

NOUVEAUTES DANS LE TRAITEMENT DES CANCERS SUPERFICIELS DU RECTUM T1

Professeur Stanislas CHAUSSADE

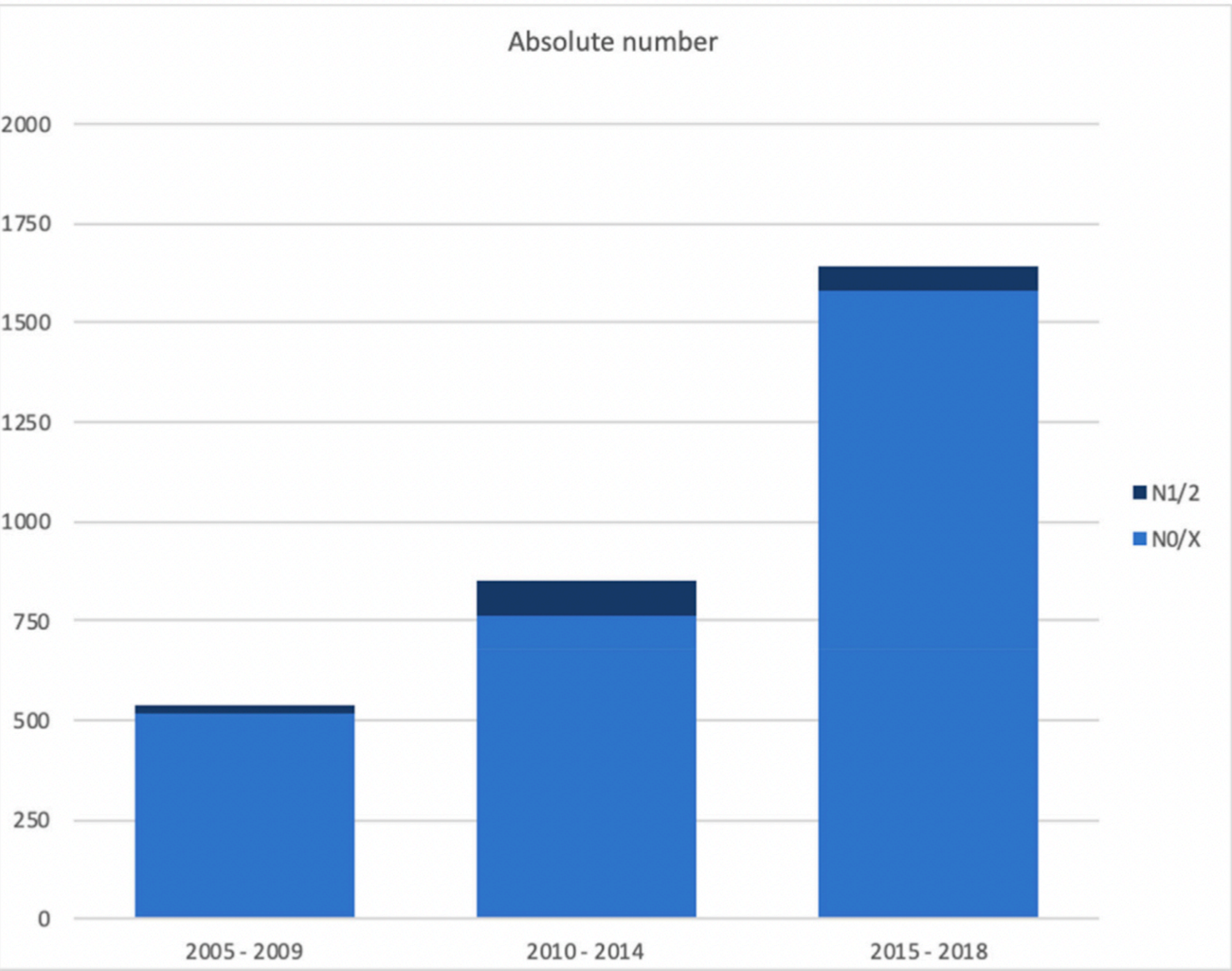
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INTRODUCTION

- Epidemiologie
- TNCD sur les cancer du rectum 2023
- Stratégie de conservation non chirurgicale
 - « watch and wait » après réponse tumorale par radiochimiothérapie pour des tumeurs plus avancées
 - Amélioration des traitements endoscopiques
 - Meilleure connaissance des risques de métastases ganglionnaires ou à distance

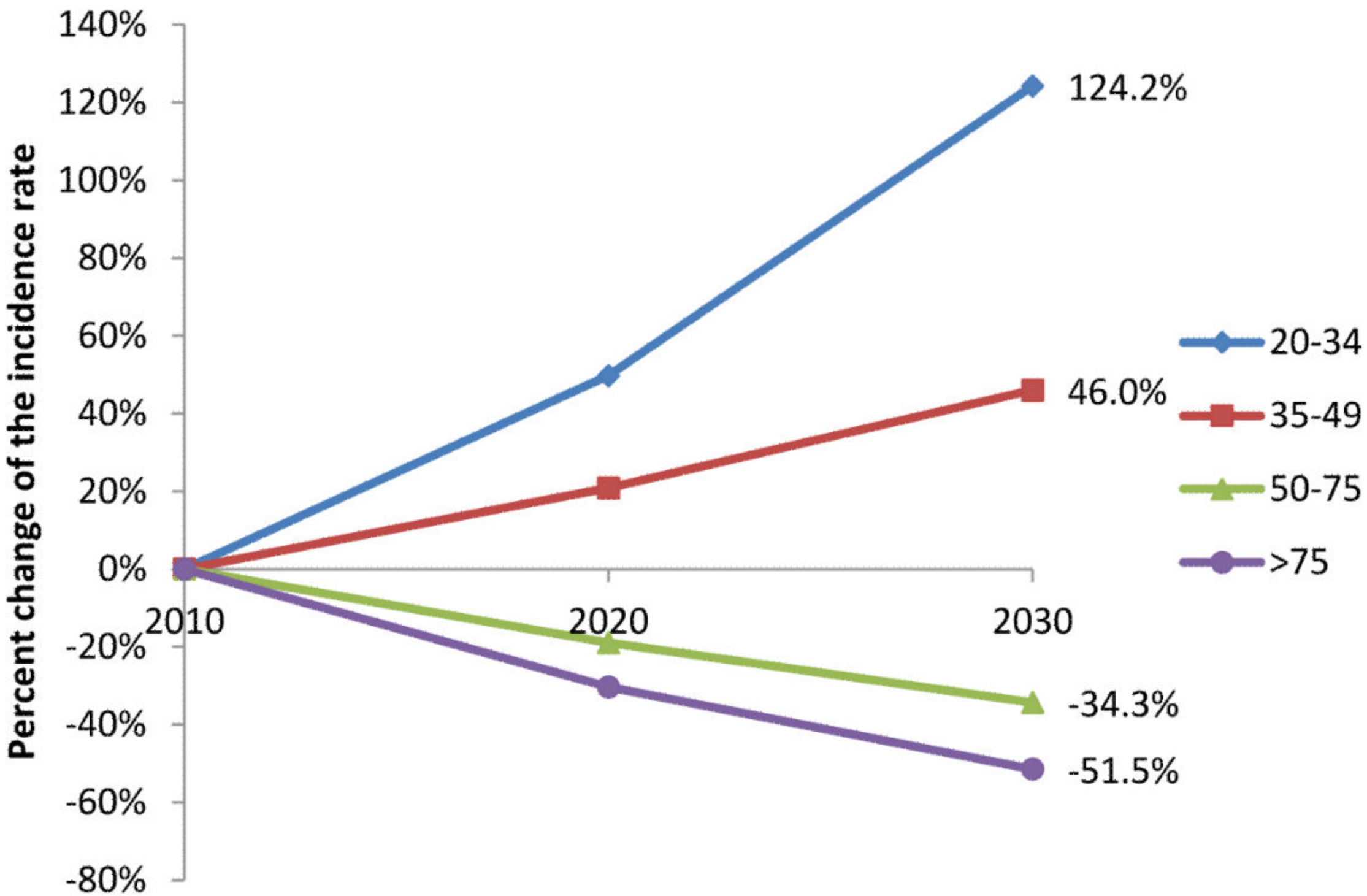
Treatment of clinical T1 rectal cancer in the Netherlands; a population-based overview of clinical practice

M. Verseveld ^{a,b,*}, D. Verver ^a, B.J. Noordman ^{a,b}, S. Pouwels ^c, M.A.G. Elferink ^d, E.J.R. de Graaf ^e, C. Verhoef ^b, P.G. Doornebosch ^e, J.H.W. de Wilt ^f



