001). The detection rate for UGI neoplasms was significantly higher among the intensive training ≥1-year group than among the < 1-year group (< 1-year: 2.2%; ≥ 1-year: 3.7%, OR=1.65, 95% CI: 1.02–2.68, P=0.041).

Conclusions There was no association between inspection times and neoplasm detection rates. The quality of EGD, as measured by neoplasm detection rates, may be improved by ≥ 1-year of intensive training.

Introduction
Gastric cancer is the fifth most common malignancy and the third leading cause of death from cancer worldwide [1,2]. Early detection is ideal for optimum patient survival, with esophagogastroduodenoscopy (EGD) being the most sensitive method for early detection [2,3]. Several studies have demonstrated the effectiveness of endoscopy screening for gastric cancer, with a 30% reduction in gastric cancer mortality as a result of cancer detection attributed to endoscopy screening [4–7]. Since then, in Korea and Japan, use of EGD has been allowed for gastric cancer screening in the national screening program [8,9]. Furthermore, EGD is also suggested to be effective for early detection of additional upper gastrointestinal (UGI) neoplasms, such as hypopharynx, esophageal, and duodenal tumors [10,11]. Although EGD is the standard procedure for diagnosing gastric cancer, inexperienced endoscopists tend to overlook gastric cancer due to subtle morphological changes in some early gastric cancer lesions, which are difficult to distinguish from background mucosa with atrophic change [12–14].
As a result, the false-negative rate for detecting gastric cancer with EGD is 4.6% to 25.8% [12, 15 – 19]. These findings indicate the importance of quality indicators (QI) for UGI neoplasm detection in EGD.

Several studies have reported quality indicator outcome measures in EGD [20 – 22]. A few previous studies found an association between inspection time during EGD and detection rates for UGI neoplasms. Results of those studies have shown that slower endoscopists detected a higher proportion of UGI neoplasms than faster endoscopists, suggesting that inspection time may be a useful quality indicator in EGD [23 – 25]. Conversely, other previous studies found that endoscopists require sufficient training and experience to detect gastric cancer properly [12 – 14]. We thus investigated inspection time and endoscopic training and experience to clarify quality improvement for UGI neoplasm detection in EGD.

Patients and methods

Study design

In this single-center, retrospective observational study, we reviewed routine EGDs for 5,091 consecutive cases at the Cancer Institute Hospital between October 2014 and March 2015. We selected this period because there was no change in the staff of endoscopists. We excluded 464 cases with EGD history within 6 months, 369 cases with endoscopic assessment by a clinician with less than 3 years of experience, 129 cases with endoscopic assessment by a clinician who performed fewer than 50 EGDs in this period, 67 cases of total gastrectomy surgery, 55 cases lacking information on stomach examination, 46 cases with residual food in the stomach, and 36 cases with ultrathin endoscopy examinations. Before undergoing EGD, all patients provided comprehensive written informed consent. The institutional review board of the Cancer Institute Hospital of the Japanese Foundation for Cancer Research approved this study (IRB no. 2016-1158) (Fig.1).

Routine EGD

Routine EGDs were performed from the hypopharynx to the horizontal part of the duodenum. Approximately 60 images were captured, including about 20 images taken in the hypopharynx and esophagus, and about 40 taken in the stomach and duodenum. Digital images were automatically saved in the endoscopic filing system. Two imaging modalities were alternated with conventional white-light imaging (C-WLI) and narrow-band imaging (NBI) when using the endoscope. NBI was used during withdrawal from the esophagus. The video processor was consistently set as follows. The structure enhancement function was set at the A5 or B8 level for C-WLI, with the color mode fixed at level 0. All procedures were carried out using an endoscope manufactured by Olympus Corporation (GIF-H260 or GIF-H290Z, Olympus Medical Systems, Tokyo, Japan) and a standard endoscopic video system (EVIS LUCERA ELITE or EVIS LUCERA SPECTRUM, Olympus Medical Systems).

As a preliminary measure prior to endoscopy screening, pharyngeal anesthesia was administered using a lidocaine viscous solution. Almost all patients underwent endoscopy under intravenous (IV) sedation, except for those with no preference for anesthesia. In accordance with the guidelines regarding sedation for gastroenterological endoscopy [26], midazolam (2.0 – 5.0 mg) was administered IV, occasionally with pethidine hydrochloride (35 mg) added for patients who were insufficiently sedated. When gastric or duodenal neoplasms were detected, a biopsy was performed after observation following NBI and dye spraying endoscopy with the application of indigo carmine. When esophageal neoplasms were detected, a biopsy was performed after observation following NBI and iodine spraying.

