Changes in fatigue and physical function with age for patients with gastrointestinal cancer in the perioperative period: a comparison between older and young patients

EISUKE KOGURE, RPT1, 2)*, TSUYOSHI HARA, RPT, PhD3), TAKAYA ISHII, RPT2), MASAHARU MAEDA, MD1)

1) Department of Rehabilitation, International University of Health and Welfare Graduate School: 1-3-3 Minami Aoyama, Minato-ku, Tokyo 107-0062, Japan
2) Division of Rehabilitation, International University of Health and Welfare Mita Hospital, Japan
3) Department of Physical Therapy, Faculty of Health Science, International University of Health and Welfare, Japan

Abstract. [Purpose] The purpose of this study was to examine changes in fatigue and physical function with age for groups of patients with gastrointestinal cancer in the perioperative period. [Subjects and Methods] The study involved 52 patients with gastrointestinal cancer (28 males and 24 females, aged 62.4 ± 12.0 years). The subjects were divided into two groups: those 65 or over (the so-called elderly group,) and those 64 or under (the so-called young group). The six minute walk distance (6MWD), Cancer Fatigue Scale (CFS), and serum albumin (ALB) were evaluated on three occasions: before surgery, after surgery and after discharge. This study considered change in physical function and fatigue, differences between the two groups and correlations between age and each evaluation on the three occasions. [Results] The 6MWD was the main difference between the elderly and young groups, with the former having significantly reduced 6MWD values as compared to the latter after surgery. Age was positively correlated with 6MWD and ALB after surgery and after discharge. Moreover, age was negatively correlated with CFS after surgery and after discharge. [Conclusion] It is suggested that elderly patients are more likely to experience a decline in exercise tolerance and an increase in fatigue.

Key words: Patients with gastrointestinal cancer in the perioperative period, Fatigue, Older and young patients

INTRODUCTION

Fatigue is a symptom experienced by cancer patients and one of the factors affecting their quality of life (QOL). The National Comprehensive Cancer Network (NCCN) defines cancer-related fatigue (CRF) as “a distressing, persistent, subjective sense of physical, emotional, and/or cognitive tiredness or exhaustion related to cancer or cancer treatment that is not proportional to recent activity and interferes with usual functioning” 1). Previous studies have shown a prevalence of fatigue of greater than 30% for cancer patients and cancer survivors2, 3), and greater than 50–80% for advanced and end-of-life cancer patients4, 5). Improvement in physical function has been reported as decreased fatigue and improved QOL in cancer patients6, 7). In addition, anxiety, depression and psychological stress have been reported as being correlated with fatigue8). Hence, fatigue is associated with physical function and the psychological factors of anxiety and depression. However, report on the occurrence of fatigue was few studies that after chemotherapy, radiotherapy and lung, prostate,
breast, terminal cancer patients is enough6–11, but gastrointestinal cancer patients is not enough. The study of physical activity and function gastrointestinal cancer patients that cancer incidence rates in Japan accounts for about 45%12 was examined preceding study13–15, but fatigue is not enough. The factor of decrease physical activity at before operation with breast and prostate cancer patients were correlated age16, 17, therefore there is a possibility as an elderly patients decrease in physical function and increase fatigue. The purpose of the present study was to examine changes in fatigue and physical function with age for patients with gastrointestinal cancer in the perioperative period, and correlation of age and each evaluation on the three occasions.

SUBJECTS AND METHODS

The study involved 52 patients with gastrointestinal cancer (28 males and 24 females, aged 62.4 ± 12.0 years) from September 2014 to May 2016. The exclusion criteria were as follows: physical and cognitive dysfunction; psychiatric disorders; not independence Activity of Daily Living (ADL), anyone who said that it was impossible of a gait based on the judgment of the doctor in charge; development of postoperative complications. The patient characteristics that were examined were age, gender, stage classification of cancer (cancer stage), operation time, blood loss, and Body Mass Index (BMI). Nutrition status was evaluated using serum albumin (ALB) levels, and inflammation was evaluated using C-reactive protein (CRP).