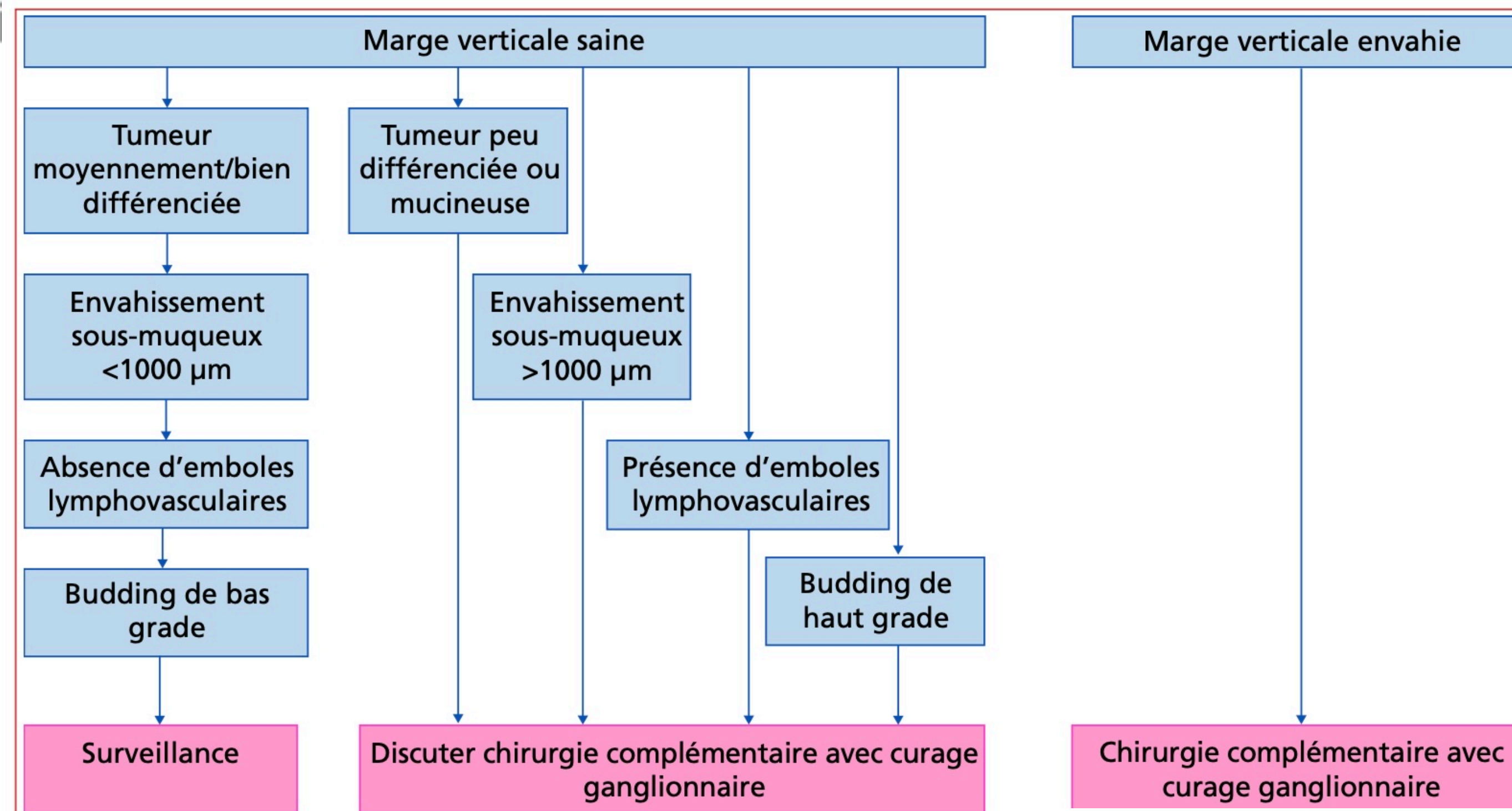
EPIDEMIOLOGIE CCR USA

ADK RECTO-SIGMOÏDE



Current problems and perspectives of pathological risk factors for lymph node metastasis in T1 colorectal cancer: Systematic review

Katsuro Ichimasa,¹ Shin-ei Kudo,¹ Hideyuki Miyachi,¹ Yuta Kouyama,¹ Kenichi Mochizuki,¹ Yuki Takashina,¹ Yasuharu Maeda,¹ Yuichi Mori,^{1,8} Toyoki Kudo,¹ Yuki Miyata,¹ Yoshika Akimoto,¹ Yuki Kataoka,^{3,4,5,6} Takafumi Kubota,^{6,7} Tetsuo Nemoto,² Fumio Ishida¹ and Masashi M

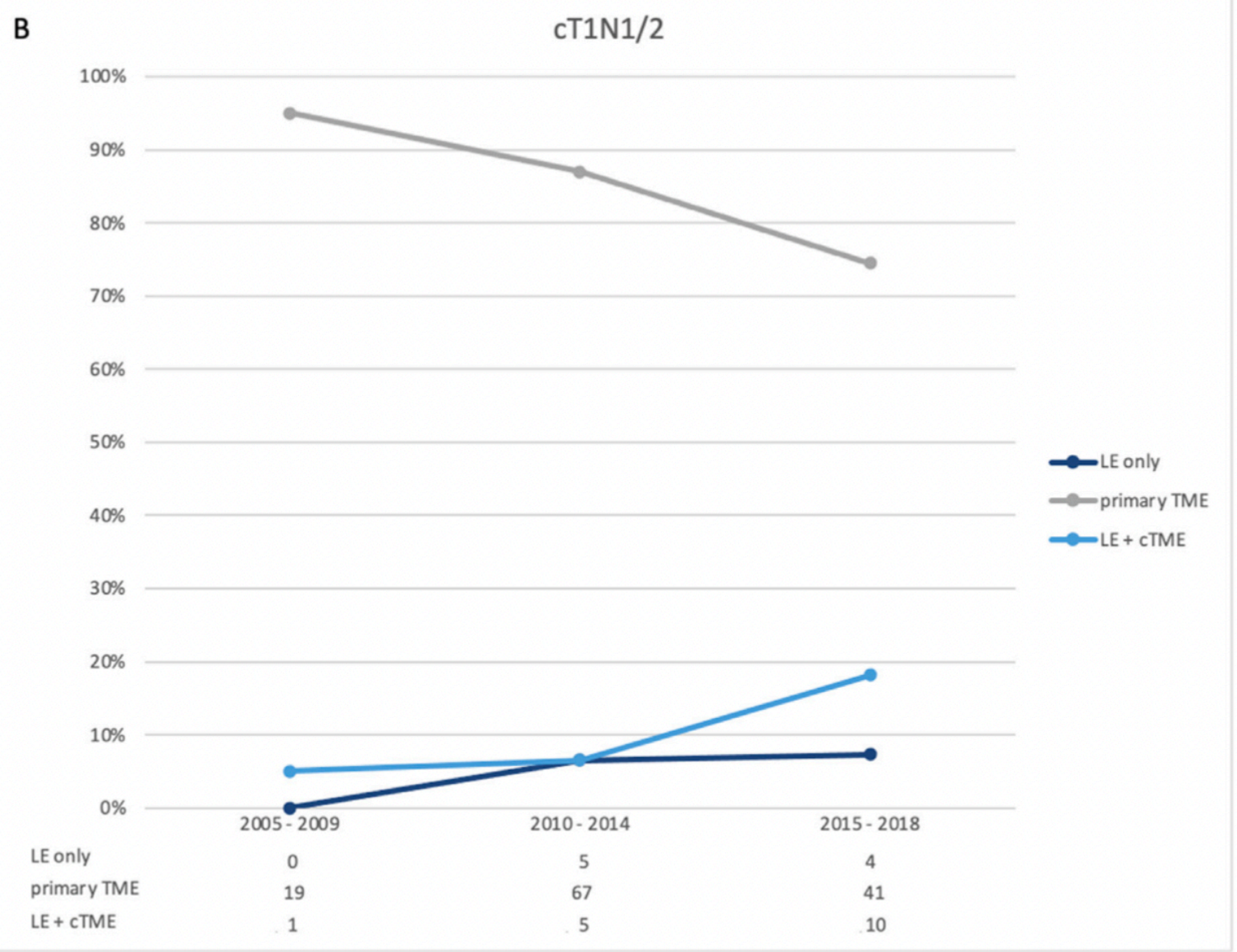
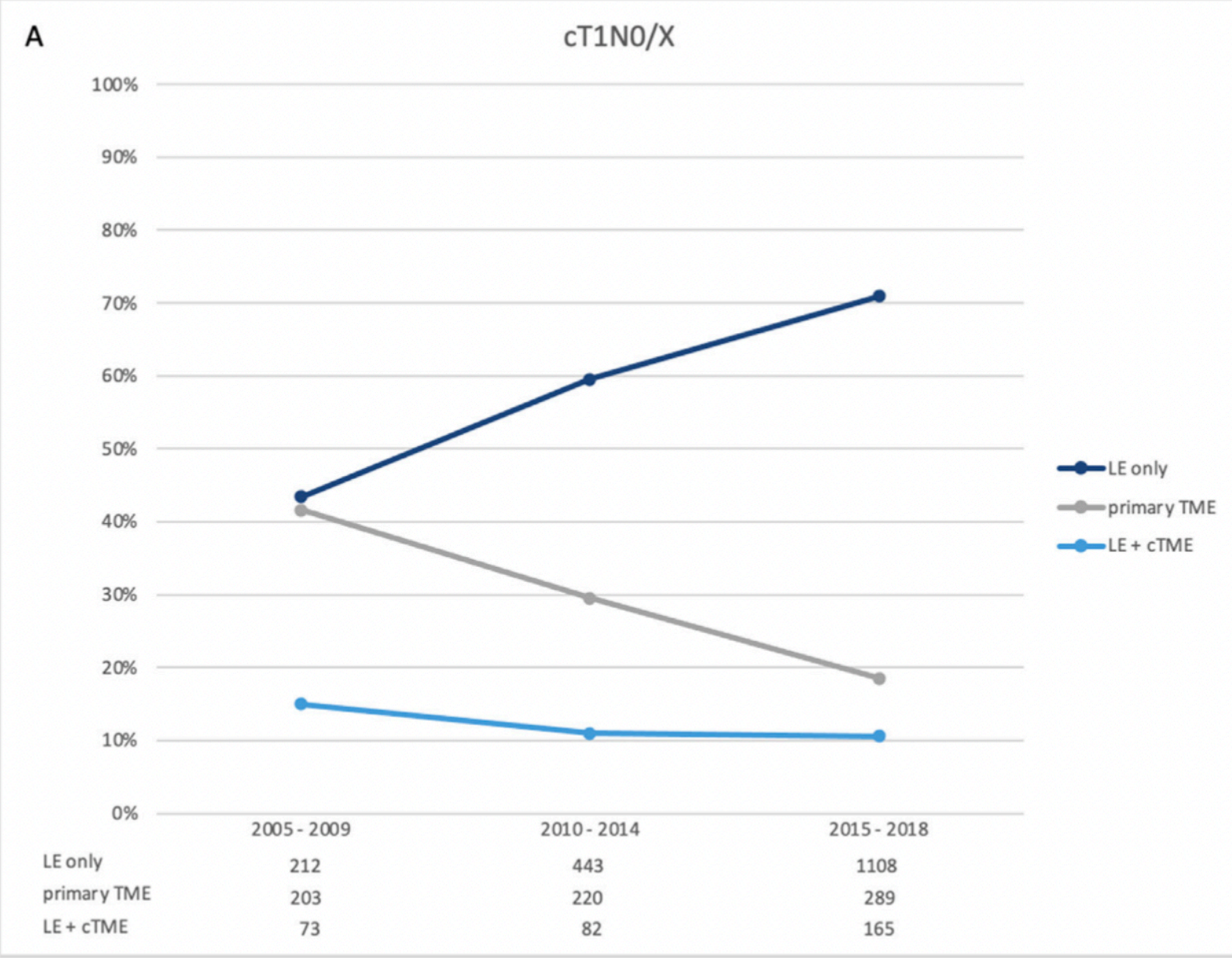
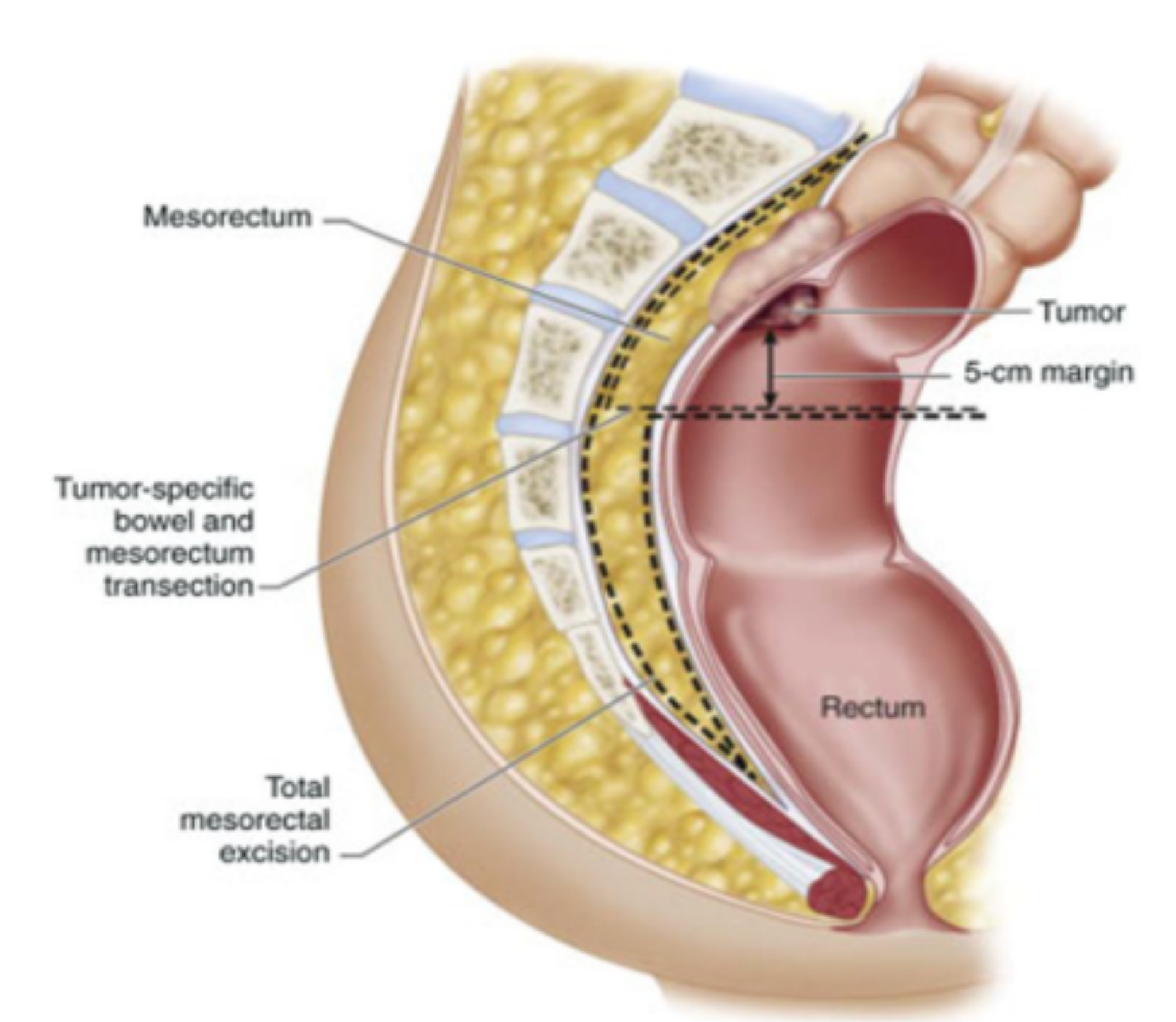


