ORIGINAL ARTICLE



The type of gastrectomy affects skeletal muscle loss and the long-term outcomes of elderly patients with gastric cancer: a retrospective study using computed tomography images

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Abstract

Purpose Sarcopenia is common in elderly gastrectomized patients and a known risk factor for postoperative complications and poor overall survival. However, the long-term outcomes of skeletal muscle loss after gastrectomy and the differences in outcomes of different gastrectomy procedures remain unclear.

Methods The subjects of this retrospective study were 136 patients who underwent various gastrectomy procedures for early gastric cancer, namely: total gastrectomy (TG; n = 20), proximal gastrectomy (PG; n = 16), distal gastrectomy (DG; n = 60), and pylorus-preserving gastrectomy (PPG; n = 40). Skeletal muscle volume (SMV), calculated as the skeletal muscle index (SMI), was measured using cross-sectional computed tomography (CT) scans preoperatively and then 1, 2, and 3 years after gastrectomy.

Results Sarcopenia developed from 2 years onwards in all the patients who underwent TG. The SMI and sarcopenia prevalence after gastrectomy deteriorated over time. Multivariate analysis revealed that TG and PG were significant risk factors for skeletal muscle loss in postoperative years 1 and 3. A decrease in the SMI after TG or PG was most remarkable in elderly patients.

Conclusions The type of gastrectomy affects skeletal muscle loss in the long term. Elderly patients who undergo TG or PG are at high risk of severe skeletal muscle loss.

Keywords Gastrectomy · Gastric cancer · Postoperative sarcopenia · Risk factor · Skeletal muscle loss

Introduction

In recent years, high response rates of chemotherapy have been reported, but radical gastrectomy remains the standard treatment for gastric cancer [1]. Studies have found that nutritional status impacts patient survival [2, 3] and quality of life [4]; therefore, nutritional therapy is critical for patients receiving oncological therapy after gastrectomy.

Rosenberg [5] defined sarcopenia as "low muscle mass plus low muscle strength and/or low physical performance in the elderly". Recent studies have shown that preoperative sarcopenia is a risk factor for perioperative complications [6–8], chemotherapy intolerance [9], and impaired overall survival after gastric cancer surgery [10]. In contrast, few reports have investigated secondary sarcopenia following gastrectomy, even though gastrectomy causes malabsorption and skeletal muscle loss. Yamaoka et al. reported that after total gastrectomy (TG), the skeletal muscle index (SMI) decreased by 6.2% (±6.8%) from the preoperative value [11]. Yet, long-term postoperative changes in skeletal muscle volume (SMV) and perioperative risk factors for sarcopenia secondary to gastrectomy remain unclear.

In this study, we investigated the patterns of skeletal muscle loss following gastrectomy for gastric cancer and tried to

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identify the patients and treatment-specific factors that are associated with long-term skeletal muscle loss contributing to a worse prognosis. We evaluated the SMV and subcutaneous and visceral fat compartments based on the findings of computed tomography (CT), performed annually for 3 years in patients who underwent one of four types of gastrectomy for early-stage gastric cancer: TG, proximal gastrectomy (PG), distal gastrectomy (DG), or pylorus-preserving gastrectomy (PPG).

