



# Mide Kanseri Cerrahi Tedavi İlkeleri Ne değişti?

Ali Güner

MD, PhDc, BA, FACS KTÜ Genel Cerrahi ABD Herhangi bir biomedikal firma ile çıkar çatışmam bulunmamaktadır.



# 3 3 WORLD CONGRESS

of International Association of Surgeons, Gastroenterologists and Oncologists

SEPTEMBER 28<sup>TH</sup> - OCTOBER 1<sup>ST</sup>, 2022

#### The General Rules for the Gastric Cancer Study in Surgery and Pathology

Part II. Histological Classification of Gastric Cancer

The Japanese Research Society Committee on Histological Classification of Gastric Cancer

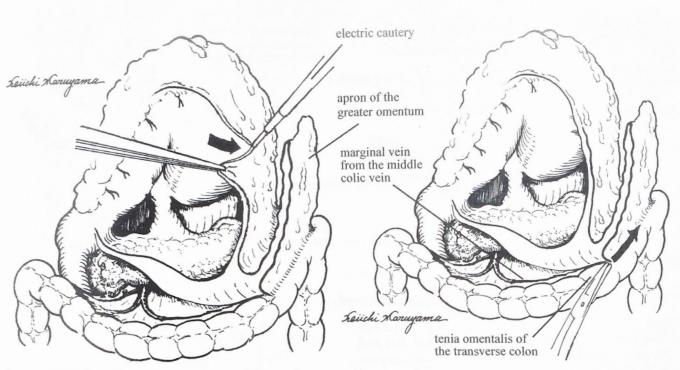


Fig. 9 Division line for the apron-preserving omentectomy

For the cases with no possibility of metastasis in the bursa and peritoneum, namely T1 and T2 tumor, the apron of the greater omentum can be preserved. It is useful to cover the small bowel and to prevent ileus.

Fig.10 Division line for total omento-bursectomy

For the cases with possibility of metastasis in the bursa and peritoneal cavity (T2 and T3 tumor) and negative washing cytology, the total omento-bursectomy is indicated. This procedure will start at separation of the greater omentum from the tenia omentalis.

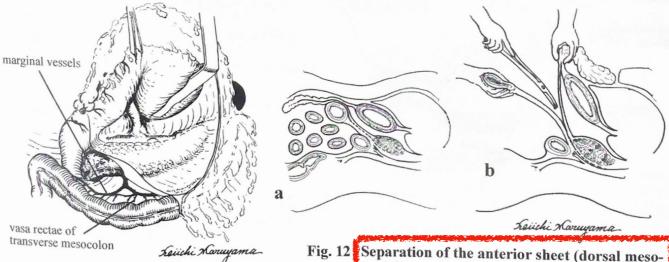


Fig. 11 Right side bursectomy to expose the infrapyloric nodes (No. 6)

Even for the apron preserving cases, this procedure is essential to remove No. 6 nodes. During the separation, the vasa rectae of the transverse colon, marginal vessels, and the middle colic vein are the useful index to trace the correct plane.

Fig. 12 Separation of the anterior sheet (dorsal mesogastrium) and posterior sheet of the mesocolon

Because of different embryonic origin, the anterior and posterior sheets have no vascular connection (a). These two membranes can be separated easily without bleeding. Assistant should tracts the transverse colon downward to make the mesocolon flat. Then the operator lift the greater omentum upward and push the mesocolon and the veins downward by scissors (b).

Kajitani T. Japan J Surg 1981





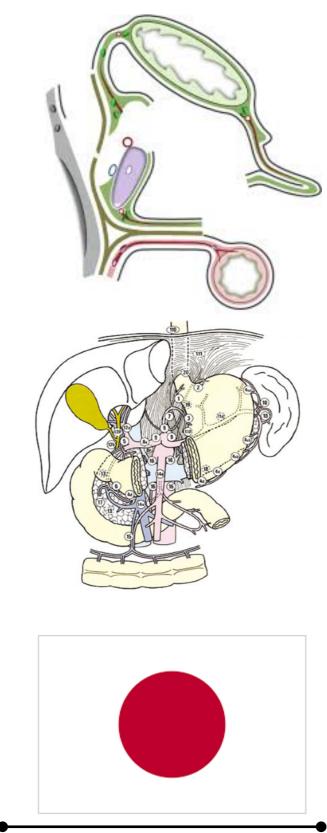




~1880 ~1950 J. Pean ■

L. Rydigier 🚄

T.Billroth



~1950-80

T. Kajitani 🕑

I. Ohashi 💷

D. Jinnai 💷

Lenfadenektomi
Bursektomi
Omentektomi
Splenektomi
Pankreatektomi
MIS
EGJ
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Neoadj/Adj Tx

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**KILAVUZLAR** 

~2000

# **Lenfadenektomi**

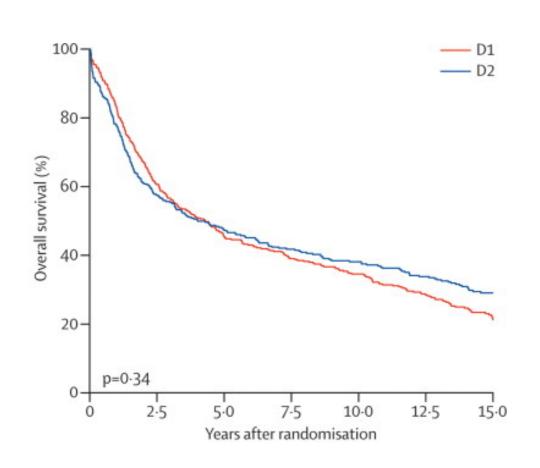


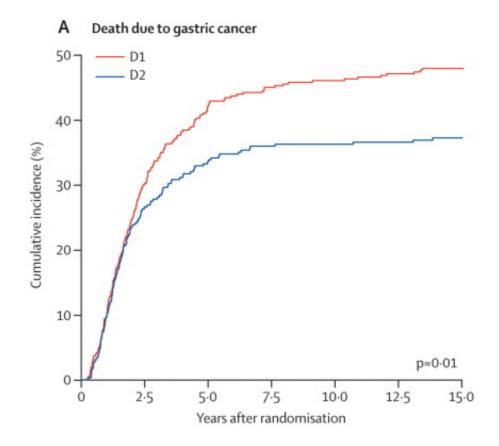
**D1 vs D2** 

D2 vs D2+PALN



# **D1 vs D2**



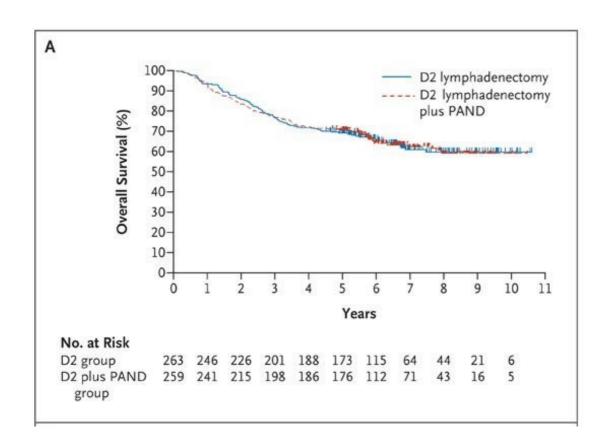


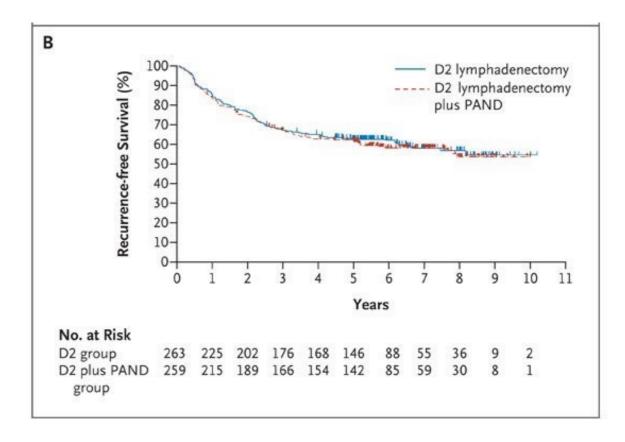
D2 lymphadenectomy is associated with <u>lower locoregional recurrence</u> and <u>gastric-cancer-related death</u> rates than D1 surgery.

D2 lymphadenectomy is the **recommended** surgical approach for patients with resectable (curable) gastric cancer.