Procedure and outcome parameters

Inspection time was defined as the time from first image capture in the pharynx to scope removal. We divided endoscopists into three groups based on median inspection time during EGD without undergoing biopsy. The cut-off median inspection times were 7 and 10 minutes (min). The three groups were as follows: fast speed endoscopists (fast group: median inspection time, < 7 min), moderate speed endoscopists (moderate group: median inspection time, ≥7 min and < 10 min), and slow speed endoscopists (slow group: median inspection time, ≥10 min).

The reason for the EGD was divided into screening and surveillance endoscopy. Screening endoscopy was defined as an inspection for cases without history of UGI neoplasms, such as examination of symptoms and a double cancer check before cancer treatment in another organ. Surveillance endoscopy was defined as periodic examination of cases with surgical and endoscopic treatment for UGI neoplasms. The Kimura-Takemoto classification is generally used for evaluating gastric mucosal atrophy [27]. We classified patients judged to have C-1 atrophic change or higher as atrophic gastritis regardless of presence of Helicobacter pylori.

Excluded (n = 1,166):

- Cases with EGD history within 6 months in our institute (n = 464)
- Cases performed by endoscopists with experience less than three years (n = 369)
- Cases performed by endoscopists with less than 50 EGDs in this period (n = 129)
- Cases that underwent total gastrectomy surgery (n = 67)
- Cases with lacking information on stomach examination (n = 55)
- Cases with residual food in stomach (n = 46)
- Cases using nasal endoscopy examinations (n = 36)

Total cases included in this study (n = 3,925)
UGI neoplasms were defined as hypopharyngeal cancer, esophageal cancer, gastric adenoma/cancer, gastric carcinoid, duodenal adenoma/cancer, and duodenal carcinoid. Gastric neoplasms were defined as gastric adenoma, cancer, and carcinoid. We analyzed the characteristics, biopsy rate, and detection rate in the three groups. The detection rate was calculated as: Detection rate = number of cases in which neoplasms were detected / number of routine EGDs. Furthermore, we compared the detection rate by duration of intensive training at the Cancer Institute Hospital.

**Intensive endoscopic training in the Cancer Institute Hospital**

The Cancer Institute Hospital is a famous high-volume center in Japan, which carries out 12,000 routine EGDs, approximately 450 gastric endoscopic submucosal dissections (ESD), and approximately 200 esophageal ESDs per year. Among the total 12,000 routine EGDs, each trainee can carry out more than 1000 routine EGDs. All patients planned for ESD and surgery undergo preoperative EGD in our hospital as well as a preoperative conference before treatment. Therefore, all trainees have opportunities to learn the endoscopic characteristics and features of approximately 550 surgeries and 450 ESD cases of gastric neoplasm, and approximately 100 surgeries and 200 ESD cases of esophageal neoplasm each year. Thus, intensive training in performing an EGD, endoscopic diagnosis, and treatment of UGI neoplasms are possible among the trainees.

**Statistical analysis**

Statistical analysis was carried out using SPSS software, version 24.0 (SPSS, Chicago, Illinois, United States). We compared categorical parameters using the chi-squared test and continuous parameters using the Mann-Whitney U-test and the Kruskal-Wallis test. P values < 0.05 were determined to be statistically significant.

**Results**

**Characteristics of endoscopists**

All included routine EGDs were carried out by 16 endoscopists. Each endoscopist had carried out a minimum of 1000 EGDs and had ≥3 years of experience before initiation of this study. Eleven were certified by the Japan Gastroenterological Endoscopy Society. A total of 1,931 of 3,925 EGDs (49.2%) were normal without undergoing biopsy. Median inspection time during EGD without undergoing biopsy was 9.3 minutes (interquartile range, 6.6–12.0 min). Three endoscopists were classified into the fast group (median 6.3 min; range, 5.0–7.5 min), five into the moderate group (median 9.0 min; range, 7.0–11.1 min), and eight into the slow group (median 11.8 min; range, 9.0–14.7 min), based on the aforementioned cut-off median inspection times. All endoscopists in the fast group were experts who had intensive training for ≥5 years. Other endoscopist characteristics are shown in Table 1.

