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## **ORIGINAL ARTICLE**

# Clinicopathologic characteristics of serosa-positive gastric carcinoma in elderly patients

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Purpose: The relationship between the prognosis and the age of patients with gastric carcinoma is controversial. This study examined the clinicopathologic features of elderly gastric carcinoma patients with serosal invasion. Methods: We reviewed the hospital records of 136 elderly gastric carcinoma patients with serosal invasion retrospectively to compare the clinicopathologic findings in the elderly (aged > 70 years) and young (aged < 36 years). Results: The 5-year survival rates of elderly and young patients with curative resection did not differ statistically (33.9% vs. 43.3%; P = 0.318). Multivariate analysis showed that two factors were independent, statistically significant parameters associated with survival: histologic type (risk ratio, 1.805; 95% confidence interval [CI], 1.041 to 3.132; P < 0.05) and operative curability (risk ratio, 2.506; 95% CI, 1.371 to 4.581; P < 0.01). Conclusion: This study demonstrated that elderly gastric carcinoma patients with serosal invasion do not have a worse prognosis than young patients. The important prognostic factor was whether the patients underwent curative resection.

Key Words: Gastric carcinoma, Serosal invasion, Elderly, Prognosis, Age

### **INTRODUCTION**

Although the incidence of gastric carcinoma is declining in the general population [1,2], its incidence in the elderly is increasing [3,4]. In conjunction with recent increases in life expectancy, more of these patients are undergoing surgery for gastric carcinoma than in the past. Despite advances in operative techniques [5,6], the prognosis of patients with gastric carcinoma invading the serosa remains poor [7,8].

Since the incidence of gastric carcinoma in the elderly is also increasing, we are interested in the clinicopathologic features and prognostic factors that affect the survival rate of elderly gastric carcinoma patients with serosal invasion. This study analyzed the clinicopathologic features of gastric carcinoma patients with serosal invasion older than 70 years and compared them with young patients.

## **METHODS**

#### Patients and specimens

From 1991 to 2004, 2,032 patients with gastric carcinoma

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were admitted to the Division of Gastroenterologic Surgery. Of these, 136 were in the elderly group (defined as older than 70 years of age). All patients had a primary adenocarcinoma of the stomach and had no evidence of any other malignancy. The clinicopathologic features of these elderly gastric carcinoma patients with serosal invasion were reviewed retrospectively. Information on each patient's age, sex, extent of lymph node dissection, operative curability, type of surgery, tumor size, tumor location, Borrmann type, histologic type, nodal involvement, hepatic metastasis, peritoneal dissemination, stage at the initial diagnosis, and survival rate was obtained from the hospital records. A histological evaluation was performed according to the Japanese General Rules for Gastric Cancer Study in Surgery and Pathology [9]. Curative resection was defined as all gross disease removed as judged by the surgeon at operation.

#### Statistical analysis

Statistical analysis was performed using the SPSS ver. 12.0 (SPSS Inc., Chicago, IL, USA). The survival rates of the patients were calculated using the Kaplan-Meier method and the relative prognostic importance of the parameters was investigated using the Cox proportional hazards model. The chi-square test was used to evaluate the stat-

Table 1. Clinical features of elderly gastric carcinoma patients with serosal invasion

Variable	Age $> 70 \text{ yr}$ (n = 136)	Age < 36 yr (n = 99)	P-value
Age (yr)	$74.2 \pm 3.4$	$30.8 \pm 4.1$	< 0.001
Gender			< 0.001
Male	91 (66.9)	41 (41.4)	
Female	45 (33.1)	58 (58.6)	
Extent of lymph			0.194
node dissection			
D1	28 (20.6)	19 (19.2)	
≥D2	108 (79.4)	80 (80.8)	
Curability			0.813
Curative	94 (69.1)	70 (70.7)	
Non-curative	42 (30.9)	29 (29.3)	
Operative type			0.014
Total	35 (25.7)	50 (50.5)	
Subtotal	91 (66.9)	44 (44.4)	
Others	10 (7.4)	5 (5.1)	

Values are presented as mean  $\pm$  SD or number (%).

istical significance of differences, and P-values less than 0.05 were considered significant.