Methods

Study population

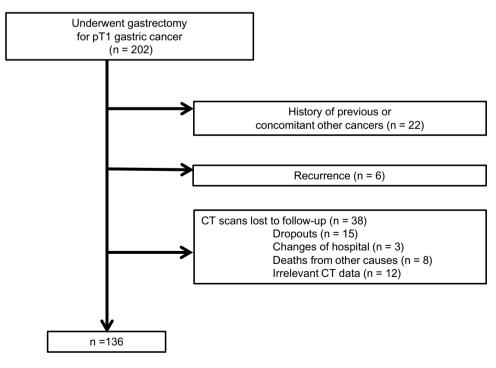
This retrospective study included 202 consecutive patients who underwent gastrectomy (TG, PG, DG, or PPG) for pathological T1-classified gastric cancer (mucosa and submucosa), according to the Japanese classification of gastric cancer [12], between January, 2012 and December, 2014, at the University of Tokyo Hospital in Tokyo, Japan. 22 patients with previous or concomitant cancers and 6 patients with recurrent disease were excluded from the study. The skeletal muscle area could not be measured at all in 38 patients whose CT scans were lost during follow-up and the skeletal muscle mass could not be measured at the level of the third lumbar vertebra (L3) on the CT scans of 12 patients. 15 patients withdrew from follow-up, 3 changed hospitals, and 8 died of diseases other than cancer. Finally,

136 patients were included in the analysis (Fig. 1). None of the patients received adjuvant or neoadjuvant chemotherapy. This retrospective study was approved by the University of Tokyo, Faculty of Medicine Ethics Committee (ID: 3962) and complies with the ethical standards of the Declaration of Helsinki.

CT image analysis of body composition

We reviewed CT scans taken preoperatively and then 1, 2, and 3 years postoperatively, which were required for the evaluation of tumor staging and screening for recurrence, for each of the patients who underwent gastrectomy. The cross-sectional abdominal CT inferior aspect of L3 was assessed to estimate the muscle mass. Images were analyzed using OsiriX (ver. 8.5.1, Pixmeo, Switzerland), which is open-source software that enables the examination of specific tissues using Hounsfield units (HUs). Measurements were recorded in a semi-automated fashion by setting the tissue of interest threshold at -29 to +150 HUs for skeletal muscle, -190 to -30 for subcutaneous adipose tissue [13], and -150 to -50 for visceral adipose tissue [11]. Skeletal muscle areas (SMA; cm²) in the L3 region computed from each image were normalized for height squared (m²) to obtain the SMI at the level of the inferior aspect of L3. Sarcopenia was defined as L3 SMI < 52.4 cm²/m² in men and $< 38.5 \text{ cm}^2/\text{m}^2$ in women [14]. At the same time, subcutaneous and visceral fat compartments for the subcutaneous fat area (SFA) and visceral fat area (VFA) were evaluated at

Fig. 1 Patient selection process



CT, computed tomography



the equivalent L3 level as the skeletal muscle area. Figure 2 shows typical transverse CT images at L3.

Statistical analyses

Statistical analyses were performed using JMP software (Pro 14, SAS Institute, Cary, NC, USA). Data are expressed as the mean \pm standard error. Categorical variables were compared using the χ^2 test or Fisher's exact test. Multivariate logistic regression was performed. A P < 0.05 was considered significant. The following perioperative factors were analyzed: sex, age, preoperative sarcopenia, comorbidity based on the Clavien–Dindo (C–D) classification (C–D \geq III) [15], surgical procedure (DG or PPG versus PG or TG), and lymphadenectomy (D1 or D1 + versus D2). In accordance with previous reports, we defined severe skeletal muscle loss (SSML) as a 10% decrease in SMI from the preoperative value [11, 16]. Operative time and blood loss were divided into two groups, using their average value as the cut-off point.

Results

Patient characteristics

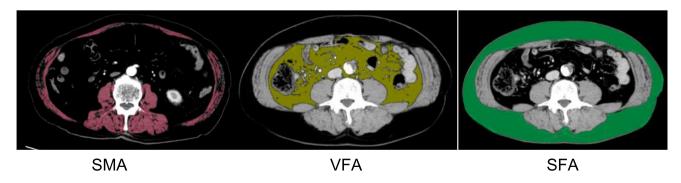
Table 1 summarizes the clinical characteristics of the 136 patients in the study cohort. The study population comprised 94 men and 42 women, with a mean age of $66.06~(\pm\,10.76)$ years. The preoperative mean SMI was $47.84~(\pm\,8.10)~\text{cm}^2/\text{m}^2$ and the prevalence of preoperative sarcopenia was 54.4%. Table 1 lists the clinical characteristics of the patients, who were divided into four groups according to the gastrectomy procedure they underwent. The average operative blood loss was higher in the TG group than in the other groups, and the average operative time was longer for the DG group than for the PG or TG groups.