# D2 vs D2+PALN





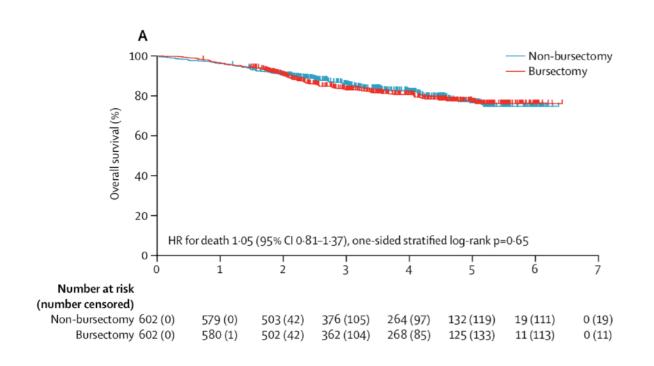
As compared with D2 lymphadenectomy alone, treatment with

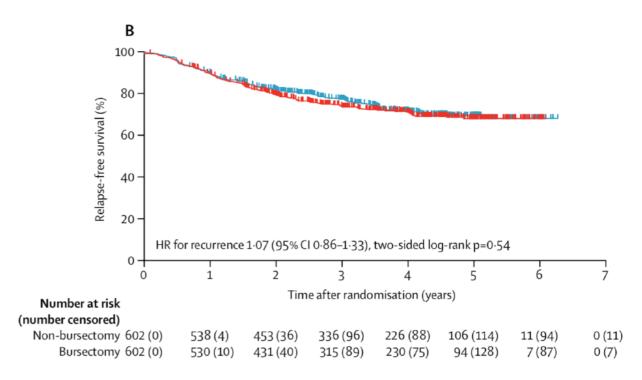
# D2 lymphadenectomy plus PAND does not improve the survival

rate in curable gastric cancer.



# Bursektomili vs Bursektomisiz





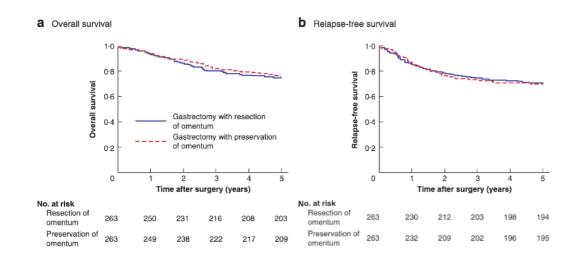
Bursectomy did not provide a survival advantage over non-bursectomy.

D2 dissection with omentectomy alone should be done as a standard surgery for resectable cT3-T4a gastric cancer.

## **Omentektomi**



# **Total vs Parsiyel**



Overall survival and disease recurrence were **comparable** in patients with cT3–4 gastric cancer who underwent GPO or GRO.

Ri M et al. BJS 2020

Recruitment status	Open public recruiting				
Unique ID issued by U MIN	UMIN000036253				
Receipt No.	R000041060				
Scientific Title	Randomized controlled phase III trial to evaluate omentum preserving gastrect omy for patients with advanced gastric cancer (JCOG1711, ROAD-GC)				
Date of disclosure of the study information	2019/03/19				

~1050 hasta Tarih: 2019-<u>2031</u>

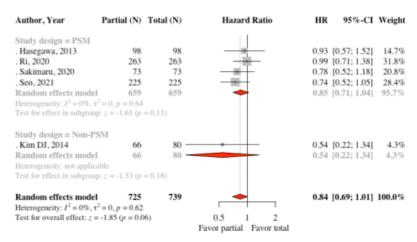


Figure 2. Forest plot comparing disease-free survival between the partial omentectomy group and the total omentectomy group.

Author, Year	Partial (N)	Total (N)	Hazard Ratio	HR	95%-CI	Weight
Study design = Non-PS	SM		i 1			
. Kim JH, 2009	52	122		1.26	[0.51; 3.10]	5.1%
. Kim DJ, 2014	66	80 -		0.61	[0.18; 2.00]	2.9%
Random effects model	118	202		0.97	[0.47; 1.98]	8.0%
Heterogeneity: $I^2 = 0\%$ , $\tau$	$^{2} = 0, p = 0.34$		i I			
Test for effect in subgroup	z = -0.09 (p	= 0.92)				
			i			
Study design = PSM			i			
. Hasegawa, 2013	98	98		0.63	[0.35; 1.13]	12.0%
. Ri, 2020	263	263	-	0.89	[0.61; 1.29]	29.8%
. Sakimaru, 2020	73	73		0.92	[0.59; 1.45]	20.4%
. Seo, 2021	225	225	- 10	0.70	[0.48; 1.01]	29.8%
Random effects model	659	659	<b>~</b>	0.79	[0.64; 0.98]	92.0%
Heterogeneity: $I^2 = 0\%$ , $\tau$	$p^2 = 0, p = 0.60$		: I			
Test for effect in subgroup	z = -2.17 (p	= 0.03)	i			
			i			
Random effects model	777	861	<b>*</b>	0.80	[0.66; 0.98]	100.0%
Heterogeneity: $I^2 = 0\%$ , $\tau$	$^{2} = 0, p = 0.69$	Г				
Test for overall effect: $z =$	-2.11 (p = 0.0)	4) 0.2	2 0.5 1 2	5		
			Favor partial Favor total			

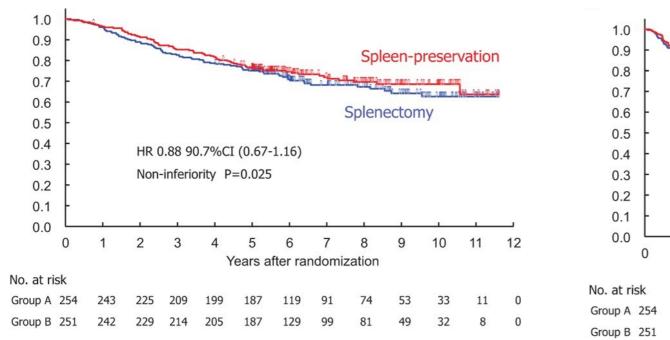
Figure 3. Forest plot comparing overall survival between the partial omentectomy and the total omentectomy group.

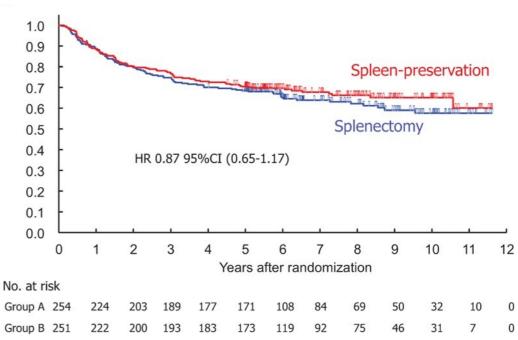
Chai SW et al. Cancers 2021

Recruitment status	Open public recruiting
Date of protocol fixation	2019 Year 01 Month 07 Day
Date of IRB	2019 Year 02 Month 28 Day
Anticipated trial start date	2019 Year 03 Month 19 Day
Last follow-up date	2031 Year 09 Month 19 Day



# Splenektomili vs Splenektomisiz





In total gastrectomy for proximal gastric cancer that <u>does not invade the greater curvature</u>, splenectomy should be avoided as it increases operative morbidity <u>without improving survival</u>.



# Splenektomili vs Splenektomisiz

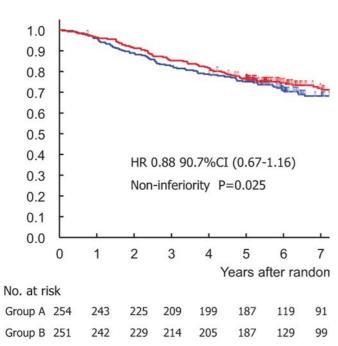


TABLE 1. Eligibility Criteria of the Trial

Inclusion criteria

Before operation

Histologically proven adenocarcinoma

A T2/T3/T4\* tumor located in the upper third of the stomach

Absence of or 3 cm or shorter esophageal invasion by endoscopy and barium fluoroscopy

Absence of tumor invasion of the greater curvature

Not a stump cancer

Not of linitis plastica type (Borrmann type 4)

N0/N1/N2\* and M0 by diagnostic imaging

Sufficient organ functions for total gastrectomy with splenectomy

Age between 20 and 75 years inclusive

No previous chemotherapy or surgery for gastric cancer

Written informed consent from the patient

During operation

Absence of or 3 cm or shorter esophageal invasion, and absence of tumor invasion of the greater curvature by inspection and palpation

T2/T3/T4\* and N0/N1/N2\* by inspection and palpation

Peritoneal lavage cytology is negative for cancer cells

Curative operation is feasible without combined resection of the pancreas and spleen

No gross lymph node metastasis along the splenic artery or splenic hilum

Exclusion criteria

Liver cirrhosis, or portal hypertension

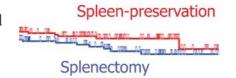
Idiopathic thrombocytopenic purpura, or other diseases for which splenectomy is a treatment option

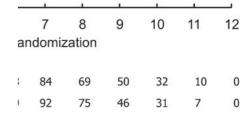
Previous splenectomy

Interstitial pneumonia, pulmonary fibrosis, or extensive pulmonary emphysema

Synchronous or metachronous malignancy in other organs except for carcinoma in situ or intramucosal tumor cured by local resection Pregnant, or possibly pregnant

Psychotic disorders inappropriate for participation in clinical trials





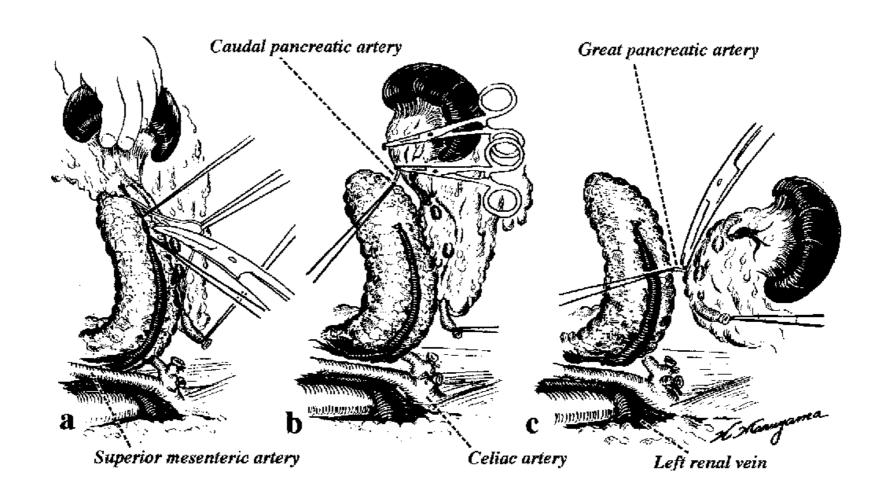
plenectomy should be avoided as it

In total gastrectomy for proximal gastriincreases operative morbidity **witho** 

<sup>\*</sup> Tumors were staged according to both Japanese Classification (13th ed, 1998) and UICC-TNM (6th ed., 2006).