Fatigue was evaluated using the Cancer Fatigue Scale (CFS)18, which is correlation of any questionnaire of fatigue. Moreover, it is possible evaluate to conveniently questionnaire of fatigue19. The CFS consists of three subscales (namely the physical, affective, and cognitive aspect of fatigue) and a 15-item self-rating scale for assessing fatigue in cancer patients. The questionnaire was answered using ratings of 1 (not at all) to 5 (very much).

The maximum total score was 60, with the higher scores indicating. The cut-off score was 19; anything higher indicated severe fatigue that more failure ADL20.

Anxiety and depression were evaluated using the Japanese version of the Hospital Anxiety and Depression Scale (HADS). This consists of a 14-item self-rating scale (assessing anxiety and depression) and is composed of seven-item anxiety and depression subscales. The associated questionnaire was answered in choices of 1 to 4. Total anxiety and depression were calculated as the sum of these aspects (7 points or fewer meant no diagnosis, between 8 and 10 points meant a suspected diagnosis, and 11 points or more meant a certain diagnosis21, 22).

Physical function was evaluated using the six minute walk distance (6MWD). This was measured using a distance meter and was performed according to the stated guidelines of the American Thoracic Society23. Evidence presented in a preceding study supports the validity of the 6MWD as a measure of recovery after elective colon resection surgery24. Moreover, the 6MWD was selected to give evaluable evidence of physical function and exercise tolerance.

Each parameter was measured on three occasions (before surgery, after surgery, and after discharge). The subjects of the present study were divided into two groups: those who were 65 years old or over (the so-called elderly group) and those who were 64 years old or under (the so-called young group). The study considered change, differences between the two groups, and correlations between age and each evaluation at each of the three occasions.

The two groups were compared based on age, operation time, blood loss, length of day were used T value test, cancer stage, and surgical site using χ² tests. Any p-value of less than 0.05 was considered statistically significant. The data were analyzed using SPSS Statistics 23.0 for Windows. Change of three occasions were used a two-way factorial analysis of variance (ANOVA) (factors: evaluation times and age) and a multiple comparison test (Bonferroni) to compare the studied parameters. Again, and p-value of less than 0.05 was considered statistically significant.

The present study was approved by the ethics committee of the International University of Health and Welfare graduate school (Approval Number: 15-Ig-43).

RESULTS

The characteristics of the subjects are listed in Table 1. No significant were characteristics of subjects. In the multiple-comparison test, 6MWD showed significant differences between the elderly and young groups after discharge, with the young group having a significantly extended 6MWD compared to the elderly group. The CFS was no main effect and interaction between the elderly and young groups, but the elderly group had higher and more dispersed CFS values compared to the young group. The young group were distance extender 6MWD compared to the elderly group at the three measurement occasions. For both groups, the 6MWD was shorter after surgery than before (Table 2). There were positive correlations between (i) age and 6MWD and (ii) age and ALB after surgery and after discharge. Moreover, age was negatively correlated with CFS after surgery and after discharge (Table 3).

DISCUSSION

The present study examined fatigue and physical function compared to age (essentially either older or younger than 65) and relationship age and each parameter of patients with gastrointestinal cancer in the perioperative period.

Fatigue was found to be not significant between those over 65 and those 64 or under. In a preceding study, fatigue was
### Table 1. Characteristics of patients with gastrointestinal cancer in the perioperative in this study

|                         | All Patients | Elderly group | Young group |
|-------------------------|--------------|---------------|-------------|
| Subjects (n)            | 52           | 23            | 29          |
| Age (years)             | 62.4 ± 12.0  | 73.2 ± 6.1    | 53.1 ± 7.8  |
| Gender (n)              |              |               |             |
| Males                   | 18           | 11            | 17          |
| Females                 | 24           | 12            | 12          |
| Cancer stage (n)        |              |               |             |
| I                       | 25           | 9             | 16          |
| II                      | 13           | 7             | 6           |
| III                     | 10           | 6             | 4           |
| IV                      | 4            | 1             | 3           |
| Surgical site (n)       |              |               |             |
| Stomach                 | 17           | 8             | 9           |
| Pancreas                | 1            | 1             | 0           |
| Colon                   | 26           | 10            | 16          |
| Rectum                  | 8            | 4             | 4           |
| Surgical procedure (n)  |              |               |             |
| Laparoscopic            | 43           | 18            | 25          |
| Laparotomy              | 9            | 5             | 4           |
| Operation time (minutes)| 246.7 ± 73.3 | 252.2 ± 74.5  | 242.3 ± 73.5 |
| Blood loss (ml)         | 156.7 ± 368.9| 215.1 ± 454.6 | 110.4 ± 283.8 |
| Length of stay (day)    | 13.9 ± 4.9   | 15.5 ± 4.8    | 12.7 ± 3.3  |

The value of age, operation time, blood loss and length of stay are shown as the mean ± standard deviation.