Japanese Society for Cancer of the Colon and Rectum (JSCCR) guidelines 2019 for the treatment of colorectal cancer

Surgery is the Standard of Care for Early Rectal Cancer

R. Hargest

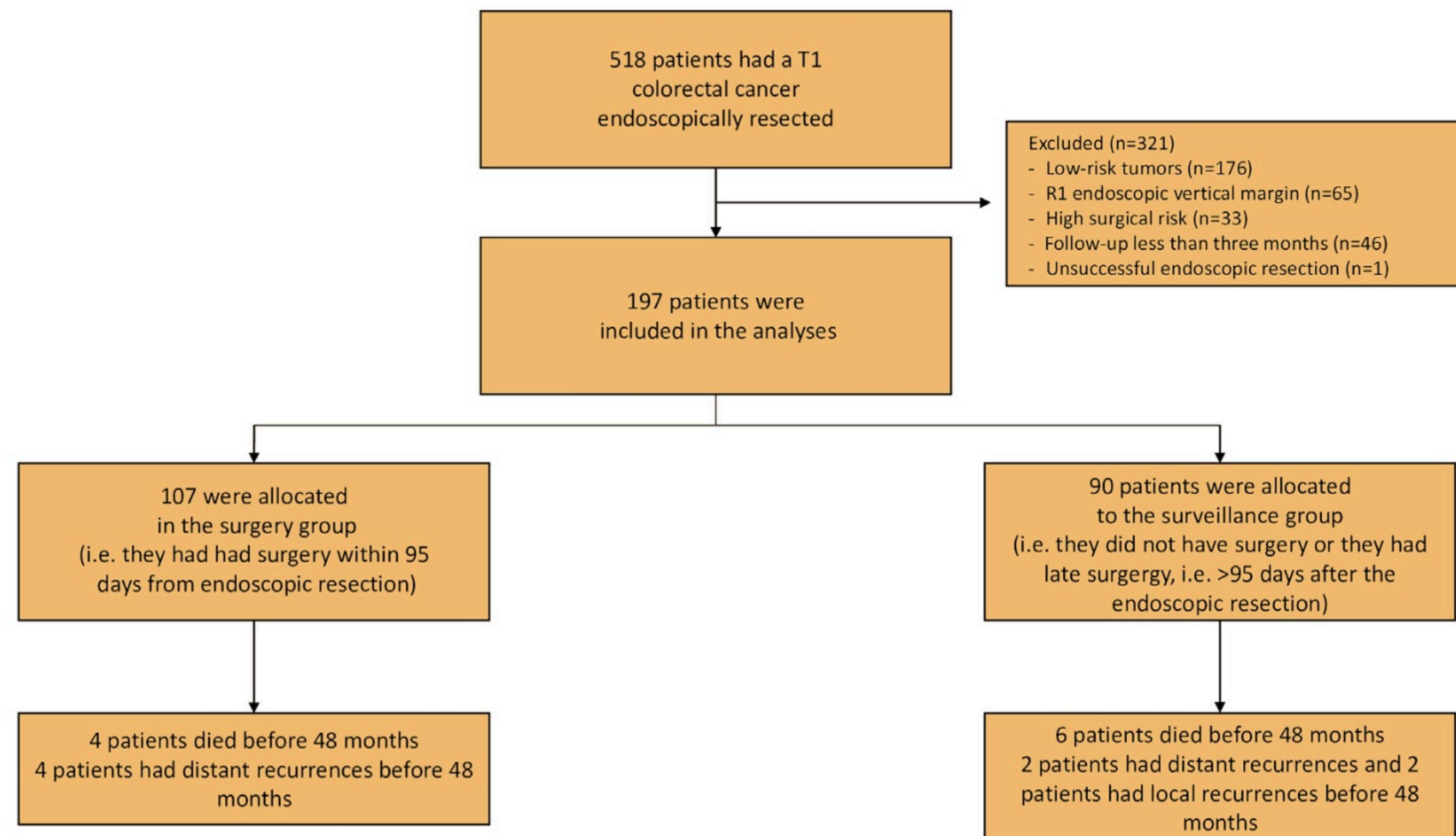
2023



Impact of surgery after endoscopically resected high-risk T1 colorectal cancer: results of an emulated target trial

2024

Félix Corre, MD, MSc,^{1,2} Jérémie Albouys, MD, MSc,³ Viet-Thi Tran, MD, PhD,⁴ Vincent Lepilliez, MD,⁵ Jean-Philippe Ratone, MD,⁶ Emmanuel Coron, MD, PhD,^{7,8} Thomas Lambin, MD, MSc,⁹ Gabriel Rahmi, MD, PhD,¹⁰ David Karsenti, MD,¹¹ Jean-Marc Canard, MD,¹² Edouard Chabrun, MD,¹³ Marine Camus, MD, PhD,¹⁴ Timothée Wallenhorst, MD,¹⁵ Jean-Baptiste Chevaux, MD,¹⁶ Marion Schaefer, MD, MSc,¹⁶ Romain Gerard, MD,¹⁷ Alexandre Rouquette, MD,^{2,18} Benoit Terris, MD, PhD,^{2,18} Romain Coriat, MD, PhD,^{1,2} Jérémie Jacques, MD, PhD,³ Maximilien Barret, MD, PhD,^{1,2} Mathieu Pioche, MD, PhD,⁹ Stanislas Chaussade, MD, PhD,^{1,2}