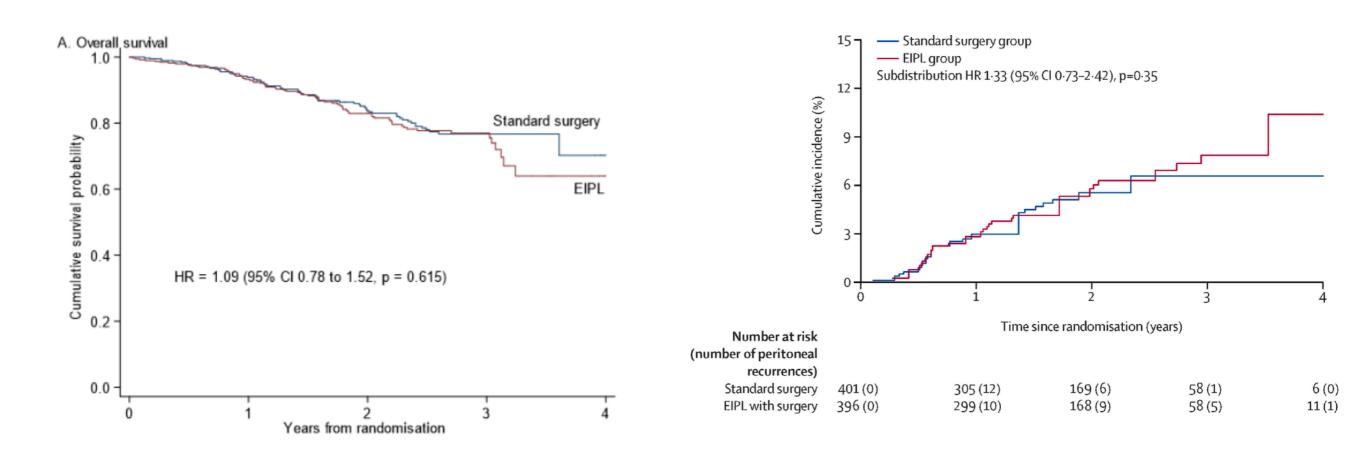


# Pancreas-preserving surgery



## **Extensive Intraperitoneal Lavage**



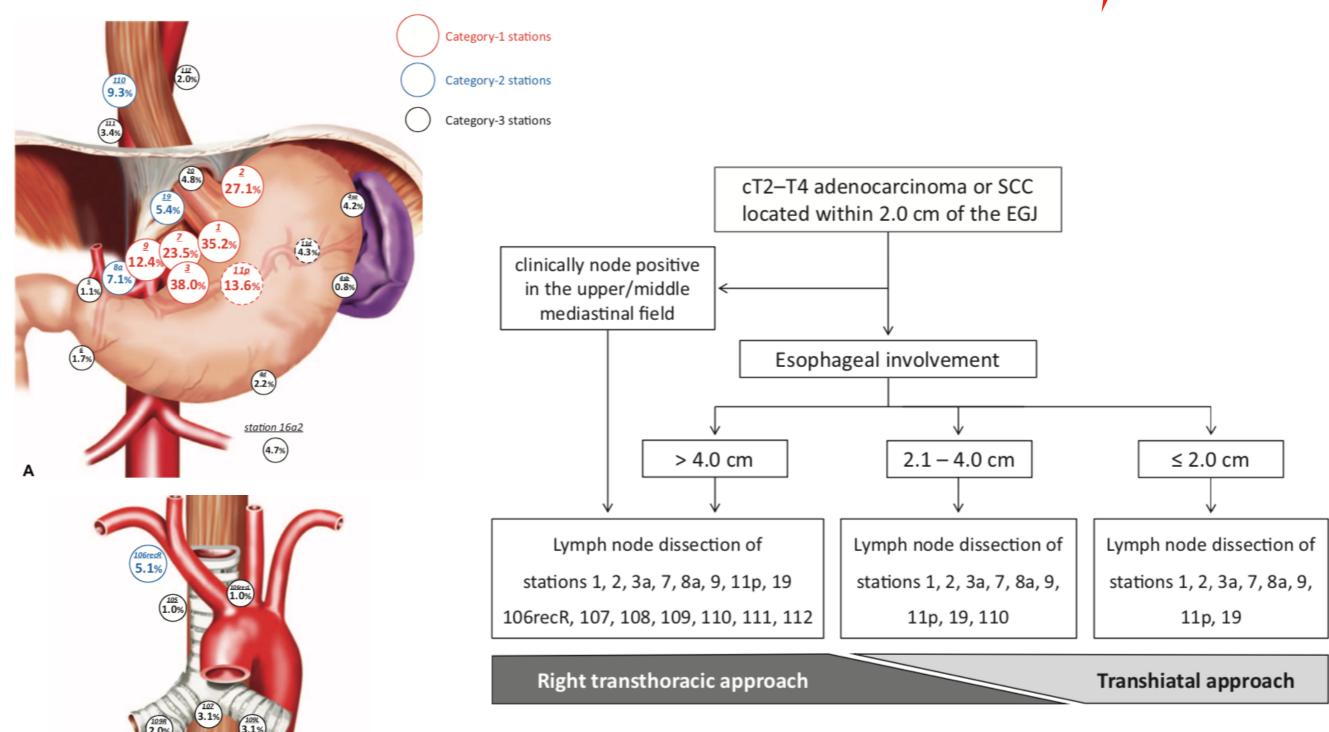


Extensive lavage with saline **did not reduce risk of peritoneal recurrence nor improve survival** after gastric cancer surgery. It may <u>increase</u> risk of postoperative complications