### Table 2. Each parameter at three occasions

|                         | Elderly group | Young group |
|-------------------------|---------------|-------------|
| CFS (score)             |               |             |
| Before surgery          | 15 (11, 20)   | 14 (9, 18)  |
| After surgery           | 15 (11, 23)   | 12 (8, 16)  |
| After discharge         | 14 (9, 20)    | 10 (5, 14)  |
| HADS Anxiety (score)    |               |             |
| Before surgery          | 4 (3, 6)      | 4 (3, 7)    |
| After surgery           | 3 (2, 6)      | 3 (2, 5)    |
| After discharge         | 3 (2, 5)      | 2 (1, 5)    |
| HADS Depression (score) |               |             |
| Before surgery          | 5 (2, 7)      | 3 (1, 6)    |
| After surgery           | 3 (1, 6)      | 2 (1, 5)    |
| After discharge         | 4 (1, 7)      | 2 (1, 4)    |
| BMI (kg/m²)             |               |             |
| Before surgery          | 22.8 ± 3.6    | 21.9 ± 3.0  |
| After surgery           | 21.7 ± 3.4    | 20.8 ± 2.8  |
| After discharge         | 22.0 ± 3.3    | 21.1 ± 2.9  |
| ALB (g/dl)              |               |             |
| Before surgery          | 4.3 ± 0.3     | 4.4 ± 0.3   |
| After surgery           | 3.6 ± 0.3     | 3.8 ± 0.3   |
| After discharge         | 4.1 ± 0.3     | 4.3 ± 0.3   |
| CRP (mg/dl)             |               |             |
| Before surgery          | 0.4 ± 0.9     | 0.2 ± 0.3   |
| After surgery           | 1.8 ± 1.9     | 1.8 ± 2.0   |
| After discharge         | 0.7 ± 2.1     | 0.2 ± 0.5   |
| 6MWD (m)                |               |             |
| Before surgery          | 530.2 ± 83.9  | 567.6 ± 112.5 |
| After surgery           | 466.3 ± 83.2  | 519.1 ± 102.8 |
| After discharge         | 514.2 ± 81.9  | 587.7 ± 103.7** |

The values of CFS, HADS Anxiety and HADS Depression are shown as the median (25th percentile, 75th percentile)
The values of BMI, ALB, CRP and 6MWD are shown as the mean ± standard deviation.

*A two-way factorial analysis of variance main effect
**Multiple comparison test (Bonferroni)

### Table 3. Correlation between age and each parameter at three occasions

| Age (years) | 6MWD (m) | CFS (score) | HADS Anxiety (score) | HADS Depression (score) | ALB (g/dl) | CRP (mg/dl) |
|-------------|----------|-------------|----------------------|-------------------------|------------|-------------|
| Before surgery | −0.225  | 0.062       | 0.054                | 0.103                   | −0.239     | 0.075       |
| After surgery  | −0.291* | 0.353*      | −0.023               | 0.113                   | −0.351*    | 0.040       |
| After discharge| −0.369* | 0.293*      | 0.146                | 0.190                   | −0.453*    | 0.265       |

*Spearman's rank correlation coefficient
found to be not significant in a comparison between those over 60 and those 59 years old or under at the preoperative stage. Moreover, fatigue was found to be significant between patients with high and low fatigue at one-month postoperative. These findings suggest that comparisons between about 60 and about 65 between high and low fatigue are not significant.

A limitation of this study was that we did not consider detail of physical activities and physical condition before hospitalization. A future study will be necessary to examine fatigue, QOL, physical and psychological function at before Hospitalization, in addition to relationship between ages.

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