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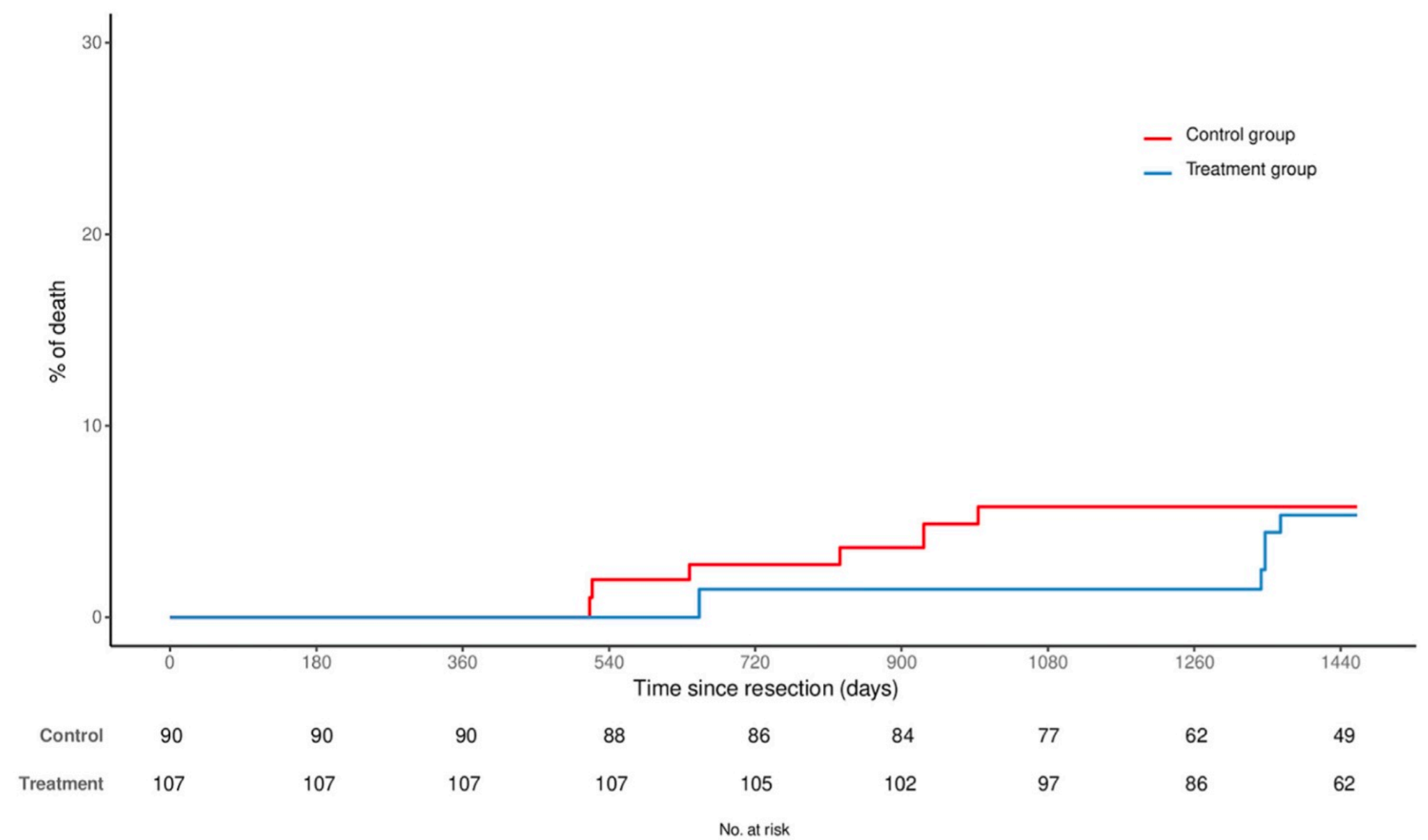
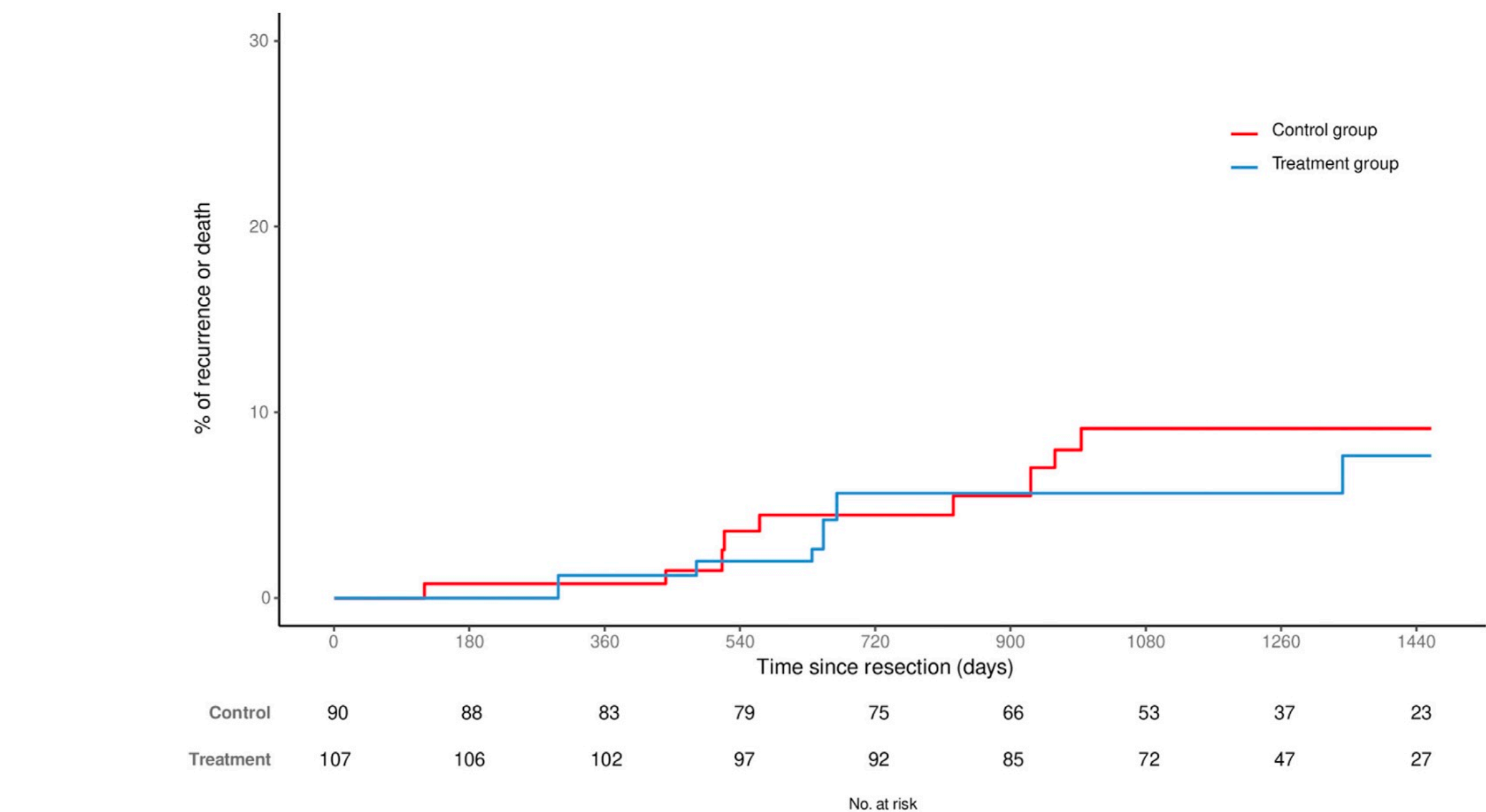
Corre et al Impact of surgery after endoscopically resected high-risk T1 CRC

Figure 3. Death rate of patients in the treatment and control groups after balancing the baseline covariates by inverse probability of treatment weighting (weighted hazard ratio, 1.19; 95% confidence interval, .49-2.88).

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Pas d'impact de la Chirurgie : pourquoi ?

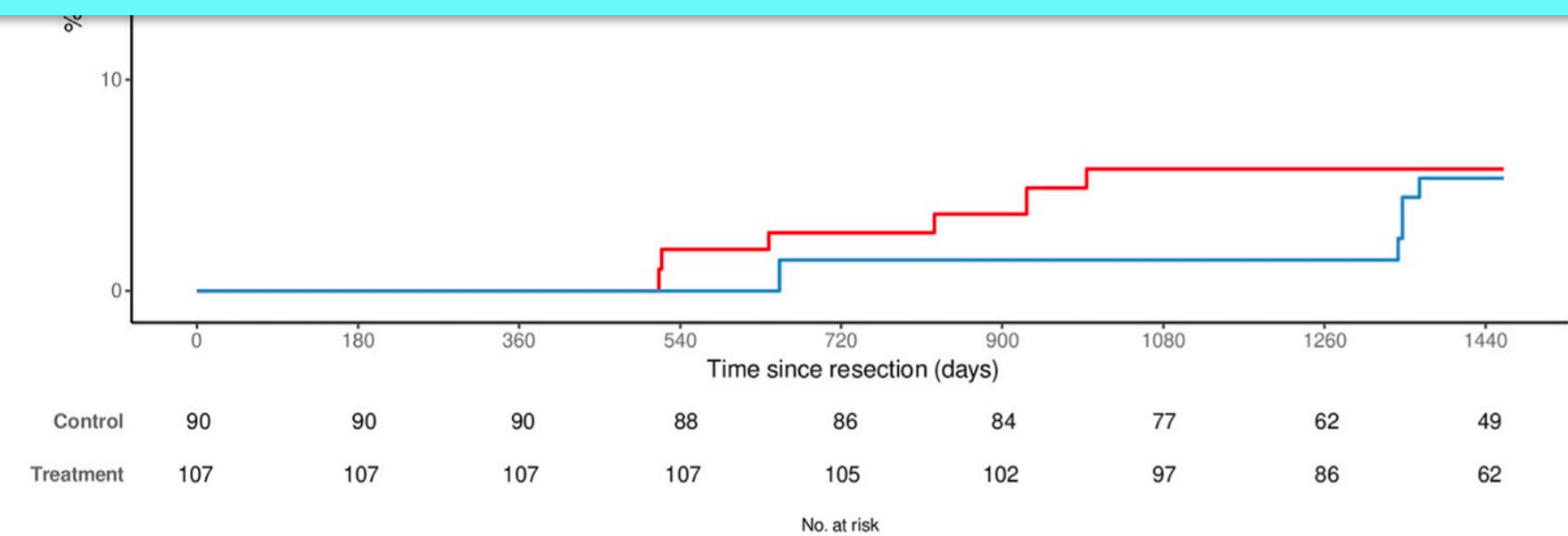
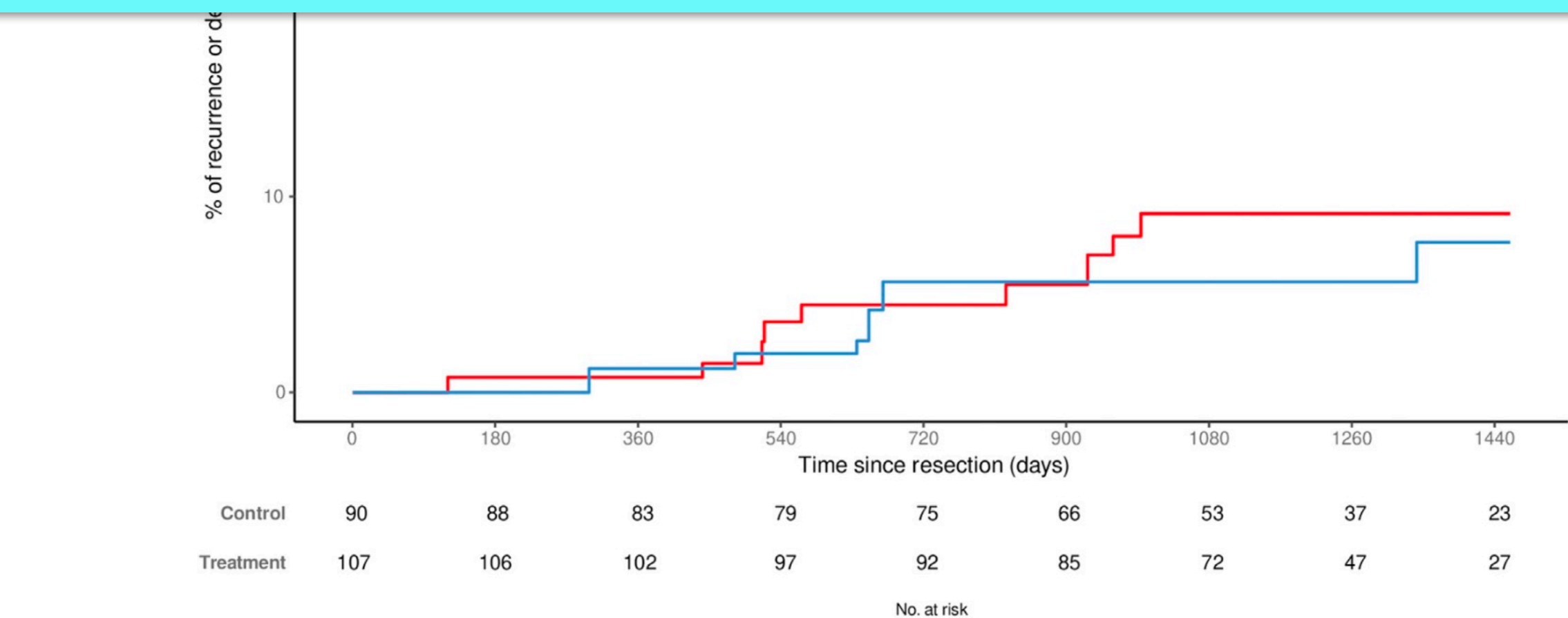


Figure 3. Death rate of patients in the treatment and control groups after balancing the baseline covariates by inverse probability of treatment weighting (weighted hazard ratio, 1.19; 95% confidence interval, .49-2.88).