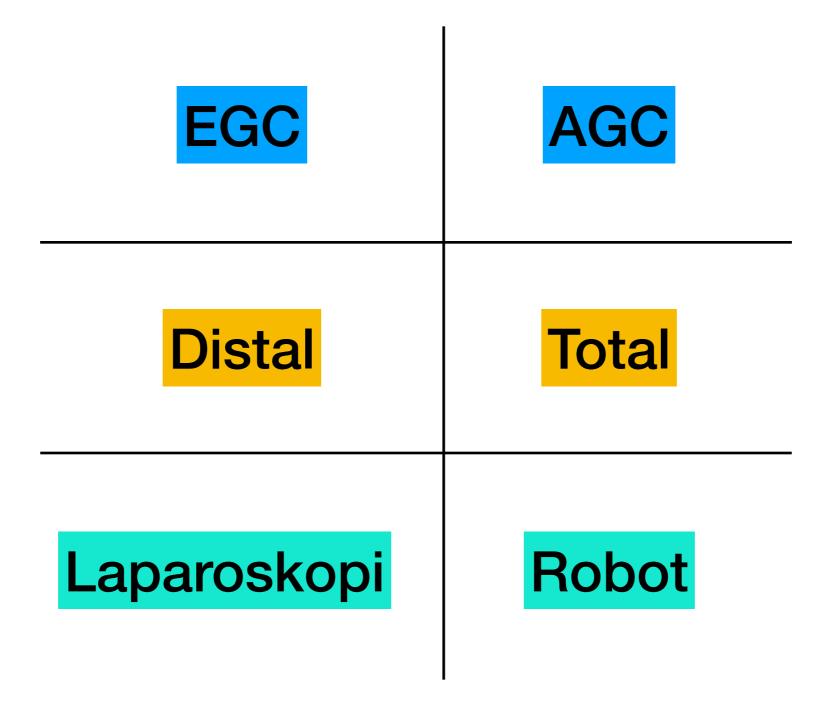
## **EGJ**

В





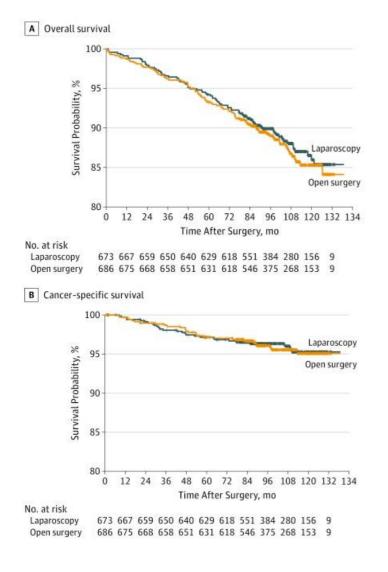
# **Minimal İnvaziv Cerrahi**



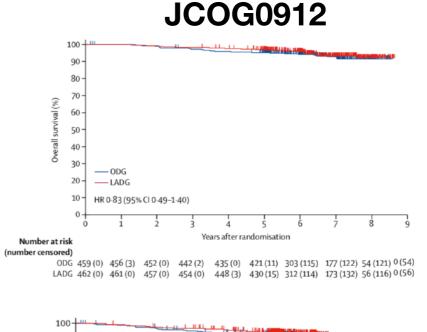
# EGC Distal Laparoskopi



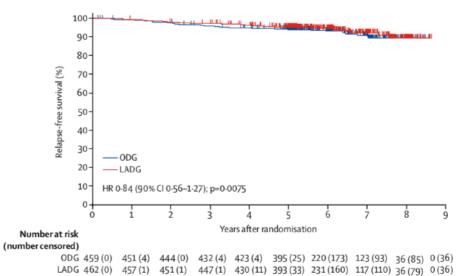
#### KLASS01



5y-OS 94.2% LDG grup 93.3% ODG grup



5y-RFS 95.1% LDG grup 94% ODG grup



# similar overall and cancer-specific

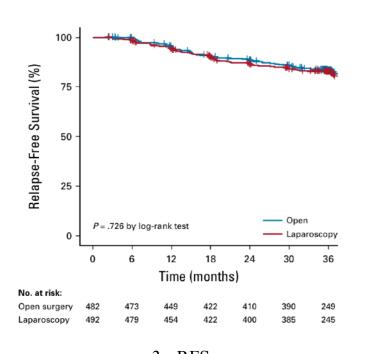
**SURVIVA** rates between patients receiving laparoscopic and open distal gastrectomy

**non-inferiority of LADG** compared with ODG for clinical stage I gastric cancer relapse-free survival, suggesting that LADG should be considered a standard treatment option when performed by experienced surgeons.



# JLSSG0901

#### CLASS01



3y-RFS 80.3% LDG grup 81.3% ODG grup

Laparoscopic distal gastrectomy with D2 lymphadenectomy was

**comparable** to open surgery in terms of relapse-free survival for patients with locally advanced gastric cancer.

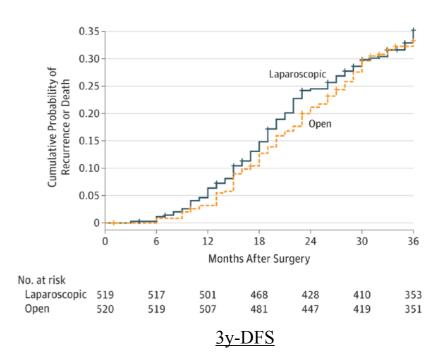
	Progress
Recruitment status	No longer recruiting
Date of B	20' D9' 10. 5 De
Anticipated trial start date	2009 Year 11 Month 01 Day
Last follow-up date	2021 Year 08 Month 01 Day

## The **technical safety** of

LADG with D2 lymph node dissection for locally advanced gastric cancer was demonstrated

Inahi N et al. World J Surg 2015

5-yr RFS Hazard ratio: 0.9556 (95%CI: 0.7226-1.2637 < 1.31) Non-inferiority P=0.0317



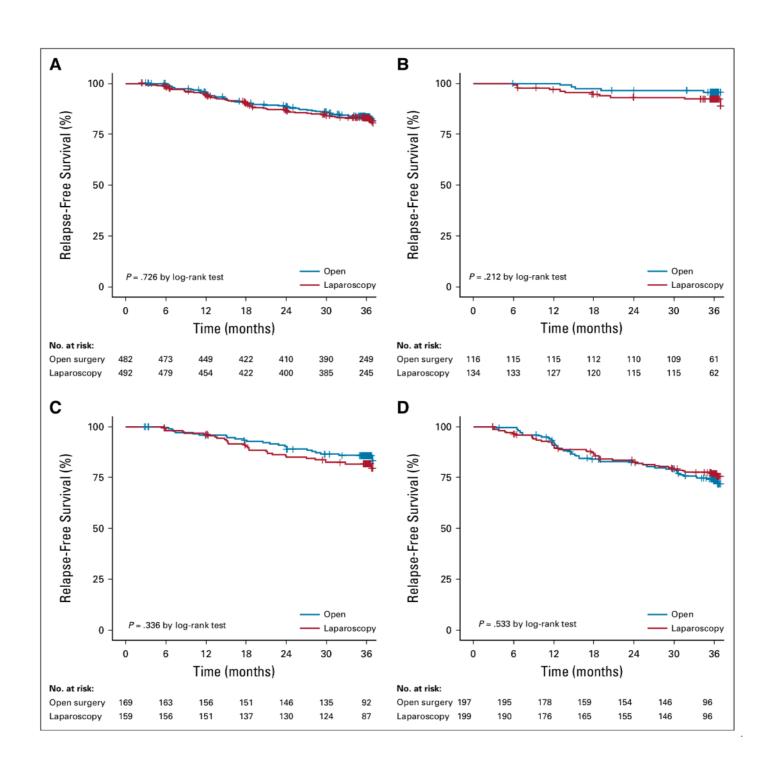
76.5% LDG grup 77.8% ODG grup

- Laparoscopic distal gastrectomy, compared with open distal gastrectomy, did not result in inferior disease-free survival at 3 years.
- Laparoscopic distal gastrectomy with D2 lymphadenectomy performed by experienced surgeons in high-volume specialized institutions resulted in similar 5year overall survival

# AGC Distal Laparoskopi

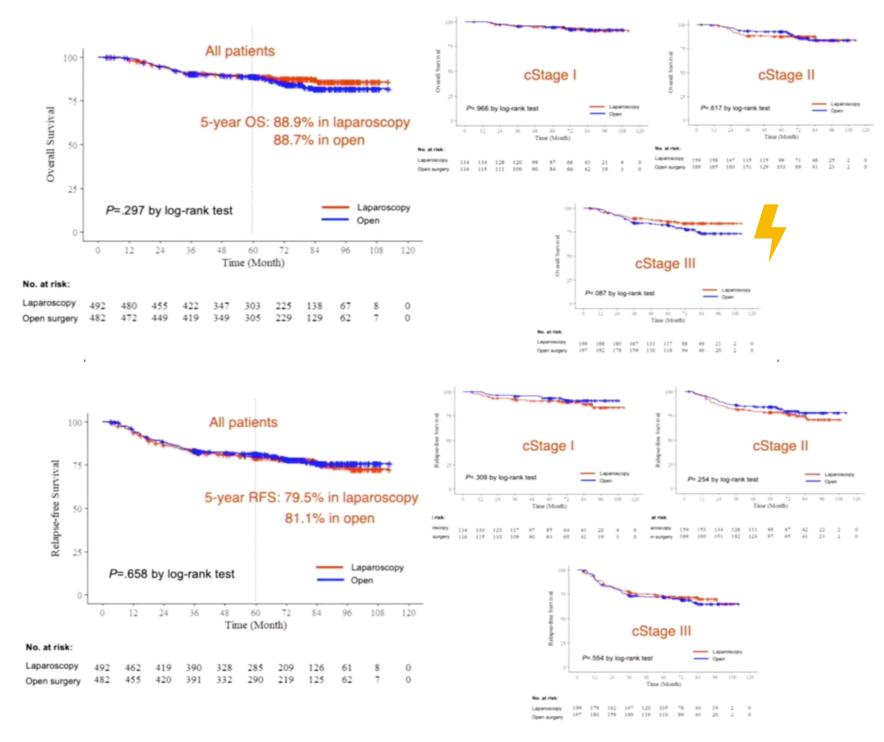


## KLASS02



3y-RFS 80.3% LDG grup 81.3% ODG grup









**JCOG1401** 

CLASS02 (RCT)

Morbidity 20.6% (33/160)

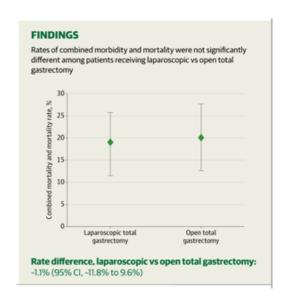
Mortality 0.6% (1/160)

Major complications 9.4% (15/160)

Reoperations 1.9% (3/160)

EJ Anastomosis leakage 2.5% (6/244) Conversion 1.7%

Mortality 0



showed **acceptable** postoperative morbidity and mortality for patients with clinical stage I gastric cancer.

confirmed the **<u>safety</u>** of LATG/LAPG

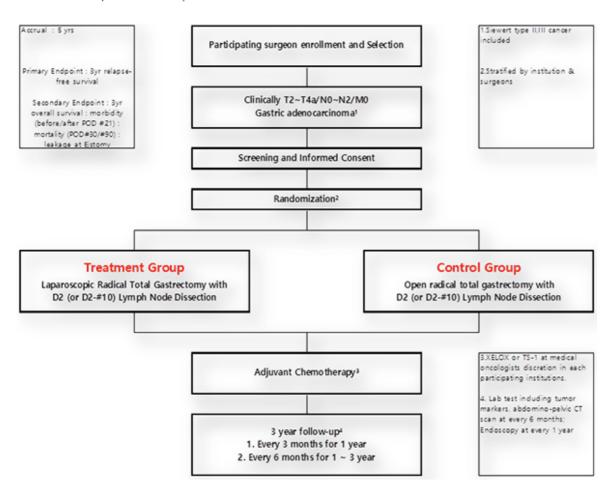
The results of the CLASS02 trial showed that the **Safety** of LTG with lymphadenectomy by experienced surgeons for clinical stage I.