Changes in SMI, VFA, and SFA

Change in SMI was calculated by the rate of change from preoperative values. Because the values for VFA and SFA were close to 0 in lean patients and the rate of change was an outlier, VFA and SFA were calculated based on the amount of change. Figure 3 shows the rate of change in SMI and changes in VFA and SFA for each group in each postoperative year. The SMI was significantly lower than the preoperative value for every group in postoperative years 1, 2, and 3. The rate of change in SMI for the DG group decreased mainly in the first 2 years after gastrectomy and stabilized in postoperative year 3. The rate of change in SMI for the PPG group decreased mainly in postoperative year 1, then stabilized in postoperative years 2 and 3. The rate of change in SMI for the TG and PG groups decreased continually until postoperative year 3. There were significant differences in the rate of change in SMI between the DG and the TG groups in postoperative year 1 (P = 0.0223), the DG and the PG groups in postoperative years 1, 2, and 3 (P = 0.0020, P = 0.00400, P = 0.0214), and the PPG and the PG groups in postoperative year 3 (P = 0.0361). The rate of change in SMI for the PPG and PG groups were not significantly different from those for the DG and TG groups. VFA and SFA decreased significantly from the preoperative values in both groups every year; mainly in postoperative years 1 and 2 for VFA and mainly in postoperative year 1 for SFA.

Prevalence of sarcopenia evaluated using SMI after the different types of gastrectomy

Figure 4 shows changes in the prevalence of sarcopenia evaluated using SMI over time for each of the four groups. The numbers of patients with sarcopenia identified in each group preoperatively and then 1, 2, and 3 years postoperatively were as follows: 32, 37, 40, and 40 in the DG group (53.3%, 61.7%, 66.7%, 66.7%); 21, 25, 25, and 25 (52.5%, 62.5%,



SMA, skeletal muscle area; VFA, visceral fat area; SFA, subcutaneous fat area

Fig. 2 Measurements of skeletal muscle area (SMA), visceral fat area (VFA), and subcutaneous fat area (SFA): axial computed tomograms were taken at the inferior level of the third lumbar vertebra (L3)



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Table 1 Clinical characteristics of all patients and of the subgroups broken down by procedures