RESULTATS DE LA CHIRURGIE APRES TRAITEMENT ENDOSCOPIQUE

Tumeurs à bas risque : 30%
0 critère histopronostique
péjoratif

Tumeurs à haut risque 70%
> = 1 critère
histopronostique péjoratif

**80% des CCR T1 NE TIRE AUCUN
BENEFICE DE LA CHIRURGIE**

2% N+, 2% R+

20% N+

80% pT0N0

Traitement chirurgical
5% R+ après traitement
chirurgical surtout pour CCR du
recto sigmoïde

QUE DIT LE TNCD ?

Date de cette version :
05/09/2023

Date de dernière mise à jour à vérifier sur
Date de dernière mise à jour à vérifier sur www.tncd.org ou www.snfge.org

Sur le plan carcinologique, une résection locale d'adénocarcinome rectal peut être considérée curative lorsque la résection est R0) et qu'elle présente les critères suivants :

- Adénocarcinomes intra-muqueux purs (T1a)**
- Adénocarcinomes à envahissement sous-muqueux superficiel (sm1)** lorsque l'invasion sous- muqueuse est inférieure à 1000 microns et qu'aucun embol vasculonerveux ni *Budding* de grade ≥ 2 ni composantes indifférenciées n'est détecté par l'anatomo-pathologiste.
- Adénocarcinomes à envahissement sous-muqueux profond ($> sm1$)** lorsque l'invasion sous-muqueuse est supérieure à 1000 microns et qu'aucun embol vasculonerveux ni *Budding* de grade ≥ 2 ni composantes indifférenciées n'est détecté par l'anatomo-pathologiste.

STRATEGIE DE CONSERVATION D'ORGANE

- Amélioration R0
- Discussion des paramètres histologiques
- Autres critères et nomogramme
- Traitement complémentaire non chirurgical :
radiochimiothérapie.

Liselotte W. Zwager, Barbara A.J. Bastiaansen, Nahid S.M. Montazeri, Roel Hompes, Valeria Barresi, Katsuro Ichimasa, Hiroshi Kawachi, Isidro Machado, Tadahiko Masaki, Weiqi Sheng, Shinji Tanaka, Kazutomo Togashi, Chihiro Yasue, Paul Fockens, Leon M.G. Moons, Evelien Dekker

Journal Pre-proof

Deep submucosal invasion is not an independent risk factor for lymph node metastasis in T1 colorectal cancer: a meta-analysis

DSI in relation to LNM in univariable analysis

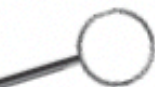
 67 studies

 21,238 patients

Overall LNM-rate: 11.2%

OR 2.58 (95% CI 2.10-3.18)

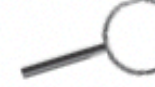
DSI in relation to histological high-risk* factors in multivariable analysis

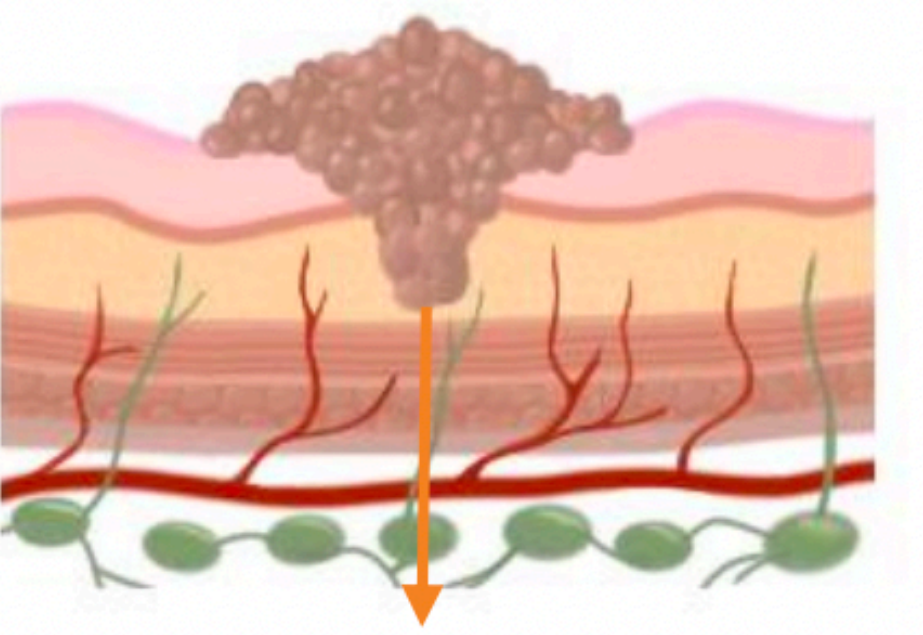
 8 studies, 3621 

Study	OR [95% CI]
Nakadoi et.al., 2011	5.88 [1.36, 25.40]
Kawachi et.al., 2015	6.40 [2.29, 17.90]
Pai et.al., 2017	2.20 [0.71, 6.86]
Shin et.al., 2018	0.88 [0.32, 2.43]
Yasue et.al., 2019	1.61 [0.60, 4.31]
Zhang et.al., 2019	1.84 [0.55, 6.20]
Mochizuki et.al., 2020	0.69 [0.25, 1.91]
Haasnoot et.al., 2020	0.78 [0.26, 2.33]
DSI overall OR	1.73 [0.96, 3.12]

OR 1.73 (95% CI 0.96-3.12)

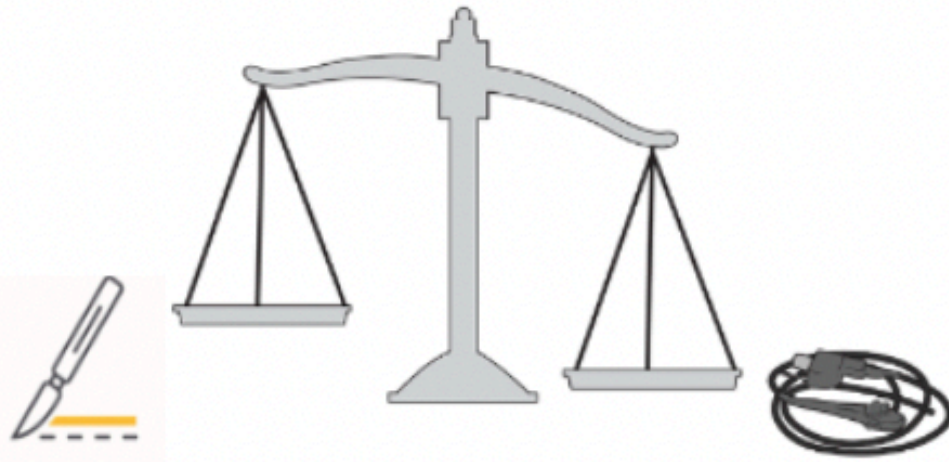
DSI as solitary risk factor for LNM

 8 studies, 1146 



Absolute risk: 2.6%

DSI should be reconsidered as *strong* indicator for oncologic surgery



DSI (deep submucosal invasion); LNM (lymph node metastasis); OR (odds ratio).
*poor differentiation grade, lymphovascular invasion and high-grade tumor budding

A risk stratification for nodal metastasis in T1 colorectal cancer after successful therapeutic endoscopy

		T1b colorectal cancer		T1a colorectal cancer	
Authors of the study	Number of cases	Any other risk factors		Any other risk factors	
		Positive	Negative	Positive	Negative
Oishi <i>et al.</i> ^{3*}	225				
		19/114 16.7%	2/70 2.9%	0/17 0%	0/24 0%
Kawachi <i>et al.</i> ⁴	806				
		79/367 21.5%	14/236 5.9%	3/82 3.7%	1/121 0.8%
Ajioka <i>et al.</i> ⁵	2057				
		191/800 23.9%	41/875 4.7%	1/39 2.6%	1/343 0.3%

Lymphovascular Infiltration, Not Depth of Invasion, is the Critical Risk Factor of Metastases in Early Colorectal Cancer

Retrospective Population-based Cohort Study on Prospectively Collected Data, Including Validation

Carl-Fredrik Rönnow, MD, PhD, Victoria Arthursson, MD,* Ervin Toth, MD, PhD,† Peter-Martin Krarup, MD, PhD,‡ Ingvar Syk, MD, PhD,* and Henrik Thorlacius, MD, PhD*✉*

TABLE 2. Risk of Lymph Node Metastases With 1 Risk Factor Only

	Sm1	Sm2	Sm3	Total
Submucosal invasion, only	17/336 (5.1%)	21/240 (8.8%)	36/424 (8.5%)	74/1000 (7.4%)
LVI*, only	7/21 (33.3%)	5/11 (45.5%)	16/40 (40%)	28/72 (38.9%)
Perineural invasion, only	1/1 (100%)	1/2 (50%)	2/4 (50%)	4/7 (57.1%)
Mucinous subtype, only	1/16 (6.3%)	2/10 (20%)	5/30 (16.7%)	8/56 (14.3%)
High-grade, only	1/17 (5.9%)	0/11 (0%)	2/25 (8%)	3/53 (5.7%)

Risk analysis of submucosal invasive rectal carcinomas for lymph node metastasis to expand indication criteria for endoscopic resection

Shiro Oka,¹ Shinji Tanaka,¹ Koichi Nakadoi,² Hiroyuki Kanao¹ and Kazuaki Chayama²

Depth of SM invasion (μm)	No. cases	Incidence of LN metastasis (%)	95% CI	No. of Limited cases	Limited cases [‡] of LN metastasis (%)	95% CI
<1000	29	1 (3.5)	0.09–17.8	14	0 (0)	0.00–19.3
<2000	54	3 (5.6)	1.16–15.4	22	0 (0)	0.00–12.7
<3000	74	7 (9.5)	3.89–18.5	29	1 (3.5)	0.09–17.2
<4000	90	10 (11.1)	5.46–19.5	37	1 (2.7)	0.07–14.2
<5000	103	11 (10.7)	5.45–18.3	40	1 (2.5)	0.06–12.9
Total	118	13 (11.0)	6.00–18.1	46	1 (2.2)	0.06–11.5