Multicenter, Prospective, Phase III trial Primary endpoint: 3 year relapse free survival Estimated sample size - 772 patients



~772 hasta Tarih: 2018-2027





Must or Cot	RG		LG		Maint	Odds Ratio	Odds Ratio
Study or Subgroup			Events			M-H, Random, 95% CI	M-H, Random, 95% CI
Nhossaini 2019	10	25	11	30	1.3%	1.15 [0.39, 3.43]	
Cianchi 2016	6	30	8	41	1.2%	1.03 [0.32, 3.36]	
Eom 2012	4	30	4	62	0.8%	2.23 [0.52, 9.62]	
3ao 2018	22	163	46	339	4.1%	0.99 [0.58, 1.72]	
Han 2015	13	68	15	68	2.1%	0.84 [0.36, 1.92]	<del></del>
Ho-Jung 2020	97	421	374	1663	8.7%	1.03 [0.80, 1.33]	
Hong 2016	30	232	32	232	4.2%	0.93 [0.54, 1.59]	
Huang 2014	9	72	6	73	1.3%	1.60 [0.54, 4.74]	
lyun 2013	18	38	32	83	2.4%	1.43 [0.66, 3.11]	<del></del>
Junfeng 2014	7	120	17	394	1.9%	1.37 [0.56, 3.40]	<del>-  </del>
Kang 2012	14	100	29	282	2.9%	1.42 [0.72, 2.81]	+
(im HI 2013	9	172	20	481	2.2%	1.27 [0.57, 2.85]	
(im HI 2016	22	185	19	185	3.2%	1.18 [0.62, 2.26]	-
Kim KM 2012	44	436	81	861	6.1%	1.08 [0.73, 1.59]	+
Kim MC 2010	0	16	1	11	0.2%	0.21 [0.01, 5.71]	
(im YW 2015	5	87	26	288	1.6%	0.61 [0.23, 1.65]	
Kong 2019	37	294	105	750	5.9%	0.88 [0.59, 1.32]	-
ee 2015	14	133	34	267	3.1%	0.81 [0.42, 1.56]	
i Z 2018	15	112	13	112	2.3%	1.18 [0.53, 2.60]	
iu 2018	5	100	9	135	1.3%	0.74 [0.24, 2.27]	
Lu 2018	14	101	38	303	3.1%	1.12 [0.58, 2.17]	
Nakauchi 2016	2	84	56	437	0.8%	0.17 [0.04, 0.69]	
Noshiro 2014	2	21	16	160	0.7%	0.95 [0.20, 4.45]	
Obama 2016	38	315	62	525	5.5%	1.02 [0.67, 1.58]	
Okumura 2015	45	370	24	132	4.1%	0.62 [0.36, 1.07]	
	30	151	19	151	3.3%		
Parisi 2017						1.72 [0.92, 3.22]	
Park 2015	12	145	46	612	3.1%	1.11 [0.57, 2.15]	
Pugliese 2010	1	16	6	48	0.4%	0.47 [0.05, 4.20]	
Seo 2014	11	40	12	40	1.6%	0.89 [0.34, 2.33]	
Shen 2016	9	93	33	330	2.4%	0.96 [0.44, 2.09]	
Son SY 2012	2	21	2	42	0.4%	2.11 [0.28, 16.10]	and the same of th
Son T 2014	8	51	13	58	1.6%	0.64 [0.24, 1.71]	
Song 2009	1	20	2	20	0.3%	0.47 [0.04, 5.69]	
Buda 2015	2	88	54	438	0.8%	0.17 [0.04, 0.69]	The second second
Jyama 2012	2	25	38	225	0.8%	0.43 [0.10, 1.89]	0
Wang WJ 2018	42	223	78	223	5.4%	0.43 [0.28, 0.67]	
Noo 2011	26	236	81	591	4.9%	0.78 [0.49, 1.25]	
rang 2016	9	173	63	511	2.7%	0.39 [0.19, 0.80]	
/oon 2012	6	36	10	65	1.3%	1.10 [0.36, 3.32]	
Total (95% CI)		5043		11268	100.0%	0.91 [0.79, 1.04]	•
Total events	643		1535				
Heterogeneity: Tau <sup>2</sup> =	0.04: Ch	i <sup>2</sup> = 49.		8 (P = 0.	10);  2 = 2	4%	0.01 0.1 1 10 1

Fig. n.10. Overall complication.

higher <u>operating time</u> [MD 44.73, (95%Cl 36.01, 53.45) p <0.00001] less <u>intraoperative blood loss</u> [MD -18.24, (95%Cl -25.21, 11.26) p <0.00001] lower rate of <u>major surgical complication</u> [OR 0.66, (95%Cl 0.49, 0.88) p =0.005] increased number of <u>retrieved lymph nodes</u> [MD 1.84, (95%Cl 0.84, 2.84) p =0.0003]



# 2018-2020 Lap vs. Robotik - **RCT** intra-abdominal infectious complications

**No** significant difference in the incidence of intra-abdominal infectious complications (per-protocol) (10 of 117 [8.5%] in the LG group vs 7 of 113 [6.2%] in the RG group)

grade II or higher was significantly higher in the LG group (23 [19.7%]) than in the RG group (10 [8.8%]) (P = .02) grade IIIa or higher significantly higher in the LG group (19 [16.2%]) than in the RG group (6 [5.3%]) (P = .01)







Japan Clinical Oncology Group(日本臨床腫瘍研究グループ) 胃がんグループ

国立がん研究センター研究開発費 2020-J-3 「成人固形がんに対する標準治療確立のための基盤研究」班

## **JCOG1907**

cT1-2N0-2 胃癌におけるロボット支援下胃切除術の腹腔鏡下胃切除術に対する

優越性を検証するランダム化比較試験実施計画書 ver. 1.1.0

Randomized controlled phase III trial to investigate superiority of

robot-assisted gastrectomy over laparoscopic gastrectomy for

clinical stage T1-2N0-2 Gastric Cancer patients

略称: MONA LISA study

グループ代表者:寺島 雅典

静岡県立静岡がんセンター 胃外科

研究代表者:寺島 雅典

静岡県立静岡がんセンター 胃外科

〒411-8777 静岡県駿東郡長泉町下長窪 1007

研究事務局:幕内 梨恵

がん研究会有明病院 胃外科 〒135-8550 東京都江東区有明 3-8-31

2019年7月8日

JCOG 運営委員会プロトコールコンセプト承認(PC1907)

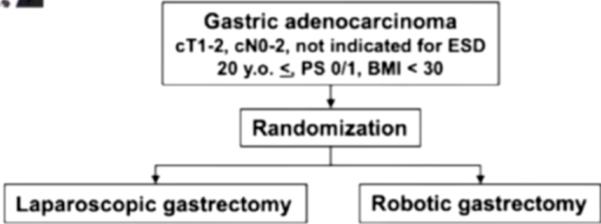
2020年1月19日

ver. 1.0.0 JCOG プロトコール審査委員会承認

2021年5月28日

ver. 1.1.0 改訂 JCOG 効果·安全性評価委員会承認 6 月 4 日発効

## JCOG1907 (MONALISA)



Primary endpoint: Intraabdominal infectious complication

(C-D Grade II or worse)

Secondary endpoints: OS, RFS, Morbidity, Mortality,

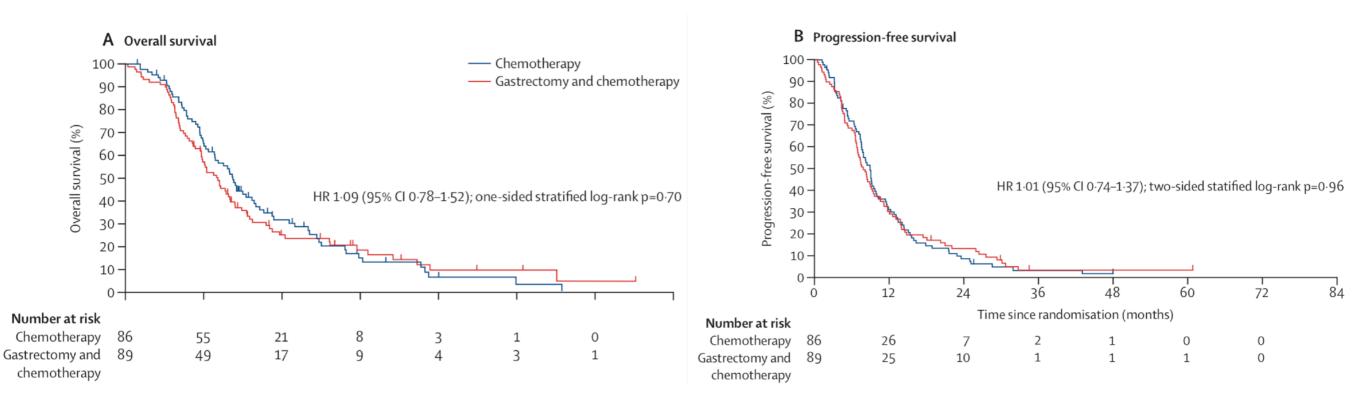
Completion rate of protocol Tx, Early

postoperative outcomes

Sample size:

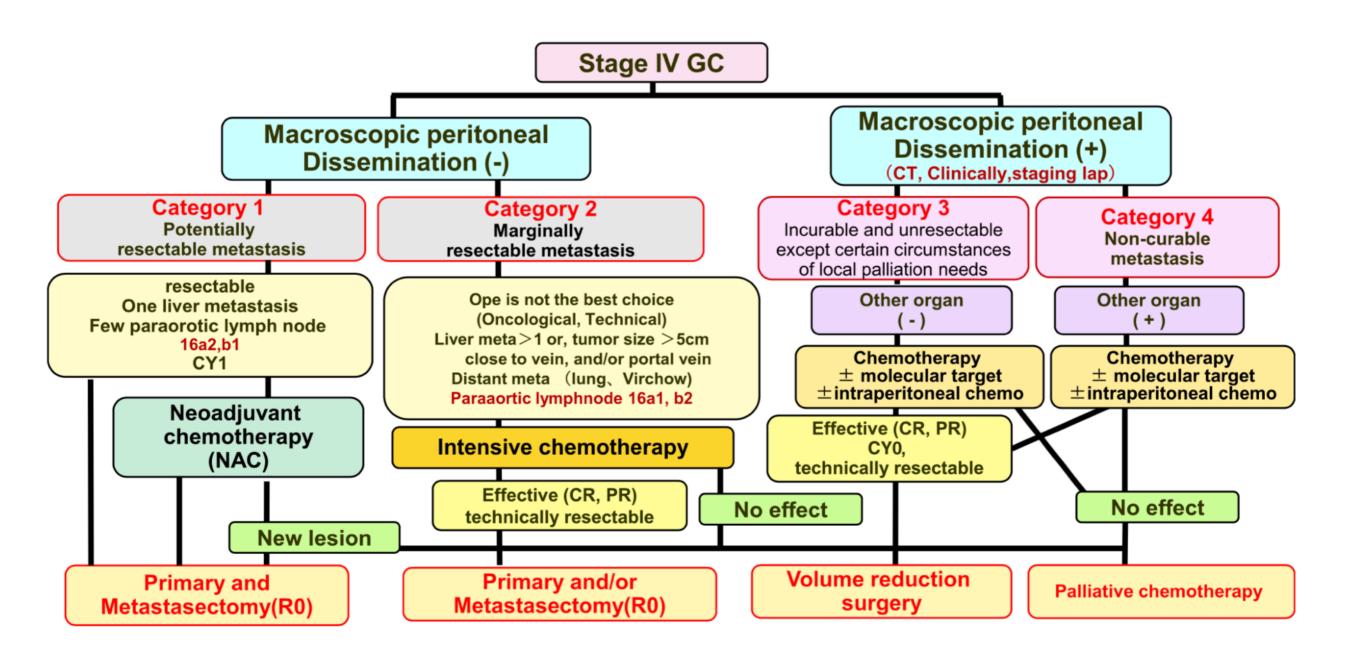
1040 pts.



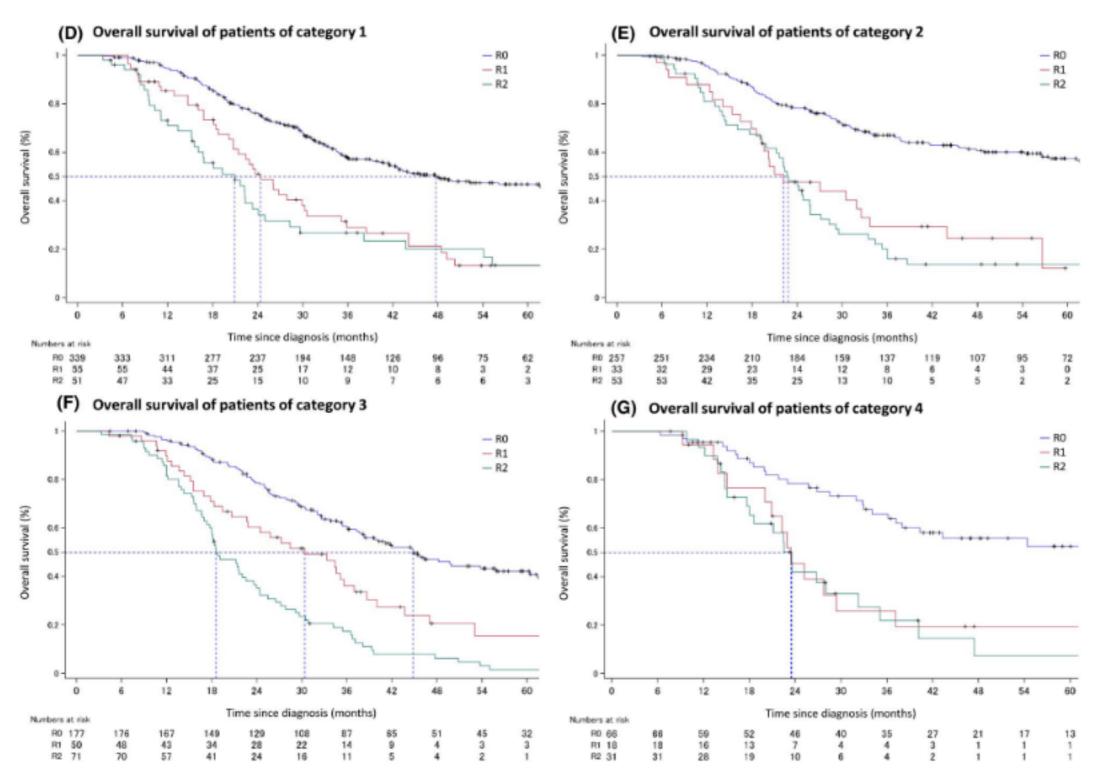


gastrectomy followed by chemotherapy **did not show any survival benefit** compared with chemotherapy alone in advanced gastric cancer with a single non-curable factor, gastrectomy cannot be justified for treatment of patients with these tumours.

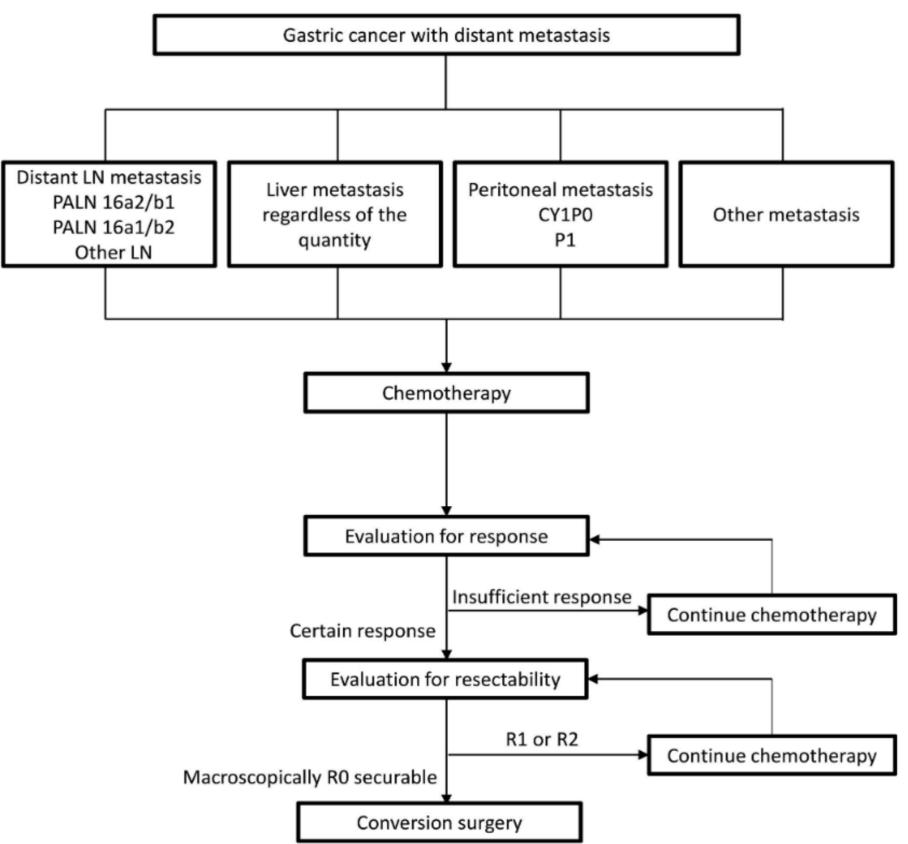






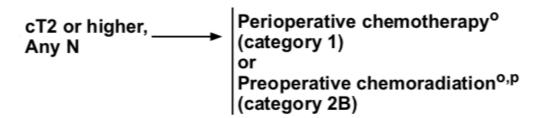






# **Gastric Cancer**

Version 2.2022 — January 11, 2022



pT3,pT4,Any N o<u>r</u> Any pT,N+

Fluoropyrimidine (fluorouracil or capecitabine)<sup>o,r</sup>
then fluoropyrimidine-based chemoradiation,<sup>o,p,r</sup>
then fluoropyrimidine (fluorouracil or capecitabine)<sup>o,r</sup>
if less than a D2 dissection (category 1)
or
Chemotherapy for patients who have undergone primary D2
lymph node dissection<sup>f,o</sup> (category 1)

#### **GAST-C** Principles of Surgery

- Siewert classification: The term "center" was clarified as "epicenter".
- Resectable tumors, Bullets and sub-bullets revised:
- ► T1b-T3: Adequate gastric resection to achieve negative microscopic margins (typically ≥4 cm from gross tumor).
- "...at least 45 16 or greater lymph nodes."
- D2 dissection is a D1 plus all the nodes along the left gastric artery, common hepatic artery, celiac artery, splenic hilum, and splenic artery.
- Routine or prophylactic splenectomy is not required. Splenectomy is acceptable when the spleen or the hilum is involved Routine splenectomy is not indicated unless the spleen is involved or extensive hilar adenopathy is noted.