**Characteristics of routine EGD between the fast- vs moderate- vs slow-speed endoscopist groups**

Among 5,091 consecutive routine EGDs, 3,925 EGDs satisfying the inclusion criteria were analyzed in this study. Characteristics of routine EGDs in three groups are shown in Table 2. There was no significant difference in age and sex among the three groups. The interval between EGDs was ≤1.5 years in approximately 75% of cases. Approximately half of EGDs were conducted for screening purposes, and half for surveillance purposes in each group. The prevalence of atrophic gastritis was >70% in each group. Approximately 35% of cases had a history of gastric cancer; approximately 10% of cases had a history of esophageal cancer. No significant differences were identified among the three groups in terms of endoscopic system used. The total UGI biopsy rate showed no significant difference among the fast group (53%), the moderate group (49%), and the slow group (51%) (P = 0.075). The proportion of endoscopists with ≥1-year intensive training in the Fast group was significantly higher than those in the other two groups (fast group: 100%; moderate group: 77%; slow group: 63%, P < 0.001).

**Neoplasm detection rate for fast vs moderate vs slow groups**

Neoplasm detection rates are shown in Table 3. The neoplasm detection rate showed no significant difference among the three groups. The detection rate for UGI neoplasms was as follows: fast group, 3.6%; moderate group, 3.3%; slow group, 3.1% (P = 0.807). The detection rate for gastric neoplasms was as follows: fast group, 2.9%; moderate group, 2.1%; slow group, 2.2% (P = 0.336).

As shown in Table 4, 133 neoplasms (131 cases; two cases had both esophageal cancer and gastric cancer) (hypopharynx cancer: seven cases; esophageal cancer: 24 cases; early gastric cancer: 68 cases; gastric adenoma: 24 cases; gastric carcinoid: one case; duodenal cancer: two cases; duodenal adenoma: six cases; duodenal carcinoid: one case) were detected in this study.

**Characteristics of routine EGD between the intensive training <1-year vs ≥1-year groups**

Characteristics of routine EGDs are shown in Table 5. Five endoscopists were classified as having intensive training <1 year and 11 as having ≥1 year, based on duration of intensive training at the Cancer Institute Hospital. The proportion of history of gastric cancer among the Intensive training <1-year group was significantly higher than that among the Intensive training ≥1-year group (<1-year group: 40%; ≥1-year: 34%; P = 0.001). The median inspection times among the Intensive training ≥1-year group were significantly shorter than that among the Intensive training <1-year group (<1-year group: median 6.3 min; range 8.2–13.9 min; ≥1-year group: median 8.9 min; range 6.4–11.4 min; P < 0.001). The other parameters were not different between the two groups.
Comparison of neoplasm detection rates between Intensive training <1-year vs ≥1-year group

As shown in Table 6, detection rates for UGI neoplasms and gastric neoplasms in the Intensive training ≥1-year group were significantly higher than that in the Intensive training <1-year group (UGI neoplasms; <1-year: 2.2%; ≥1-year: 3.7%, odds ratio = 1.65, 95% CI: 1.02–2.68, P=0.041; gastric neoplasms; <1-year: 1.5%; ≥1-year: 2.6%, odds ratio = 1.83, 95% CI: 1.01–3.30, P=0.045).

Discussion

Early detection of UGI neoplasms is ideal for ensuring optimum patient survival. Furthermore, EGD is the most sensitive method for early detection. Although quality control of EGD is necessary in early detection of UGI neoplasms, few studies have assessed quality indicators in EGD. Several quality indicators have been identified in colonoscopy for colorectal cancer screening, such as adenoma detection rate (ADR), inspection time, and surveillance intervals [28–30]. Measuring the ADR of individual colonoscopists is a priority in the quality improvement process for colonoscopy [30]; the same is also true in EGD. However, the detection rate can be affected by prevalence, so simply comparing the detection rate in each hospital is not a good method. Prevalence also varies widely based on geographic location, race, and socioeconomic status [31]. Especially, the predominant risk factor for gastric cancer is Helicobacter pylori infection, and major risk factors for esophageal squamous cell cancer include consumption of alcohol and smoking. Thus, in the current study, we compared the detection rate in the same hospital, as there is no associated bias with prevalence.