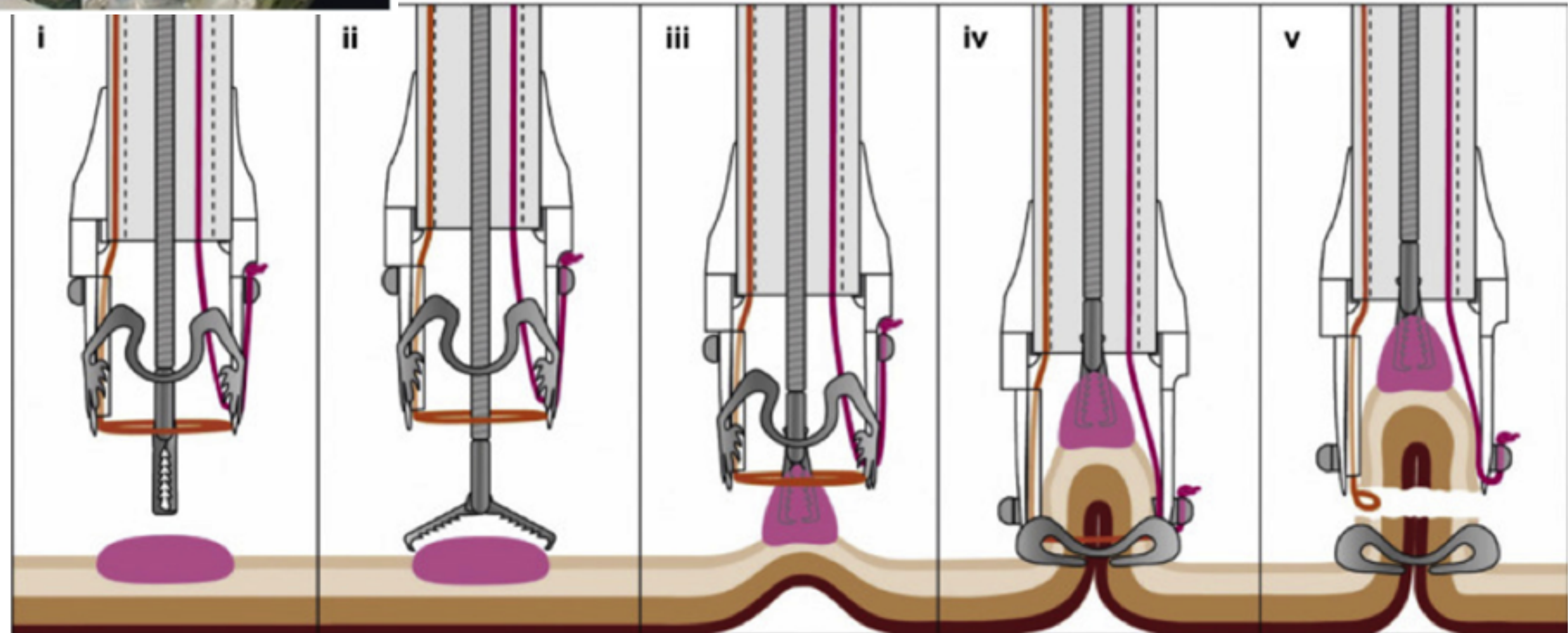
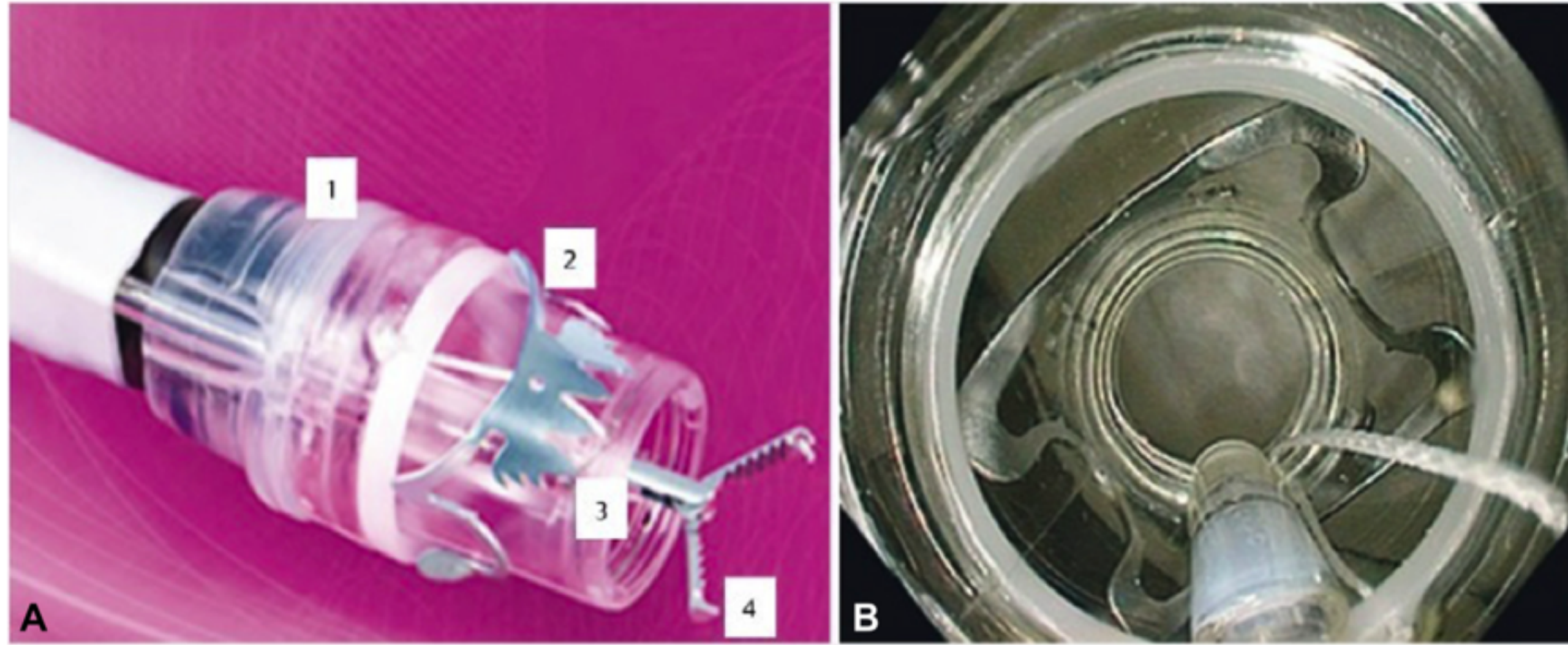
[†]*n* = 118.

[‡]Limited submucosal invasive carcinomas that were well-/moderately differentiated/papillary adenocarcinoma, with no vascular invasion and only grade 1 tumor budding.

OBTENTION D'UNE MARGE PROFONDE R0 quelque soit l'infiltration dans la sous muqueuse +++++ :

- ***Amélioration du taux de résection R0***
 - **Dissection endoscopie > Mucosectomie** quelque soit la taille +++
 - 50% des cancers T1 ont un diamètre < 20 mm
 - Etude Hollandaise sur **FTRD** en 1^e intention des cancers T1 ont un diamètre < 20 mm
 - **Dissection intermusculaire**
 - **Résection endoscopique transpariétale (TEM endoscopique)**
- ***Rattrapage d'une Marge R1 (FTRD)***

Relevance of polyp size for primary endoscopic full-thickness resection of suspected T1 colorectal cancers



Relevance of polyp size for primary endoscopic full-thickness resection of suspected T1 colorectal cancers

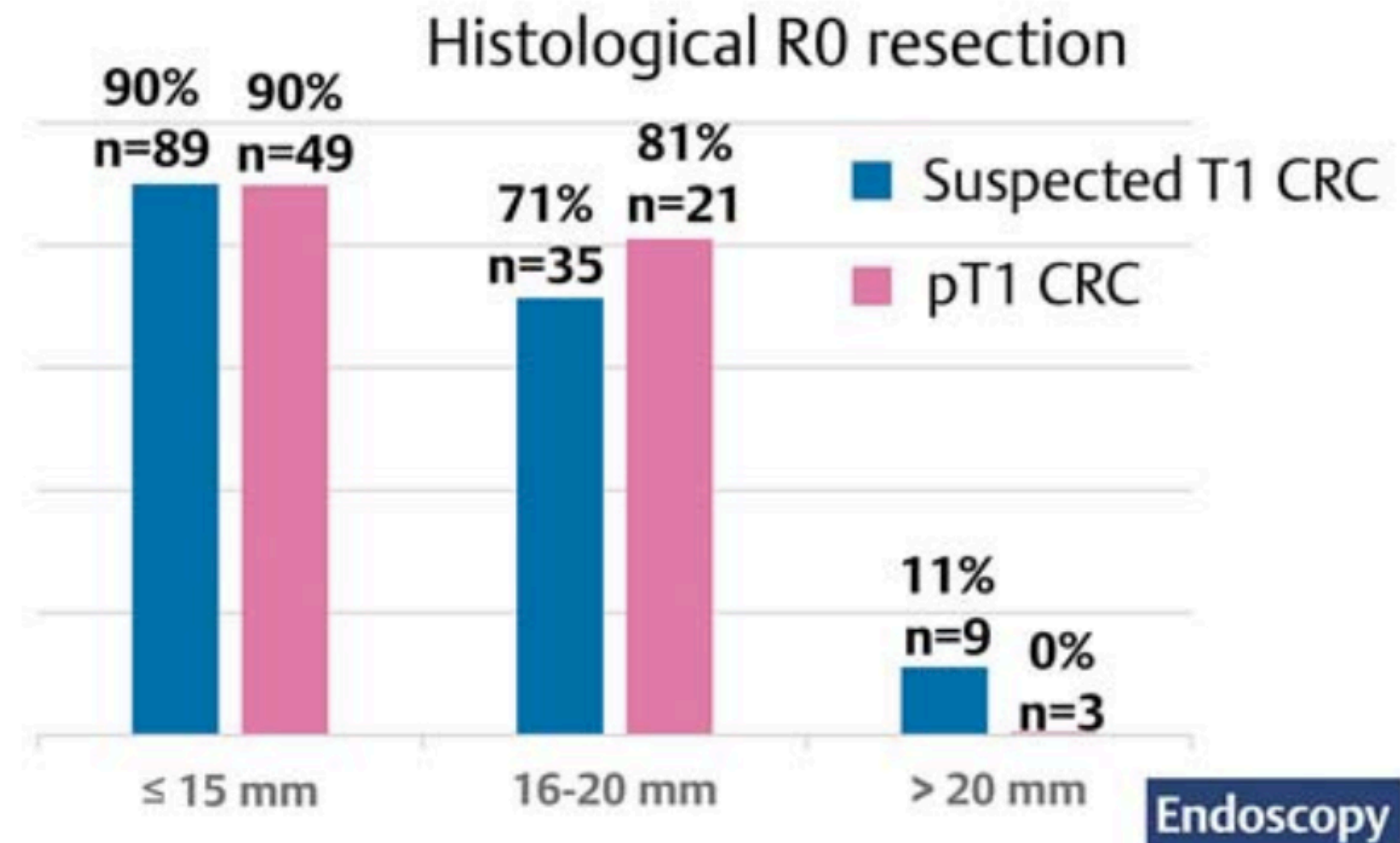
Endoscopic full-thickness resection for suspected T1 CRC