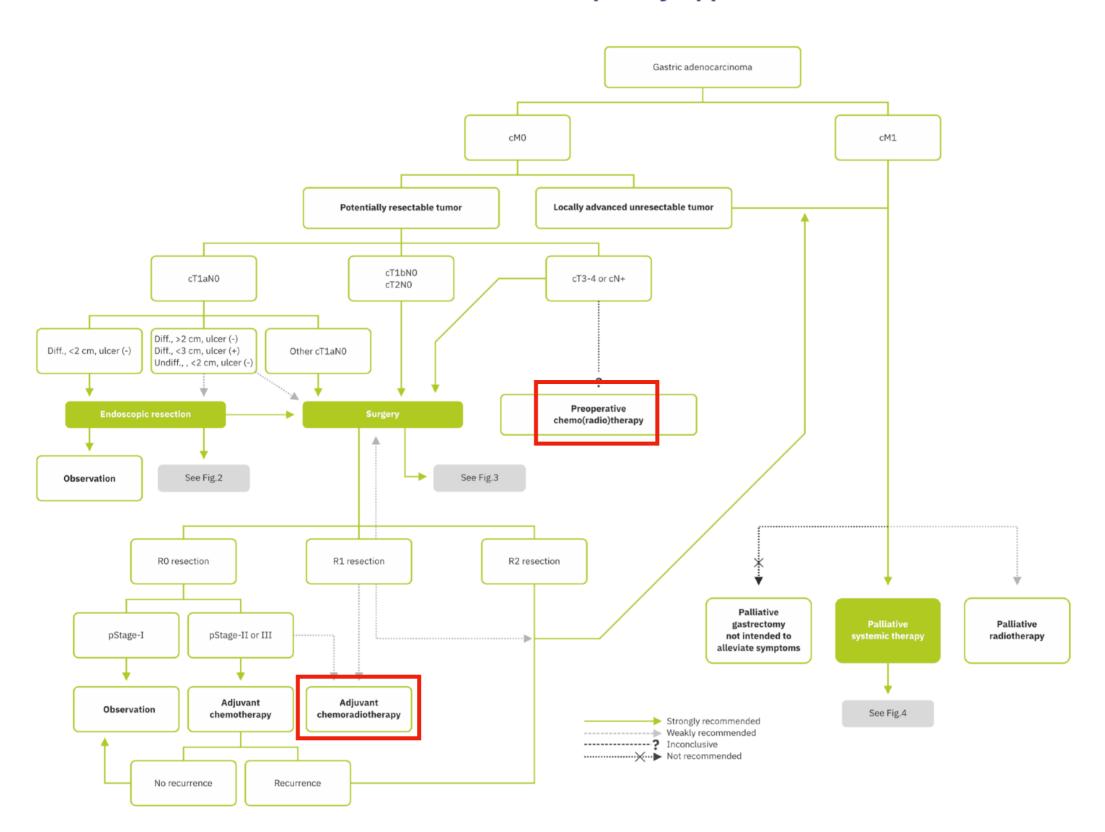
Hyperthermic intraperitoneal chemotherapy (HIPEC) or laparoscopic HIPEC may be a therapeutic alternative for carefully selected stage IV
patients in the setting of ongoing clinical trials and is under further clinical investigation.<sup>18-20</sup>



#### Special Article



## Korean Practice Guideline for Gastric Cancer 2018: an Evidence-based, Multi-disciplinary Approach

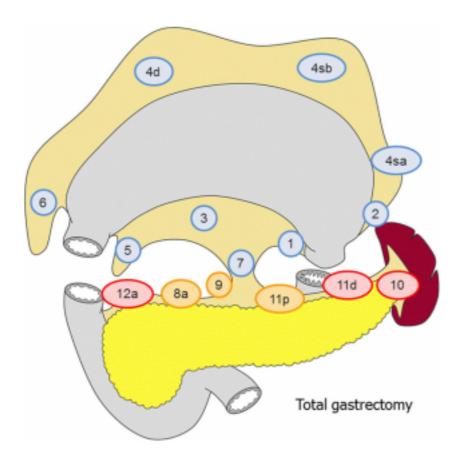


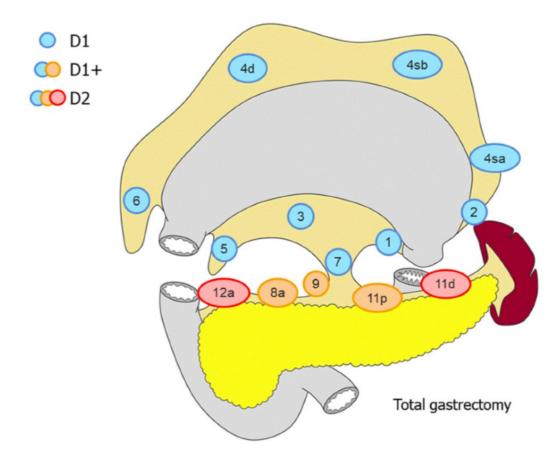
#### SPECIAL ARTICLE

#### Japanese gastric cancer treatment guidelines 2018 (5th edition)

#### Japanese Gastric Cancer Association<sup>1</sup>

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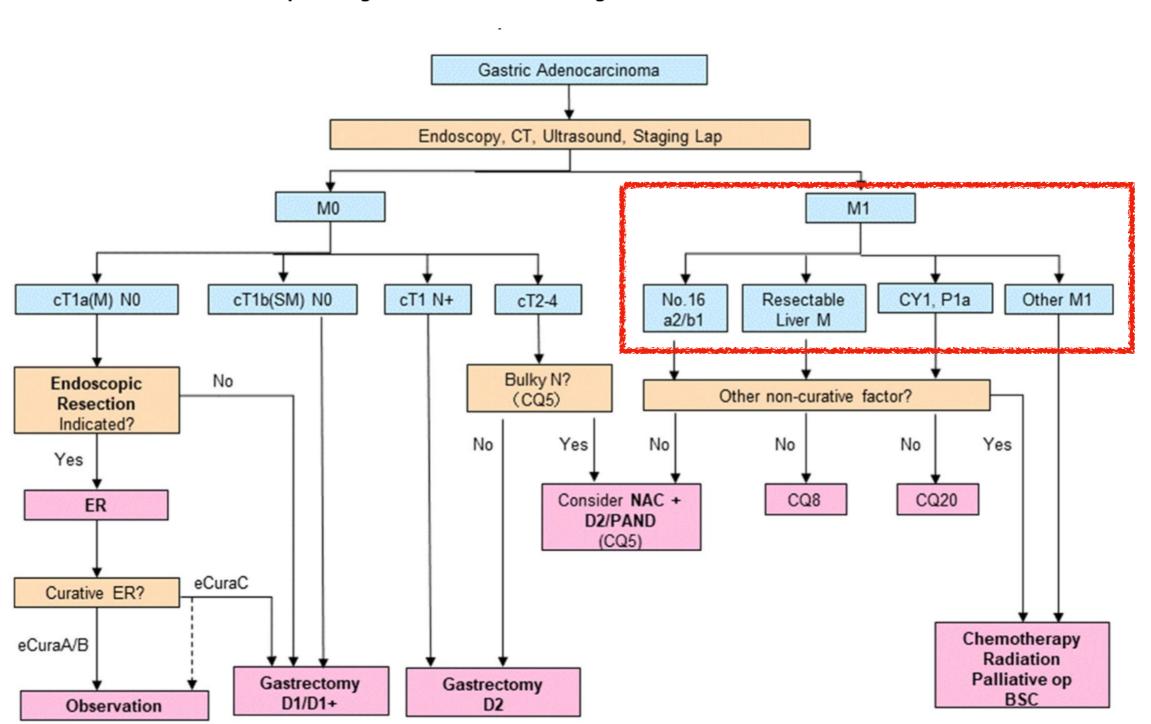