Previous studies have shown that slow-speed endoscopists had a higher detection rate for UGI neoplasms than fast-speed endoscopists [23–25]. In this study, the neoplasm detection rates showed no significant difference among the fast, moderate, and slow groups. There was no association between inspection time and neoplasm detection rates. However, median inspection time in EGD without undergoing biopsy was 9.3 minutes, which was longer than that in previous reports. Moreover, median inspection time in the fast group was 6.3 minutes, and even the fastest endoscopist had a median inspection time ≥6 minutes. Meanwhile, previous reports have indicated that in-
inspection time is a quality indicator for EGD; Kawamura et al. [23] (fast-speed endoscopist group mean inspection time, 4.4 min), Teh et al. [24] (fast-speed endoscopist group mean inspection time, 5.5 min), and Park et al. [25] (fast-speed endoscopist group mean inspection time, 2.38 minutes, defined as the time from when the endoscope reached the duodenum to the time it was withdrawn) have suggested that EGD be carried out with a shorter inspection time than that seen in this study. Thus, according to the cut-off inspection times shown in previous reports, the endoscopists classified as fast in this study (median inspection time ≥ 6 min) had a suitable inspection time to decrease the likelihood of overlooking UGI neoplasms. However, all endoscopists in the fast group in this study were expert endoscopists who had sufficient experience (intensive training at our hospital for ≥ 5 years).

In terms of endoscopic training and experience, previous studies reported that inexperienced endoscopists tend to overlook gastric cancer [12–14]. This means that endoscopists re-
### Table 4 Characteristics of the all detected neoplasms in the fast- vs moderate- vs Slow-speed endoscopist groups.

<table>
<thead>
<tr>
<th>Neoplasm</th>
<th>Fast-speed endoscopists group (n = 1,052)</th>
<th>Moderate-speed endoscopists group (n = 1,314)</th>
<th>Slow-speed endoscopists group (n = 1,559)</th>
<th>Total (n = 3,925)</th>
</tr>
</thead>
<tbody>
<tr>
<td>UGI neoplasms (%)</td>
<td>38 (3.6)</td>
<td>44 (3.3)</td>
<td>49 (3.1)</td>
<td>131 (3.3)</td>
</tr>
<tr>
<td>Hypopharynx cancer (%)</td>
<td>0 (0)</td>
<td>4 (0.30)</td>
<td>3 (0.19)</td>
<td>7 (0.18)</td>
</tr>
<tr>
<td>Esophageal cancer (%)</td>
<td>8 (0.76)</td>
<td>9 (0.67)</td>
<td>7 (0.45)</td>
<td>24 (0.61)</td>
</tr>
<tr>
<td>Gastric neoplasm (%)</td>
<td>31 (2.9)</td>
<td>27 (2.1)</td>
<td>35 (2.2)</td>
<td>93 (2.4)</td>
</tr>
<tr>
<td>Gastric cancer (%)</td>
<td>24 (2.3)</td>
<td>20 (1.5)</td>
<td>24 (1.5)</td>
<td>68 (1.7)</td>
</tr>
<tr>
<td>Gastric adenoma (%)</td>
<td>7 (0.67)</td>
<td>6 (0.46)</td>
<td>11 (0.71)</td>
<td>24 (0.61)</td>
</tr>
<tr>
<td>Gastric carcinoid (%)</td>
<td>0 (0)</td>
<td>1 (0.08)</td>
<td>0 (0)</td>
<td>1 (0.03)</td>
</tr>
<tr>
<td>Duodenal cancer (%)</td>
<td>0 (0)</td>
<td>1 (0.08)</td>
<td>1 (0.06)</td>
<td>2 (0.05)</td>
</tr>
<tr>
<td>Duodenal adenoma (%)</td>
<td>1 (0.10)</td>
<td>2 (0.15)</td>
<td>3 (0.19)</td>
<td>6 (0.15)</td>
</tr>
<tr>
<td>Duodenal carcinoid (%)</td>
<td>0 (0)</td>
<td>1 (0.08)</td>
<td>0 (0)</td>
<td>1 (0.03)</td>
</tr>
</tbody>
</table>