Retrospective multi-center cohort study

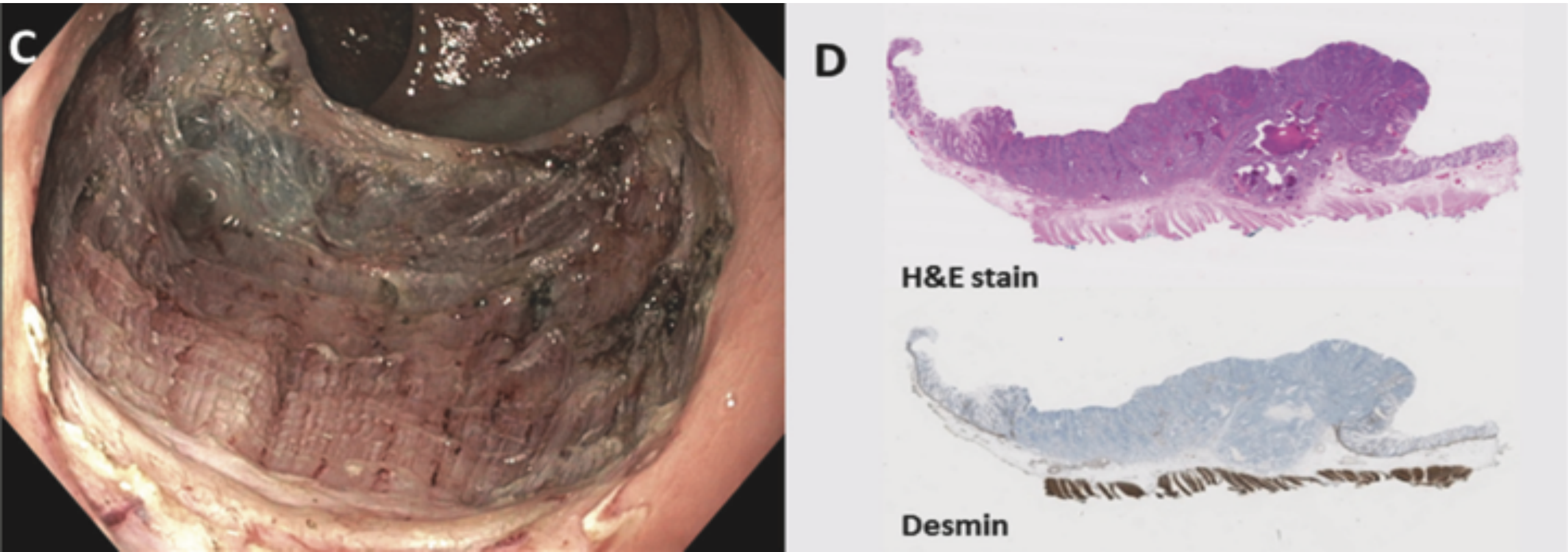
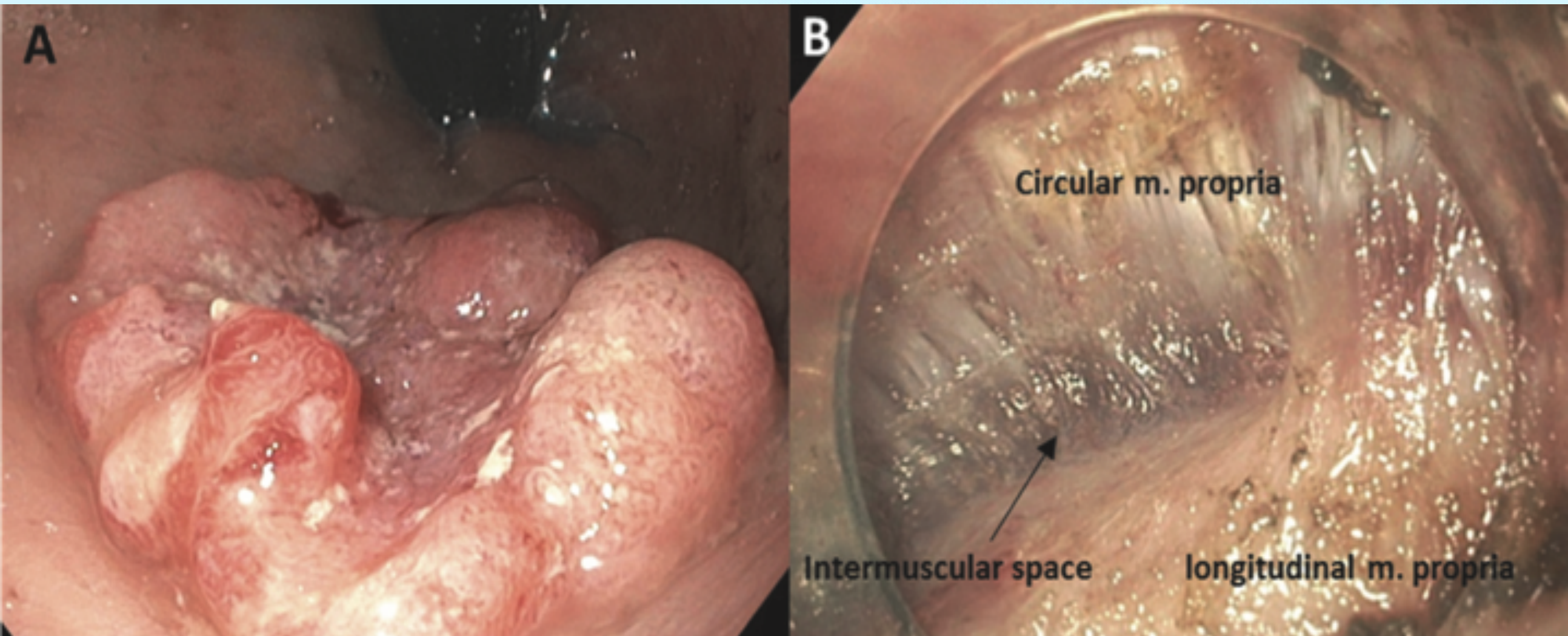
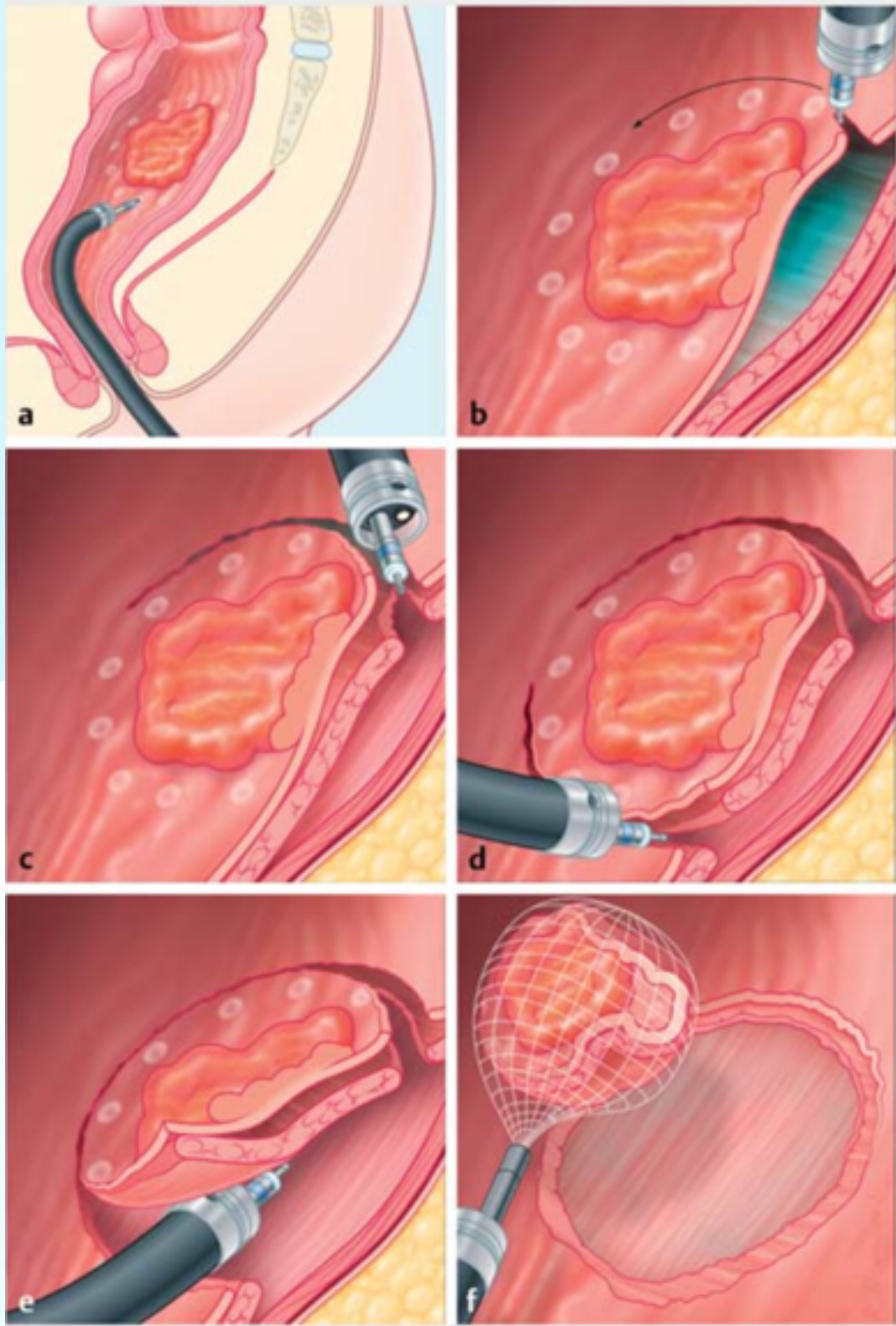
- 136 suspected T1 CRCs
- Median size 15 mm
- Included 75 pT1 CRCs

Technical success: 88%

Histological R0 resection: 80%



DISSECTION INTER MUSCULAIRE



Endoscopic intermuscular dissection for deep submucosal invasive cancer in the rectum: a new endoscopic approach