	=					
	ALL	TG	PG	DG	PPG	P value
	n = 136	n = 20	n = 16	n = 60	n = 40	
Sex (M/F)	94/42	20/0	12/4	38/22	24/16	0.0085
Age $(\pm SD)$	$66.06 \ (\pm 10.76)$	$70.5 (\pm 8.56)$	$64.5 (\pm 11.58)$	$67.60 (\pm 10.49)$	$62.5 (\pm 11.11)$	0.0284
Stage (%)						0.439
IA	120	17 (80.9%)	14 (87.5%)	54 (90.0%)	35 (87.5%)	
IB	9	2 (14.2%)	1 (6.2%)	3 (5.0%)	3 (7.5%)	
IIA	6	1 (4.7%)	0 (0%)	3 (5.0%)	2 (5.0%)	
IIB	1	0 (0%)	1 (6.2%)	0 (0%)	0 (0%)	
Lymphadenectomy (%)						< 0.0001
D1 (includeD1+)	113	18	16	39	40	
D2	23	2	0	21	0	
Open/laparoscopic surgery	68/68	20/0	16/0	28/32	4/36	< 0.0001
Reconstruction methods		R-Y=20	EG:JI=7:9	B-I: $R-Y = 6:54$		
Operative time $(\pm SD)$ (min)	$248.43 \ (\pm 67.79)$	$213.4 (\pm 9.03)$	$212.18 (\pm 18.81)$	$265.53 (\pm 9.64)$	$254.8 (\pm 8.42)$	0.0022
Operative blood loss $(\pm SD)$ (ml)	$181.98 (\pm 264.91)$	$437.75 (\pm 100.20)$	$219.68 \ (\pm 60.58)$	$131.50 (\pm 31.28)$	$114.75 (\pm 38.31)$	< 0.0001
Comorbidity						0.9130
C-D 0	106	15(75.0)	13 (81.2)	48 (80.3)	30(75.0)	
C–D I	6	2 (10.0)	0 (0)	3 (4.9)	1 (2.5)	
C–D II	18	3(15.0)	2 (12.5)	6 (9.8)	6 (15.0)	
C–D III	6	0 (0)	1 (6.2)	3 (4.9)	2 (5.0)	
Preoperative BMI (±SD)	$22.67 (\pm 3.11)$	$23.40 (\pm 2.66)$	$24.23 (\pm 4.81)$	$22.41 (\pm 2.94)$	$22.03 (\pm 2.46)$	0.0671
Preoperative SMI (±SD)	$47.84 (\pm 8.10)$	$49.14 (\pm 7.43)$	$51.44 (\pm 9.19)$	$46.92 (\pm 7.53)$	$47.12 (\pm 8.62)$	0.921
Preoperative prevalence of sarcopenia (%)	74/136 (54.4%)	14/21 (66.7%)	7/16 (43.8%)	32/61 (52.5%)	21/40 (52.5%)	0.4269
Preoperative VFA (±SD) (cm ²)	93.39 (±74.72)	$113.96 (\pm 69.60)$	130.92 (±8403)	$89.65 (\pm 76.71)$	$73.16 (\pm 63.08)$	0.0332
Preoperative SFA $(\pm SD)$ (cm ²)	$105.27 (\pm 50.75)$	$104.51 (\pm 43.29)$	125.96 (±84.02)	99.42 (± 45.40)	$105.21 (\pm 44.02)$	0.3513

TG, total gastrectomy; PG, proximal gastrectomy; DG, distal gastrectomy; PPG, pylorus-preserving gastrectomy; R-Y, Roux-en-Y reconstruction; EG, esophagogastrostomy; JI, jejunal interposition; B-I, Billroth-I reconstruction; C-D, Clavien-Dindo classification; BMI, body mass index; SMI, skeletal muscle index; VFA, visceral fat area; SFA, subcutaneous fat area

62.5%, 62.5%) in the PPG group; 14, 16, 20 and 20 (70.0%, 80.0%, 100%, 100%) in the TG group; and 7, 9, 9, and 10 (43.8%, 56.3%, 56.3%, 62.5%) in the PG group. The prevalence of sarcopenia increased by approximately 10%–20% in the DG, PPG, and PG groups. However, in the TG group, the prevalence of sarcopenia was significantly higher than in any of the other groups, increasing from 71.4% to 100% in postoperative years 2 and 3.

Risk factors for postoperative SSML

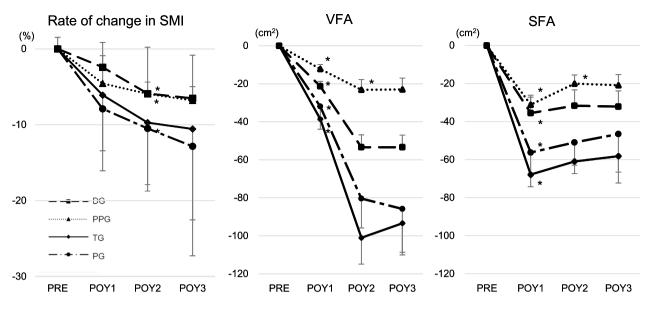
Severe skeletal muscle loss of more than 10% from the preoperative value was seen 1, 2, and 3 years after surgery in 20 patients (14%), 46 patients (33%), and 53 patients (38%), respectively. Table 2 summarizes the results of multivariate analyses of six clinical risk factors for SSML: sex, age, preoperative sarcopenia, comorbidities, TG or PG, and lymphadenectomy. Multivariate analysis identified TG and PG as independent risk factors for SSML in postoperative years 1 and 3 (odds ratio [OR] 4.08, 95% confidence interval