#### **RESULTS**

Of the 2,032 patients, 136 aged > 70 years were classified as elderly patients. There were 91 males and 45 females; the gender ratio was 2.02:1. The age of the patients at the time of the initial diagnosis ranged from 71 to 84 years old, with a mean age of 74.2 years old. Subtotal gastrectomy was the procedure most frequently performed (66.9% of cases) in elderly patients, but total gastrectomy was frequently performed in the younger group (50.5% vs. 25.7%; P < 0.05) (Table 1).

Table 2. Histopathologic features of elderly gastric carcinoma patients with serosal invasion

Variable	Age > 70  yr $(n = 136)$	Age < 36  yr $(n = 99)$	P-value
Tumor size (cm)	5.09 ± 3.21	6.01 ± 3.15	0.488
Location			0.006
Upper	14 (10.3)	28 (28.3)	
Middle	41 (30.1)	30 (30.3)	
Lower	81 (59.6)	41 (41.4)	
Borrmann type	` /	, ,	0.015
1	6 (4.4)	3 (3.0)	
2	22 (16.2)	8 (8.1)	
3	92 (67.6)	66 (66.7)	
4	16 (11.8)	22 (22.2)	
Nodal involvement	,	,	0.774
N (-)	37 (27.2)	25 (25.3)	
N (+)	99 (72.8)	74 (74.7)	
Hepatic metastasis	,	,	0.618
H(-)	127 (93.4)	94 (94.9)	
H(+)	9 (6.6)	5 (5.1)	
Peritoneal dissemination	, ,	` ,	0.482
P(-)	107 (78.7)	74 (74.7)	
P(+)	29 (21.3)	25 (25.3)	
Stage	,	,	0.894
2	24 (17.6)	19 (19.2)	
3	62 (45.6)	47 (47.5)	
4	50 (36.8)	33 (33.3)	
Histologic type	,	,	< 0.001
Well-differentiated	15 (11.0)	7 (7.1)	
Moderately differentiated		11 (11.1)	
Poorly differentiated	72 (52.9)	70 (70.7)	
Mucinous	12 (8.8)	2 (2.0)	
Signet ring cell	2 (1.5)	9 (9.1)	

Values are presented as mean ± SD or number (%).

Table 2 summarizes the histopathological features of elderly gastric carcinoma patients with serosal invasion. The lower third of the stomach was the most common site of gastric carcinoma in both groups, and the upper third was more frequently involved in the young than in the elderly (28.3% vs. 10.3%; P < 0.01). Significantly more old patients had a well- or moderately differentiated histology and more young patients had a poorly differentiated histology and signet ring cell carcinoma (P < 0.001). Borrmann type IV lesions were more common in younger patients than in the elderly group (22.2% vs. 11.8%; P < 0.05). Multivariate analysis showed that two factors were independent, statistically significant parameters associated with survival: histologic type (risk ratio, 1.805; 95% confidence interval [CI], 1.041 to 3.132; P < 0.05) and oper-

Table 3. Elderly gastric carcinoma with serosal invasion: multivariate analysis

Variable	Risk ratio	95% CI	P-value
Gender (male vs. female)	0.683	0.370-1.259	0.222
Location (upper vs. distal)	0.839	0.338-2.081	0.705
Tumor size (mm) ( $<$ 50 vs. $>$ 50)	1.677	0.917-3.068	0.093
Histologic type (differentiated	1.805	1.041-3.132	0.036
vs. undifferentiated)			
Curability (curative vs.	2.506	1.371-4.581	0.003
non-curative)			
Lymph node metastasis	1.863	0.945-3.675	0.073
(negative vs. positive)			

CI, confidence interval.

