ENDOSCOPIC TREATMENT OF EGC INCLUDING OUTCOME AT SMC

IM R3 배주환 IM1 Prof. 이혁

Introduction

- Standard treatment of gastric cancer is surgery.
- However, Endoscopic resection(ER) is an option for selected patients with EGC without known lymph node involvement who meet specific criteria.
- Endoscopic resection is associated with less treatment-related morbidity than gastrectomy and the available data suggest similar outcomes for appropriately selected patients with EGC.

Endoscopic resection

• ER includes endoscopic mucosal resection (EMR) and endoscopic submucosal dissection (ESD).

* EMR



- (A) Simulated dysplastic lesion (red mark depicts dysplasia).
- (B) Isolation of the lesion with submucosal fluid injection.
- (C, D) Snare excision of isolated dysplastic lesion.

Courtesy of Christopher Gostout, MD.

* ESD



EMR vs ESD

• ESD, compared with EMR, had higher en bloc and curative resection rates (OR 13.9 and 3.5, respectively), as well as lower rates of local recurrence (OR 0.09).

Surg Endosc. 2011 Aug;25(8):2666-77. Epub 2011 Mar 18. Gastrointest Endosc. 2012 Oct;76(4):763-70. Epub 2012 Aug 9.

• Patients who underwent ESD had lower recurrence rates than patients who underwent EMR (4 versus 18 percent)

Surg Endosc. 2010;24(11):2842.

Absolute indication of ER

- I. Well and/or moderately differentiated adenocarcinoma confined to the mucosa.
- II. Less than 20 mm in diameter, without ulceration.
- III. Absence of venous or lymphatic invasion.

심평원 ESD 인정기준 (시행일자 2012.4.1) "점막에 국한된 궤양이 없는 2㎝이하의 분화형 조기암"

Expanded indication of ER

Histology	Depth							
	M cancer				SM cancer			
	No ulceration		Ulcerated		SM1	SM2		
	≤20 mm	> 20 mm	≤ 30 mm	> 30 mm	≤ 30 mm	Any size		
Differentiated	Α	B1	B2	D	B3	D		
Undifferentiated	С	D	D	D	D	D		

- A guideline indication for EMR/ESD
- B expanded indications for EMR/ESD
- C consider surgery
- D surgery (gastrectomy + lymph node dissection)

Modified from Soetikno, Kaltenbach, Yeh, Gotoda. JCO 2005;23:4490-4498



Fig.2 Flowchart of clinical procedure for patients with early gastric cancer. *: In patients with early gastric cancer consisting of intestinal-type histology: (1) intramucosal cancer more than 2cm in size without ulcer finding, (2) intramucosal cancer less than 3cm in size with ulcer finding, (3) minute submucosal invasive cancer($\leq 500 \mu$ m) less than 3cm in size. In patients with early gastric cancer consisting of diffuse-type histology. (4) intramucosal cancer less than 2cm in size without ulcer finding.

Chika Kusano, Takuji Gotoda et al. Stomach Intestine 2008;43:73-79

Definition of curative resection

- Curative resection(When all of the following conditions were fulfilled)
 - ① Grossly complete resection (by the endoscopist):
 - ② En-bloc resection
 - ③ Well or moderately differentiated histology
 - ④ Negative resection margin
 - 5 No lymphovascular invasion

And

- I. Tumor size ≤ 2 cm, mucosal cancer, no ulcer in tumor(AI), or
- II. Tumor size > 2 cm, mucosal cancer, no ulcer in tumor, or
- III. Tumor size ≤ 3 cm, mucosal cancer, ulcer in tumor, or
- IV. Tumor size \leq 3 cm, sm1 cancer (submucosal invasion depth < 500 um from muscularis mucosa layer).

Overall survival of patients with EGC who received curative ESD at SMC



Fig. 2 Kaplan–Meier overall survival curve of patients with early gastric cancers meeting absolute indication or expanded indication criteria, and treated with curative endoscopic submucosal dissection (ESD). The 5-year overall survival rates of patients with absolute-indication and expanded-indication cancers were 97.3% and 96.4%, respectively. Fifteen Patients (n = 15) with both absolute-indication and expanded-indication early gastric cancers were included in the expanded-indication group.

Endoscopy 2015 Sep;47(9):784-93.

Metachronous recurrence after curative ESD at SMC



Fig. 3 Metachronous recurrence after curative endoscopic submucosal dissection (ESD) for early gastric cancer: cumulative incidence curve. During the 60-month surveillance period after curative ESD, the cumulative incidence curve showed a nearly linear increase, implying a constant incidence rate for metachronous recurrence.

Endoscopy 2015 Sep;47(9):784-93.

Table 2 Univariate and multivariate analysis of factors associated with metachronous recurrence after curative endoscopic submucosal dissection (ESD) for differentiated-type early gastric cancer.