[CI] 1.44–12.05; OR 2.32, 95% CI 1.06–5.19). Age was an independent risk factor for SSML in postoperative year 2 (OR 2.42, 95% CI 1.51–5.34), and this tendency was also observed in postoperative years 1 and 3.

Rate of change in SMI as stratified by age

Sarcopenia is associated with compromised outcomes in the elderly, so we subdivided patients into an elderly group (age \geq 65 years [n=79]) and a younger group (under 65 years old [n=57]) for comparison. The preoperative SMI of the elderly patients was 46.89 (\pm 7.87) cm²/m² and that of the younger patients was 48.52 (\pm 8.25) cm²/m². There was no significant difference in preoperative SMI values between the elderly and younger patients. Figure 5 shows the rate of change in SMI in the elderly and younger patients according to the four types of gastrectomy. The surgical procedure had a greater impact on the elderly patients than on the younger patients. In postoperative year 3, the rate of change in SMI of the TG and PG patients in the elderly group decreased





*:Significant decrease compared to the previous year (*P*<0.05)

	Rate of change in SMI (%)				VFA (cm ²)			SFA (cm ²)	
	POY1	POY2	POY3	POY1	POY2	POY3	POY1	POY2	POY3
DG	-2.4 (±5.6)	-5.8 (±8.6)	-6.5 (±9.2)	-21.3 (±2.6)	-53.4 (±6.5)	-53.4 (±6.4)	-35.6 (±3.8)	-31.7 (±4.0)	-32.2 (±3.9)
PPG	-4.5 (±5.4)	-5.9 (±6.1)	-6.7 (±5.9)	-12.2 (±2.3)	-23.2 (±5.3)	-23.0 (±6.0)	-31.1 (±5.0)	-20.0 (±4.6)	-20.9 (±5.6)
TG	-6.1 (±7.3)	-9.7 (±8.1)	-10.5 (±11.9)	-38.6 (±5.3)	-101.0 (±13.9)	-93.4 (±16.7)	-67.9 (±6.4)	-61.0 (±6.4)	-58.2 (±8.4)
PG	-7.8 (±8.1)	-10.4 (±8.2)	-12.8 (±14.4)	-31.9 (±6.0)	-80.4 (±15.5)	-85.9 (±22.9)	-56.3 (±11.8)	-51.0 (±12.0)	-46.5 (±25.8)

SMI, skeletal muscle index; VFA, visceral fat area; SFA, superficial fat area; TG, total gastrectomy; PG, proximal gastrectomy; PG, distal gastrectomy; PPG, pylorus-preserving gastrectomy; PRE, preoperative year; POY, postoperative year

Fig. 3 Average rate of change in the skeletal muscle index (SMI) decreased every postoperative year in each group. The visceral fat area (VFA) began to increase from 2 years postoperatively. The subcutaneous fat area (SFA) began to increase from the first postoperative year

prominently to -19.3% and -12.4%, whereas the rate of change in SMI of the TG and PG patients in the younger group returned to -2.9% and -4.4%, respectively.