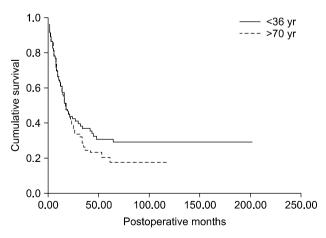


Fig. 1. Survival curves of the young and elderly groups. The 5-year survival rates of young and elderly patients did not differ statistically.

ative curability (risk ratio, 2.506; 95% CI, 1.371 to 4.581; P  $\,<$  0.01) (Table 3).

The 5-year survival rates of the young and elderly patients with serosal invasion did not differ statistically (37.5% vs. 27.8%; P = 0.295) (Fig. 1). The 5-year survival rates of young and elderly patients with curative resection did not differ statistically (43.3% vs. 33.9%; P = 0.318) (Fig. 2). The elderly patients with curatively resected gastric carcinoma had a better survival rate than the elderly patients with non-curatively resected gastric carcinoma (52.1% vs. 7.8%; P < 0.001) (Fig. 3).

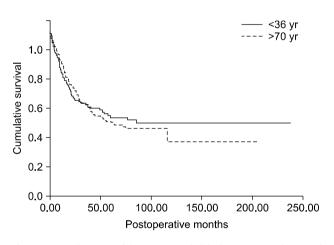


Fig. 2. Survival curves of the young and elderly groups with serosal invasion; curative resection patients.

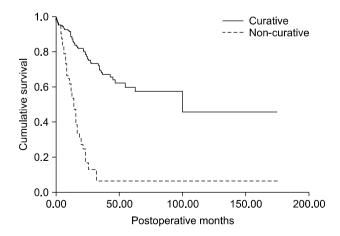


Fig. 3. Survival curves of elderly patients according to resectability. Survival curves of elderly patients with curative resection had better than those of patients with non-curative resection.

#### **DISCUSSION**

Gastric carcinoma is usually a disease of the aged, and patients have a mean age over 60 years [10,11] and the incidence of gastric carcinoma is increasing in very old patients (>70 years) [2-4]. There is controversy as to whether gastric carcinoma in elderly patients differs from that in young patients. Some authors have reported an inverse relationship between age and prognosis in gastric carcinoma [3,5]. We reviewed patients with serosa-positive gastric carcinoma retrospectively to compare the clinicopathologic features in elderly and young patients.

In the present study, there was a significant difference in the sex ratio between the elderly and young patients. In the elderly patients, there was a higher proportion of male patients (2.02:1 in this study). Several studies have obtained the same results [1,12]. The causes of this sexual imbalance are not yet clear. We proposed that possibly, sex hormones may play a role to gender predilection. Some authors suggested that male patients may have more frequent and prolonged exposure to environmental carcinogens than females, which might explain the male predominance among elderly patients [13]. By contrast, there was a higher proportion of female patients in young patients.

Concerning the tumor location, the incidence in the lower third of the stomach was higher in elderly patients than in young patients. It was reported that the same results [14]. Some investigators found that the location of tumor had a significant impact on survival [12]. In contrast to their result, we found that tumor location was not a significant prognostic factor in elderly gastric carcinoma patient with serosal invasion by multivariate analysis.

By histologic type, we found that significantly more elderly patients had a well or moderately differentiated histology, and more young patients had a poorly differentiated histology (P < 0.001). Other studies have reported similar results [1,3,4,12,14], and this may due to lack of carcinoma cell differentiation in the young patients. It was reported that the histologic type of early gastric carcinoma in elderly patients; 45.5% of early gastric carcinoma was well-differentiated adenocarcinoma [15]. In contrast to elderly patients, the higher incidence of poorly differentiated adenocarcinoma in young patients

found in this study is consistent with the literature. In this study, histologic type was one of significant prognostic indicator in multivariate analysis.