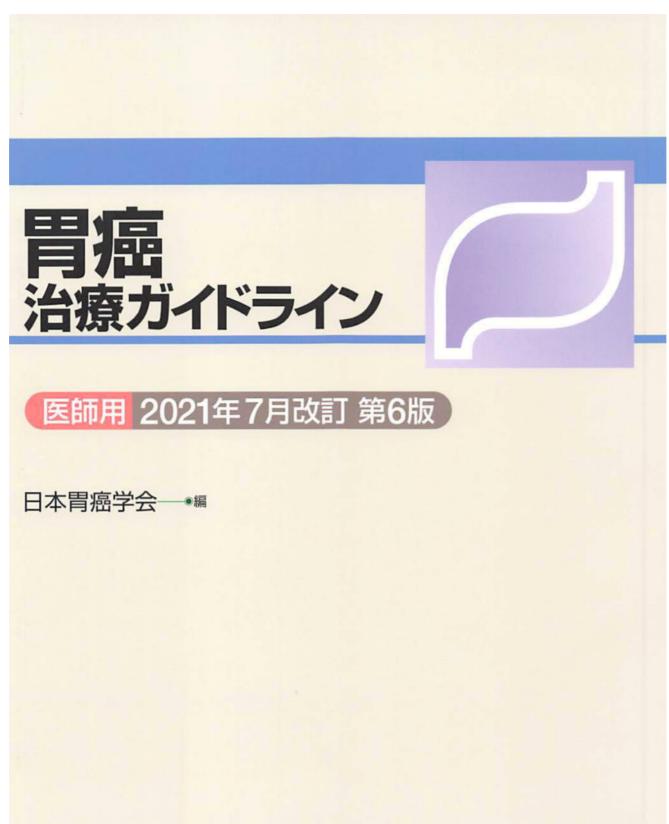


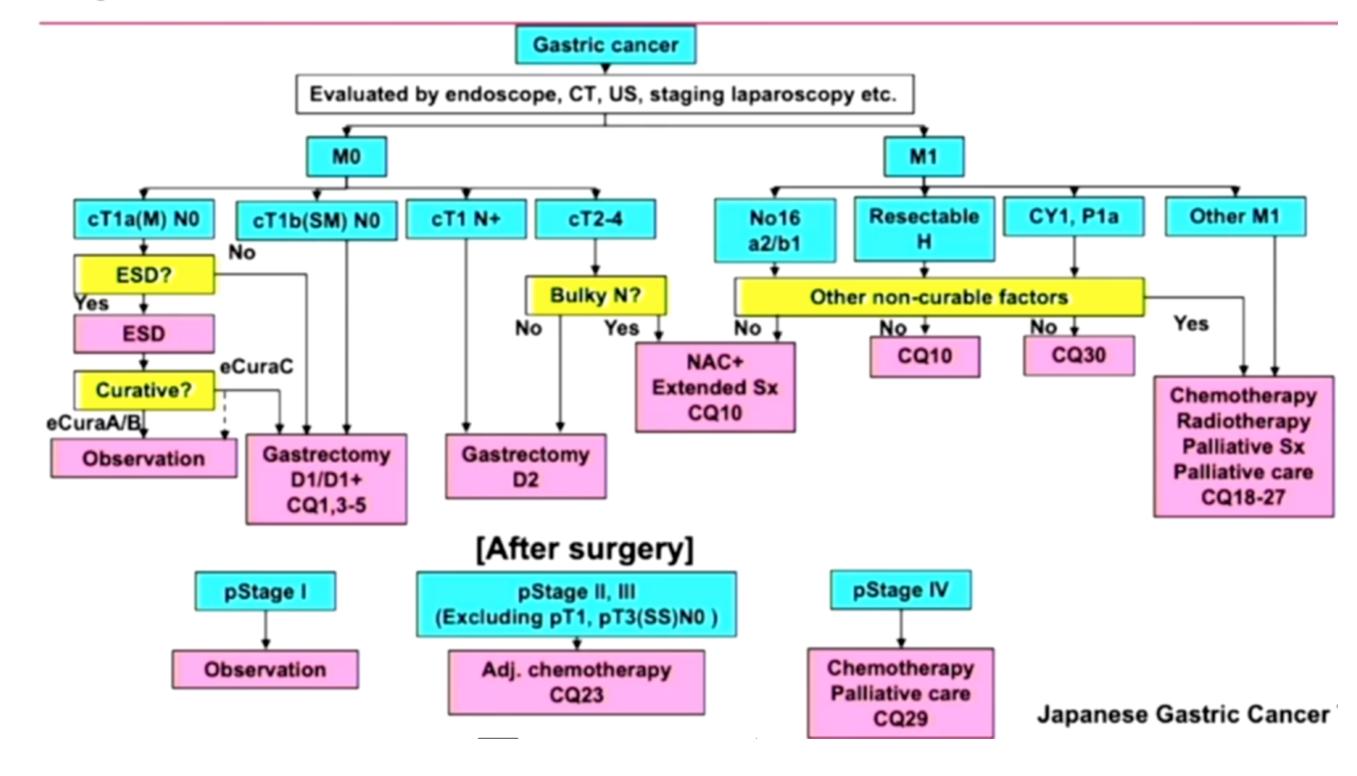
4th 5th

#### SPECIAL ARTICLE

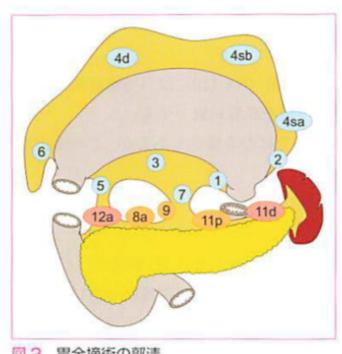
#### Japanese gastric cancer treatment guidelines 2018 (5th edition)







6th



4sb 4d 3 12a 8a 9

図2 胃全摘術の郭清

図3 幽門側胃切除術の郭清



4sb 3a 図 5 噴門側胃切除術の郭清

1) 胃全摘術 (図2)

D0: D1 に満たない郭清

D1: No. 1~7

D1+: D1+No. 8a, 9, 11p

D2: D1+No. 8a, 9, 11p, 11d, 12a

食道浸潤癌では D2 には No. 19. 20. 110\*を追加する。

2) 幽門側胃切除術 (図3)

D0: D1 に満たない郭清

D1: No. 1, 3, 4sb, 4d, 5, 6, 7

D1+: D1+No. 8a, 9

D2: D1+No. 8a, 9, 11p, 12a



3) 幽門保存胃切除術 (図4)

D0: D1 に満たない郭清

D1: No. 1, 3, 4sb, 4d, 6\*\*, 7

D1+: D1+No. 8a, 9

4) 噴門側胃切除術 (図5)

D0: D1 に満たない郭清

D1: No. 1, 2, 3a, 4sa, 4sb, 7

D1+: D1+No. 8a, 9, 11p

D2: D1+No. 8a, 9, 11p, 11d





6th

**IGCC 2022** 

Laparoscopic distal gastrectomy is strongly recommended as a standard treatment

(Consensus rate 100%(8/8), Evidence level A)

Laparoscopic total or proximal gastrectomy is weakly recommended

(Consensus rate 100% (8/8), Evidence level C)

We cannot decide the recommendation of laparoscopic gastrectomy for cStage II, III gastric cancer (Agreement rate 71.4% (5/7), Evidence level C)

Laparoscopic distal gastrectomy for cStage II, III gastric cancer is strongly recommended as a standard treatment.

Robotic gastrectomy is weakly recommended cStage I gastric cancer.

Omentectomy is weakly recommended for cT3-T4 gastric cancer

(Consensus rate 100% (8/8), Evidence level C)

It is strongly recommended not to perform splenectomy or splenic hilar lymph node dissection for tumors without greater curvature invasion.

(Consensus rate 100% (8/8), Evidence level A)

Splenectomy or splenic hilar lymph node dissection is weakly recommended for tumors with greater curvature invasion

(Consensus rate 87.5% (7/8), Evidence level C).

In surgery for esophagogastric junction cancer deeper than cT2, lower mediastinal lymph node dissection is weakly recommended if the esophageal invasion length is more than 2 cm, and upper, middle, and lower mediastinal lymph node dissection is weakly recommended if the esophageal invasion length is more than 4 cm. (Consensus rate 100% (9/9), Evidence level C)

Surgical resection after preoperative chemotherapy is weakly recommended for a small number of paraaortic lymph node metastases confined to No. 16a2 / b1.

In addition, surgical resection is weakly recommended for solitary liver metastasis without other non-curative factors

(Consensus rate 100% (7/7), Evidence level C)

Conversion surgery for patients with Stage IV gastric cancer is weakly recommended with the condition that chemotherapy provides a certain antitumor effect, the response is maintained, and R0 resection is possible

(Consensus rate 100% (7/7), Evidence level D)

No clear recommendation for neoadjuvant chemotherapy for curatively resectable advanced gastric and esophagogastric junctional cancer (Consensus rate 71.4 %, 5/7, Strength of evidence B).

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## Lenfadenektomi

- D2; standart
- D2+PALN; rutinde gereksiz.

# **Splenektomi**

• rutinde gereksiz.



#### **Bursektomi**

• rutinde gereksiz.



## **Omentektomi**

• 2031



## Metastatik

Konversiyon tedavisi 😂



#### **EIPL**

Faydasız

# Laparoskopi

EGC-Distal



- AGC-Distal
- AGC-Total