Discussion

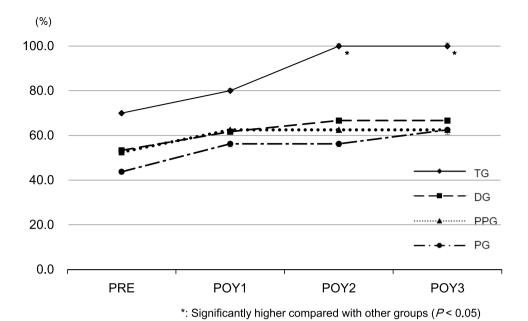
We reviewed the long-term SMI changes after four types of gastrectomy in patients with early gastric cancer and sought to identify the perioperative risk factors. The results of this study revealed annual progressive skeletal muscle loss and exacerbation of the prevalence of sarcopenia. TG and PG were both risk factors for a significant loss of skeletal muscle in postoperative years 1 and 3. The type of gastrectomy,

especially in elderly patients, appeared to have a major impact on the SMI decrease.

Many reports have demonstrated that preoperative sarcopenia is a risk factor for perioperative complications [7, 8], impaired overall survival of patients with gastrointestinal and hepatopancreatobiliary malignancies [10], and increased postoperative morbidity [6]; yet only a few reports have described the progression of postoperative sarcopenia. It was reported that 57.7% of patients have postoperative sarcopenia after gastrectomy [17], and that 25.5% of patients showed significant skeletal muscle loss after TG [11]. Kugimiya et al. also reported that skeletal muscle loss after gastrectomy is an independent predictor of poor prognosis [18]. However, these reports were limited to TG and/or DG.



Fig. 4 All patients in the total gastrectomy group had sarcopenia, evaluated by the skeletal muscle index (SMI) in postoperative years 2 and 3. Definition of sarcopenia: Male SMI < 52.4 cm²/m², Female SMI < 38.5 cm²/m²



SMI, skeletal muscle index; TG, total gastrectomy; PG, proximal gastrectomy; DG, distal gastrectomy; PPG, pylorus-preserving gastrectomy; PRE, preoperative year; POY, postoperative year

To our knowledge, the present study is the first to examine long-term skeletal muscle loss after each of the four major types of gastrectomy.

Limited resection approaches, such as PPG and PG, are now being used more frequently to treat early gastric cancer in the middle-third and upper-third sections of the stomach, for their potential to maintain postoperative function and quality of life. PG resulted in better outcomes than TG in relation to postoperative morbidity and nutrition [19, 20] and body weight loss [21, 22]. Moreover, PPG is superior to DG because it ameliorates post-gastrectomy syndrome and maintains quality of life [23]. However, our study indicates that neither PG nor PPG has any advantage over TG or DG for skeletal muscle loss management. Further studies on large numbers of gastrectomy patients, with due consideration of the reconstructive techniques used and other relevant factors, are needed.

In this study, the SMI in the PG and TG groups continued to decrease throughout the observation period, in contrast to the pattern exhibited by the DG and PPG groups, in which the SMI decreased mainly in postoperative years 1 and 2. Recent studies show the nutritional benefit of subtotal gastrectomy, in which part of the stomach is left, even if it is small, over TG and PG [24, 25]. Since our DG group had less SMI reduction than the TG and PG groups, we consider subtotal gastrectomy the procedure of choice for upper gastric cancer.

Heneghan et al. reported that the prevalence of sarcopenia was 81.4%, 18 to 24 months after esophagectomy and total

or subtotal gastrectomy [26]. This does not conflict with our findings, which showed that all patients who underwent TG had sarcopenia from 2 years postoperatively onward. Although the PG patients had the same rate of SMI reduction as the TG patients, the prevalence of sarcopenia in the TG patients might have been higher due to lower preoperative SMI values or variations in skeletal muscle loss in the PG patients. The postoperative SMI after every type of gastrectomy decreased each year, while the SFA and VFA began to increase from 2 years onward. A previous study noted that body weight decreased 6 months postoperatively, but then gradually increased [21]. The present study found that the SMI decreased over 3 years postoperatively, which is a different trend from the change in body weight. This finding suggests that bodyweight recovery reflects fat tissue recovery.