Some investigators found that the presence of lymph node metastasis plays an important role in predicting the prognosis in patients with gastric carcinoma [16,17]. In the current study, there were no significant differences in either lymph node invasion between the two groups and lymph node metastasis had no affect on prognosis of elderly gastric carcinoma patients with serosal invasion. Some authors demonstrated that serosal invasion is another important prognostic factor in addition to lymph node metastasis [16,18]. However, our previous results found that serosal invasion did not emerge as an independent, statistically significant prognostic parameter for nodepositive gastric carcinoma patients with curative resection who showed long-term survival [4].

Surgery is the only potentially curative modality for localized gastric carcinoma. In accordance with most literature reports [11,14,19], curative resection offered the only chance of long-term survival. It was reported that when the tumor was curatively resected, the prognosis was favorable in patients with gastric carcinoma patients [19]. Nevertheless, some authors concluded that the extent of surgery should be considered, especially as total gastrectomy and extended node dissection are associated with higher operative mortalities [20]. Many investigators have reported a low curative resection rate in elderly patients with gastric carcinoma [12,21,22]. In our series, however, the curative resection rate (69.1%) in the elderly group was much higher than previously reported in Western countries. It was reported that surgery should not be avoided based solely on the age of patient [23]. We also agree with their recommendation and perform gastrectomy with D2 lymph node dissection in elderly patients with advanced gastric carcinoma who have no medical illness, such as cardiovascular or respiratory problems to achieve curative resection. In contrary, some investigators recommended that the less extensive gastric surgery for the very old patients with gastric carcinoma to improve quality of life [24].

It was reported that there was a trend to more cases with stage IV in the group of young patients than in older pa-

tients (76% vs. 64%), although the difference was not statistically significant [12]. In contrast to their result, we found that stage IV gastric carcinoma was more in the group of elderly patients than in young patients (36.8% vs. 33.3%), and also the difference was not statistically significant in this study. This was due to delay in diagnosis in elderly gastric carcinoma patients.

Although the young patients presented aggressive histologic patterns, it was reported that there was no statistical difference in survival rates between the elderly and young gastric carcinoma patients [1]. In this study, the 5-year survival rates of the elderly and young patients did not differ statistically (46.5% vs. 52.8%). These findings suggest that elderly patients with gastric carcinoma can tolerate radical treatment well. The elderly patients with curatively resected gastric carcinoma had a better survival rate than the elderly patients with non-curatively resected gastric carcinoma (68.1% vs. 6.5%). By contrast, others [3,20,24] reported that the survival rates of the elderly were worse, both overall and after curative resection, than those of younger patients. Delay in diagnosis and a more advanced stage of gastric carcinoma in elderly patients have been suggested as possible causes. In a few reports, however, the prognosis of elderly patients who underwent curative resection was the same as that of young patients. Many investigators [4,22,25] also reported similar survival rates in the two age groups when the same tumor stages were compared.

In conclusion, elderly patients represent more differentiated histology compared to young patients. This study demonstrated that there was no significant differences in outcome of gastric carcinoma with serosal invasion between elderly and young patients. The prognosis of serosa-positve gastric carcinoma is not related to the patients' age.

## **CONFLICTS OF INTEREST**

No potential conflict of interest relevant to this article was reported.