UGI, upper gastrointestinal

1 Two cases in the fast-speed endoscopists group had double cancers of esophageal cancer and gastric cancer.

### Table 5 Characteristics of routine EGD between the Intensive training < 1-year vs ≥ 1-year groups.

<table>
<thead>
<tr>
<th>Variable</th>
<th>Intensive training &lt; 1-year group (n = 891)</th>
<th>Intensive training ≥ 1-year group (n = 3,034)</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age, median (range)</td>
<td>67 (21–88)</td>
<td>67 (18–93)</td>
<td>0.521</td>
</tr>
<tr>
<td>Sex (%) Male/female</td>
<td>542 (61)/349 (39)</td>
<td>1894 (62)/1140 (38)</td>
<td>0.388</td>
</tr>
<tr>
<td>Interval between EGDs (%)</td>
<td></td>
<td></td>
<td>0.606</td>
</tr>
<tr>
<td>≤ 1.5 years</td>
<td>699 (79)</td>
<td>2368 (78)</td>
<td></td>
</tr>
<tr>
<td>&gt; 1.5 years</td>
<td>135 (15)</td>
<td>447 (15)</td>
<td></td>
</tr>
<tr>
<td>Unknown</td>
<td>56 (6)</td>
<td>219 (7)</td>
<td></td>
</tr>
<tr>
<td>Purpose of EGD (%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Screening/Surveillance</td>
<td>405 (46)/486 (54)</td>
<td>1479 (49)/1555 (51)</td>
<td>0.084</td>
</tr>
<tr>
<td>Gastric mucosal atrophy (%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Atrophy/non-atrophy</td>
<td>647 (73)/244 (27)</td>
<td>2169 (72)/865 (28)</td>
<td>0.512</td>
</tr>
<tr>
<td>Past history of gastric cancer (%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>With/without</td>
<td>359 (40)/532 (60)</td>
<td>1033 (34)/2001 (66)</td>
<td>0.001</td>
</tr>
<tr>
<td>Past history of esophageal cancer (%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>With/without</td>
<td>91 (10)/800 (90)</td>
<td>361 (12)/2673 (88)</td>
<td>0.166</td>
</tr>
<tr>
<td>Endoscope model (%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>GIF-H260/GIF-H290Z</td>
<td>432 (49)/459 (51)</td>
<td>1367 (45)/1667 (55)</td>
<td>0.071</td>
</tr>
<tr>
<td>Light source model (%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>SPECTRUM/ELITE</td>
<td>425 (48)/466 (52)</td>
<td>1351 (44)/1683 (56)</td>
<td>0.095</td>
</tr>
<tr>
<td>UGI biopsy cases (%) Done/Not done</td>
<td></td>
<td></td>
<td>0.274</td>
</tr>
<tr>
<td>Inspection time, min, median (range)</td>
<td>11.0 (8.2–13.9)</td>
<td>8.9 (6.4–11.4)</td>
<td>&lt;0.001</td>
</tr>
</tbody>
</table>

EGD, esophagogastroduodenoscopy

1 Interquartile range
require sufficient training to detect gastric cancer properly. In this study, although all trainees had experienced more than 1000 EGDs in another hospital, the detection rate for UGI and gastric neoplasms by endoscopists with ≥ 1-year intensive training was significantly higher than that by endoscopists with < 1-year intensive training. In addition, endoscopists with ≥ 1-year intensive training have significantly shorter inspection times than endoscopists with < 1-year intensive training. Thus, intensive training for ≥ 1 year made it possible to detect more neoplasms within a shorter examination time. This indicates that the quality of EGD, as measured by neoplasm detection rates, may be improved by ≥ 1-year intensive training. Each trainee at our hospital can carry out approximately 1000 EGDs each year. Moreover, each trainee has opportunities to learn from the endoscopic findings of approximately 1000 cases of gastric neoplasm, and 300 cases of esophageal cancer each year. To improve the detection rate, it is also necessary to gain experience identifying many tumor characteristics and features in addition to technical training in EGD.

In this study, the biopsy rate (50.8 %) was slightly high, which may be because we tend to perform many biopsies, as it is a cancer specialty hospital visited by many patients with a high cancer risk. Approximately 35 % of cases showed gastric cancer history, approximately 10 % of cases showed esophageal cancer history, and more than 70 % of cases showed atrophic gastritis, which has a high risk of progressing to gastric cancer. In these patients, we usually perform more careful examinations to avoid overlooking small lesions. Accordingly, nearly all neoplasms found in this study were determined to be early-stage cancer.

The current study had some limitations. First, it was a non-randomized, retrospective, single-center study. Second, this study included a small number of cases. Third, a patient-selection bias might have been present. To reduce selection bias, we included all consecutive EGDs performed in a period where the staff of endoscopists was unchanged. Further, we planned the study after the case selection period. Therefore, we believe information bias from the endoscopists is relatively low. However, future prospective study with randomization in multiple centers is necessary to validate our findings.

### Conclusion

In conclusion, there was no association between inspection time and UGI neoplasm detection rates. Endoscopists with ≥ 1-year intensive training have significantly higher UGI neoplasm detection rates than endoscopists with < 1 year of intensive training, although inspection times were significantly shorter. Accordingly, our results demonstrated that the quality of EGD, as measured by UGI neoplasm detection rates, may be improved by intensive training.

### Competing interests

None

### References