The elderly patients in the TG and the PG groups had severe skeletal muscle loss, which is associated with a worse prognosis [18, 27]. Takeshita and Liang et al. reported that elderly patients had a poorer overall survival rate than younger patients, although the disease-specific survival rates of the two groups were similar [28, 29]. TG and PG, which are selected based on the tumor location, cannot be avoided when performing curative gastrectomy. To improve the prognosis of elderly patients after TG and PG, we must evaluate the efficacy of surgical techniques and postoperative supportive care, such as oral nutritional support with training intervention. It is thought that gastrectomy-induced malnutrition can be compensated through nutritional support. The

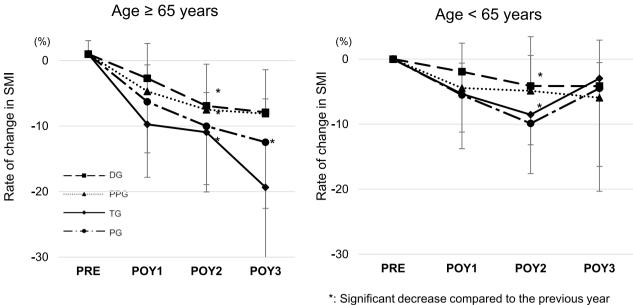


Table 2 Univariate and multivariate analyses of clinical factors for significant loss of skeletal muscle in postoperative years 1, 2, and 3

Factor	POY1	1.			POY2			POY3		
	и	Number of patients in	Univariate analysis	Multivariate analysis		Univariate analysis	Multivariate analysis	Number of patients in	Univariate analysis	Multivariate analysis
		SSML	Odds (95% P value CI)	Odds (95% P value CI)	SSML	Odds (95% P value CI)	Odds (95% P value CI)	SSML	Odds (95% P value CI)	Odds (95% P value CI)
Sex										
Male	94	19	10.38 (1.34–0.0072 80.40)	5.19 (0.65–0.0653 98.06)	37	2.38 0.0502 (1.02–5.54)		38	1.22 0.7044 (0.57–2.59)	
Female	42	1			6			15		
Age (vears)										
<65	57	3			13			16		
> 65	79	17	4.935 (1.37–0.0126 17.75)	3.38 (0.98–0.0527 15.73)	33	2.42 0.0274 (1.13–5.20)	2.42 0.0019 (1.51–5.34)	37	2.25 0.0329 (1.09–4.67)	2.07 0.0503 (0.99–4.42)
Preopera- tive sar- conenia										
Ī	62	6	1.02 0.2100 (0.39–2.66)		22	0.87 0.7199 (0.42–1.77)		28	0.61 0.2100 (0.30–1.24)	
+ (74	11			24			25		
<u>_</u>	9	2	3.11 (0.53-0.2142 18.24)		4	4.19 (0.73- 0.1793 23.79)		4	3.30 (0.58- 0.2081 18.72)	
Π≥	130	18			42			49		
Type of surgery										
DG PPG	100	8			29			33		
TG PG	36	12	5.75 (2.11–0.0006 15.64)	4.08 (1.44–0.0078 12.05)	17	2.19 0.0641 (1.00–4.79)		20	2.53 0.0275 (1.16–5.52)	2.32 0.0358 (1.06–5.19)
Lymphad- enectomy										
D2	21	1			5			8		
D1, D1+	113	20	0.24 (0.031–0.1984 1.95)		41	1.67 0.32455 (0.65–4.27)		45	1.06 1 (0.47–2.38)	

SMI, skeletal muscle index; TG, total gastrectomy; PG, proximal gastrectomy; DG, distal gastrectomy; PPG, pylorus-preserving gastrectomy; POY, postoperative year; SSML, severe skeletal muscle > 10%; C-D, Clavien-Dindo classification; CI, confidence interval





*: Significant decrease compared to the previous year (*P* < 0.05)