	Metachronous recurrence ¹		Odds ratio	95 %CI	P value
	None (n=1259)	Present (n = 47)			
Age, mean±SD, y	61.5±9.7	63.1±8.8	1.015	0.983 - 1.047	0.364
Gender, n (%)					0.427
Male	1004 (79.7)	40 (85.1)			
Female	255 (20.3)	7 (14.9)	0.714	0.311-1.640	
Number of lesions, n (%)					0.025
Single	1229 (97.6)	43 (91.5)			
Multiple	30 (2.4)	4 (8.5)	3.691	1.177-11.574	
Tumor site, n (%)					0.238
Antrum/angle	994 (79.0)	34 (72.3)			
Body/fundus/cardia	265 (21.0)	13 (27.7)	1.491	0.768-2.896	
Tumor shape, n (%)					0.683
Elevated	715 (56.8)	28 (59.6)			
Flat or depressed	544 (43.2)	19 (40.4)	0.882	0.482-1.613	
Tumor size, mean ± SD, cm	1.4±0.8	1.3±0.8	0.724	0.409-1.280	0.267
Tumor depth (%)					0.516
Mucosa	1194 (94.8)	45 (95.7)			
sm1 ²	65 (5.2)	2 (4.3)	0.556	0.094-3.274	
Differentiation, n (%)	and a state				0.016
Well differentiated	506 (40.2)	28 (59.6)			
Moderately differentiated	753 (59.8)	19 (40.4)	0.477	0.262-0.869	
Indication, n (%)					0.595
Absolute	994 (79.0)	38 (80.9)			
Expanded	265 (21.0)	9(19.1)	1.406	0.400-4.937	

Endoscopy 2015 Sep;47(9):784-93.

Extragastric recurrence after curative ESD



Prognosis of non-curative endoscopic resection of EGC at SMC



(P = 0.013)

BJS 2015; **102**: 1394–1401

Overall survival



Median duration of follow-up after ER: 60.5 mo(6-141)

BJS 2015; 102: 1394–1401

Results of multivariable Cox proportional hazards analysis to determine predictors of overall survival

		n	Five-year overall survival rate (%)	Hazard ratio	95% Confidence interval	P value
Age	<65 years	140	95.6			
	\geq 65 years	134	87.0	2.048	0.717-5.849	0.181
Gender	Male	205	90.3			
	Female	69	95.0	0.428	0.144-1.273	0.127
Charlson comorbidity index	< 4	157	96.1			
	≥4	117	85.5	1.381	0.525-3.629	0.513
Additional surgery	No treatment	80	84.7			
	Surgery	194	94.3	0.425	0.181-0.998	0.049
Tumor depth	Mucosa or SM1*	76	92.8			
	SM2 or SM3	198	90.9	1.564	0.618-3.962	0.345
Lymphovascular invasion	Negative	138	92.1			
	Positive	136	90.7	1.718	0.763-3.871	0.191
Tumor size	<2 cm	126	94.0			
	\geq 2 cm	148	89.0	1.018	0.457-2.270	0.965

*SM1, submucosal invasion depth < 500 µm from muscularis mucosa layer

Complications

- Perforation : o~5%
- Bleeding : 3~10%, most bleeding occurs within 3days after ESD.
- Post ESD pain
 - There is a article that PPI can reduce moderate to severe pain after ESD (44.9% vs 62.6%)
- Stricture after ESD
 - Subclinical stricture is quite common after ESD for lesions close to the cardia or the pyloric ring

References

- http://endotoday.com
- 이준행, 김재규, 정혜경 et al. 근거 기반 위암 진료 권고안. Korean J Gastroenterol Vol. 63 No. 2, 66-81.
- Park YM, Cho E, Kang HY, Kim JM. The effectiveness and safety of endoscopic submucosal dissection compared with endoscopic mucosal resection for early gastric cancer: a systematic review and metaanalysis. Surg Endosc. 2011 Aug;25(8):2666-77. Epub 2011 Mar 18.
- Lian J, Chen S, Zhang Y, Qiu F. A meta-analysis of endoscopic submucosal dissection and EMR for early gastric cancer. Gastrointest Endosc. 2012 Oct;76(4):763-70. Epub 2012 Aug 9.
- Park JC, Lee SK, Seo JH et al. Predictive factors for local recurrence after endoscopic resection for early gastric cancer: long-term clinical outcome in a single-center experience. Surg Endosc. 2010;24(11):2842.
- Min BH, Kim ER, Kim KM et al. Surveillance strategy based on the incidence and patterns of recurrence after curative endoscopic submucosal dissection for early gastric cancer. Endoscopy. 2015 Sep;47(9):784-93.
- Kim ER, Lee H[,] Min BH et al. Effect of rescue surgery after non-curative endoscopic resection of early gastric cancer. Br J Surg. 2015 Oct;102(11):1394-401