#### REFERENCES

- Wang JY, Hsieh JS, Huang CJ, Huang YS, Huang TJ. Clinicopathologic study of advanced gastric cancer without serosal invasion in young and old patients. J Surg Oncol 1996;63:36-40.
- 2. Roder DM. The epidemiology of gastric cancer. Gastric Cancer 2002;5 Suppl 1:5-11.
- 3. Kubota H, Kotoh T, Dhar DK, Masunaga R, Tachibana M, Tabara H, et al. Gastric resection in the aged (> or = 80 years) with gastric carcinoma: a multivariate analysis of prognostic factors. Aust N Z J Surg 2000;70:254-7.
- 4. Kim DY, Joo JK, Park YK, Ryu SY, Kim YJ, Kim SK. Predictors of long-term survival in node-positive gastric carcinoma patients with curative resection. Langenbecks Arch Surg 2007;392:131-4.
- 5. Morel P, Egeli RA, Wachtl S, Rohner A. Results of operative treatment of gastrointestinal tract tumors in patients over 80 years of age. Arch Surg 1989;124:662-4.
- 6. Schöön IM, Arvidsson S. Surgery in patients aged 80 years and over. A retrospective comparative study from 1981 and 1987. Eur J Surg 1991;157:251-5.
- 7. Bizer LS. Adenocarcinoma of the stomach: current results of treatment. Cancer 1983;51:743-5.
- 8. Korenaga D, Okamura T, Baba H, Saito A, Sugimachi K. Results of resection of gastric cancer extending to adjacent organs. Br J Surg 1988;75:12-5.
- 9. Japanese Gastric Cancer Association. Japanese Classification of Gastric Carcinoma, 2nd English Edition. Gastric Cancer 1998;1:10-24.
- 10. Grabiec J, Owen DA. Carcinoma of the stomach in young persons. Cancer 1985;56:388-96.
- 11. Mitsudomi T, Matsusaka T, Wakasugi K, Takenaka M, Kume K, Fujinaga Y, et al. A clinicopathological study of gastric cancer with special reference to age of the patients: an analysis of 1,630 cases. World J Surg 1989;13:225-30.
- 12. Medina-Franco H, Heslin MJ, Cortes-Gonzalez R. Clinico-pathological characteristics of gastric carcinoma in young and elderly patients: a comparative study. Ann Surg Oncol 2000;7:515-9.
- 13. Ershler WB, Longo DL. The biology of aging: the current research agenda. Cancer 1997;80:1284-93.
- 14. Fujimoto S, Takahashi M, Ohkubo H, Mutou T, Kure M, Masaoka H, et al. Comparative clinicopathologic features of early gastric cancer in young and older patients. Surgery 1994;115:516-20.
- 15. Nakamura T, Yao T, Niho Y, Tsuneyoshi M. A clinicopathological study in young patients with gastric carcinoma. J Surg Oncol 1999;71:214-9.
- 16. Adachi Y, Mori M, Maehara Y, Kitano S, Sugimachi K. Prognostic factors of node-negative gastric carcinoma: univariate and multivariate analyses. J Am Coll Surg 1997; 184:373-7.
- 17. Ikeguchi M, Oka S, Gomyo Y, Tsujitani S, Maeta M, Kaibara N. Prognostic benefit of extended radical lymphadenectomy for patients with gastric cancer. Anticancer Res 2000;20(2B):1285-9.

- 18. Mori M, Sugimachi K. Clinicopathologic studies of gastric carcinoma. Semin Surg Oncol 1990;6:19-27.
- 19. Maehara Y, Emi Y, Tomisaki S, Oshiro T, Kakeji Y, Ichiyoshi Y, et al. Age-related characteristics of gastric carcinoma in young and elderly patients. Cancer 1996;77:1774-80.
- 20. Katai H, Sasako M, Sano T, Maruyama K. The outcome of surgical treatment for gastric carcinoma in the elderly. Jpn J Clin Oncol 1998;28:112-5.
- 21. Damhuis RA, Tilanus HW. The influence of age on resection rates and postoperative mortality in 2773 patients with gastric cancer. Eur J Cancer 1995;31A:928-31.
- 22. Winslet MC, Mohsen YM, Powell J, Allum WH, Fielding

- JW. The influence of age on the surgical management of carcinoma of the stomach. Eur J Surg Oncol 1996;22:220-4.
- 23. Otani Y, Kubota T, Kumai K, Ohgami M, Hayashi N, Ishikawa Y, et al. Surgery for gastric carcinoma in patients more than 85 years of age. J Gastroenterol Hepatol 2000; 15:507-11.
- 24. Eguchi T, Fujii M, Takayama T. Mortality for gastric cancer in elderly patients. J Surg Oncol 2003;84:132-6.
- 25. Piso P, Bektas H, Werner U, Becker T, Aselmann H, Schlitt HJ, et al. Comparison between treatment results for gastric cancer in younger and elderly patients. Zentralbl Chir 2002;127:270-4.