		Ag	ge ≥ 65 years	<u>.</u>		Ag	ge < 65 years	1
	N	POY1	POY2	POY3	N	POY1	POY2	POY3
DG	38	-2.7 (±6.2)	-6.9 (±9.1)	-7.8 (±10.1)	22	-1.9 (±4.3)	-4.1 (±7.5)	-4.1 (±7.0)
PPG	16	-4.6 (±7.3)	-7.5 (±6.9)	-8.0 (±6.6)	24	-4.4 (±3.8)	-4.8 (±5.4)	-5.9 (±5.3)
TG	16	-9.7 (±8.0)	-10.9 (±9.1)	-19.3 (±13.1)	4	-5.3 (±5.8)	-8.5 (±4.6)	-2.9 (±17.3)
PG	9	-6.2 (±7.8)	-10.0 (±8.9)	-12.4 (±10.1)	7	-5.5 (±8.2)	-9.8 (±7.6)	-4.4 (±12.0)

SMI, skeletal muscle index; TG, total gastrectomy; PG, proximal gastrectomy; DG, distal gastrectomy; PPG, pylorus-preserving gastrectomy; PRE, preoperative year; POY, postoperative year

Fig. 5 The skeletal muscle index (SMI) in the proximal gastrectomy and total gastrectomy groups decreased more in the elderly patients than in the younger patients

estimates of the amount of such support needed are based on body weight [30, 31]. We reviewed our patients' CT scans, but records of their body weight were unreliable. In a study on upper-third gastric cancer, Yoo et al. reported that food intake and body weight decreased for 6 months after TG or PG, but then increased gradually thereafter [21]. In our study, the SMI continued to decrease in all groups from 1 to 3 years postoperatively, while the SFA and VFA began to increase from 2 years onward. The apparent discrepancy between food intake and fat tissue changes and the persistence of the SMI decrease suggests that skeletal muscle loss is related to additional factors. Hatao et al. reported that oral nutritional supplements after TG diminish postoperative weight loss but not skeletal muscle loss. Interventions such as progressive resistance training, where participants

exercise against an increasing load [30, 31], offer potential protection against skeletal muscle loss after gastrectomy and should be evaluated in future studies.

This study has several limitations. First, it was a single-center retrospective observational study with a small sample size, especially for the TG group (20 patients) and PG group (16 patients). Second, it aimed to evaluate skeletal muscle loss but could not assess sufficient data such as albumin and pre-albumin, body weight, or the amount of food intake during the perioperative period. Third, we estimated skeletal muscle only from the CT images, which cannot always specifically diagnose sarcopenia, defined as low muscle mass plus reduced muscle function. The literature contains many studies of CT-based assessment of muscle mass and studies that measure the total cross-sectional area of muscle mass



use distinct sex-specific cut-off values [32–37]. The cut-off values for sarcopenia, based on the study of sarcopenic obesity in patients with solid tumors by Prado et al. [14] from a Canadian institution, might overestimate those values for Eastern patients. Therefore, we should consider appropriate cut-off values for different ethnic groups. Prospective and larger-scale studies that include muscle-function evaluation and nutritional data are needed to establish the prevalence of postoperative sarcopenia and its influence on the overall survival of gastric cancer patients.

The findings of this long-term retrospective study of CT images showed that all patients who underwent TG had sarcopenia from 2 years postoperatively onward. TG and PG were risk factors for long-term severe skeletal muscle loss, especially in the elderly.

Declarations

Conflict of interest Yasuyuki Seto received research grants from Taiho Pharmaceutical Co., Ltd and Chugai Pharma Manufacturing CO., Ltd. Asami Okamoto, Susumu Aikou, Ryohei Iwata, Shuichiro Oya, Koichiro Kawasaki, Yasuhiro Okumura, Kotaro Wakamatsu, Masayuki Takegami, Koichi Yagi, Masato Nishida, Hiroharu Yamashita, and Sachiyo Nomura have no conflicts of interest to declare.

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