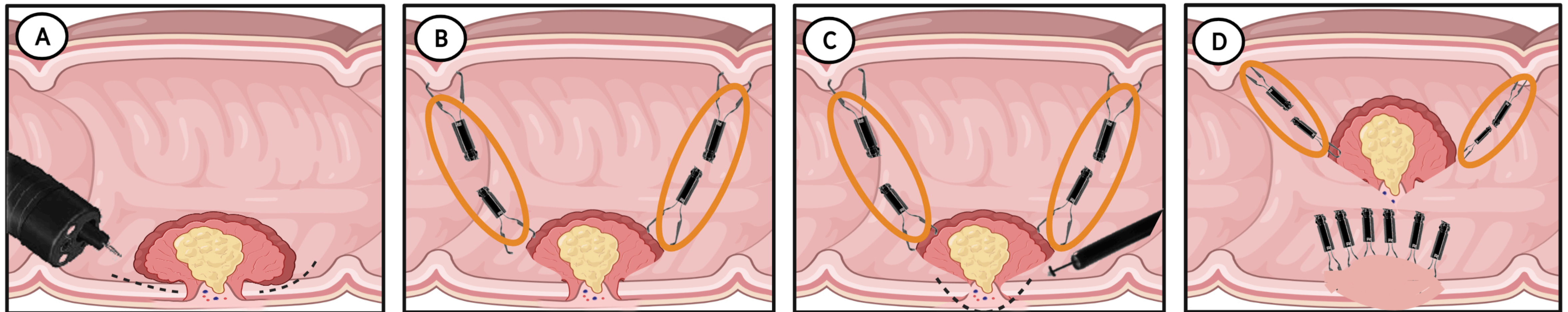
DISSECTION INTER MUSCULAIRE

Technical success		
▪ Overall	64/67 (96 %)	89 %–99 %
▪ pT1	44/45 (98 %)	90 %–100 %
▪ pT1 with deep submucosal invasion	39/40 (98 %)	89 %–100 %
R0 rate		
▪ Overall	54/67 (81 %)	70 %–89 %
▪ Technically successful cases	54/64 (84 %)	74 %–92 %
▪ pT1	41/45 (91 %)	80 %–97 %
▪ pT1 with deep submucosal invasion	36/40 (90 %)	78 %–97 %
Curative resection rate		
▪ Overall	30/67 (45 %)	33 %–57 %
▪ Technically successful cases	30/64 (47 %)	35 %–59 %
▪ pT1	22/45 (49 %)	35 %–63 %
▪ pT1 with deep submucosal invasion	18/40 (45 %)	30 %–60 %

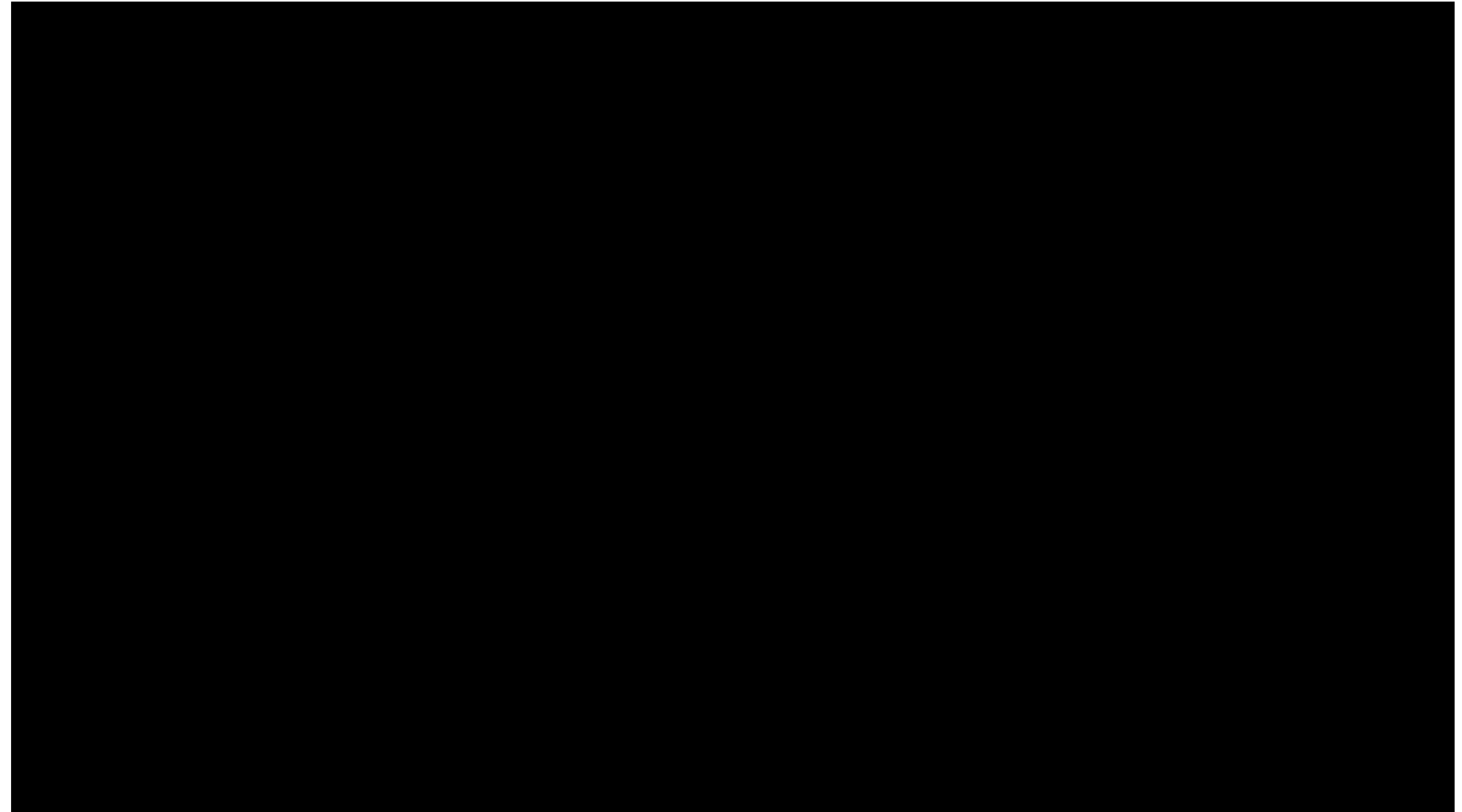
RESECTION TRANSPARIETALE ou « FLEXIBLE TEM »

2023

Graphical representation of flexible endoscope
Transanal Endoscopic Microsurgery

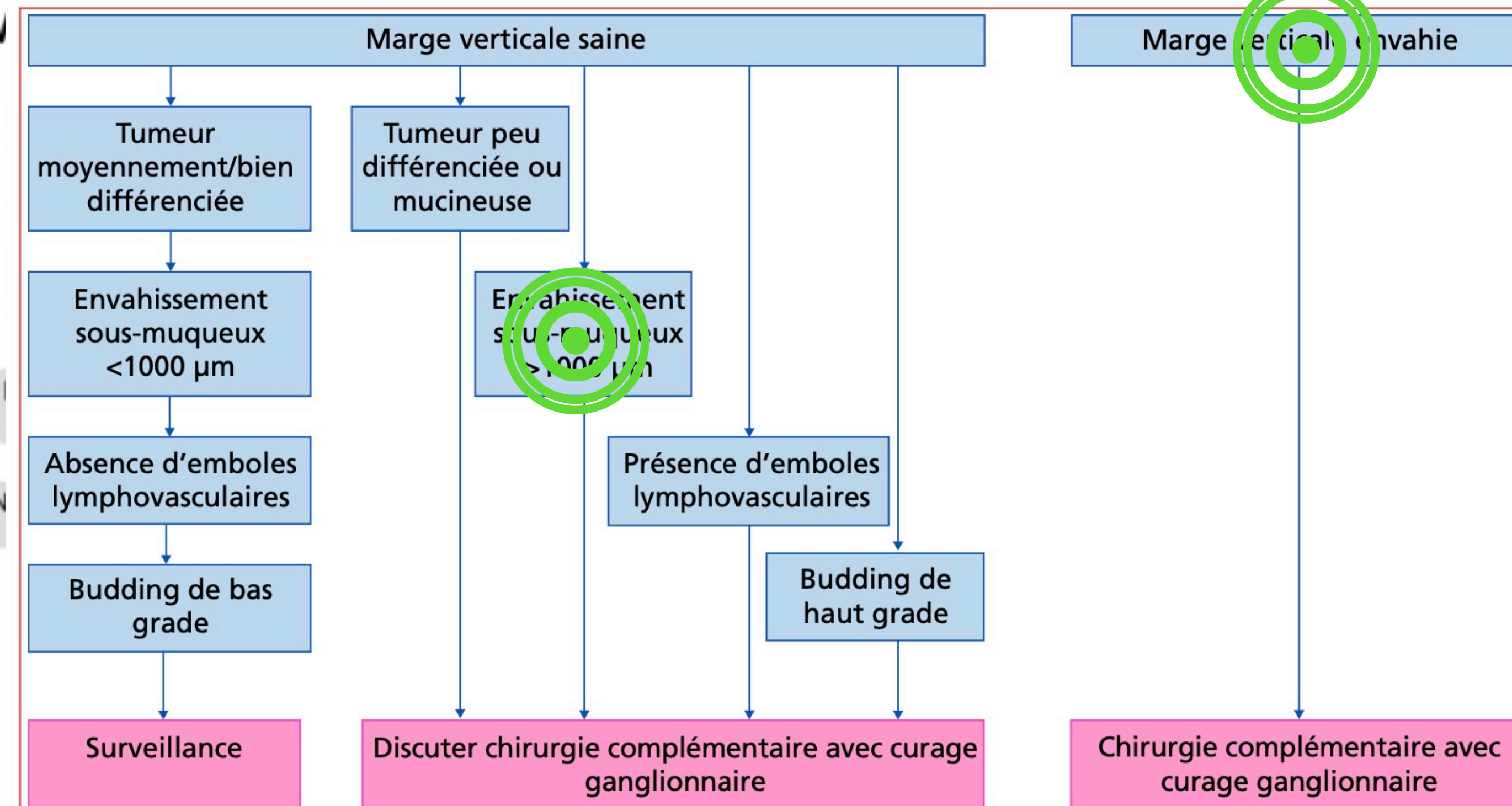


RESECTION TRANSPARIETALE ou FLEXIBLE TEM



Current problems and perspectives of pathological risk factors for lymph node metastasis in T1 colorectal cancer: Systematic review

Katsuro Ichimasa,¹ Shin-ei Kudo,¹ Hideyuki Miyachi,¹ Yuta Kouyama,¹ Kenichi Mochizuki,¹  Yuki Takashina,¹ Yasuharu Maeda,¹ Yuichi Mori,^{1,8} Toyoki Kudo,¹ Yuki Miyata,¹ Yoshika Akimoto,¹ Yuki Kataoka,^{3,4,5,6} Takafumi Kubota,^{6,7} Tetsuo Nemoto,² Fumio Ishida¹ and Masashi Misawa¹



AUTRES FACTEURS : ?

SIEGE DE LA TUMEUR

	C (<i>n</i> = 61)	A (<i>n</i> = 168)	T (<i>n</i> = 137)	D (<i>n</i> = 49)	S (<i>n</i> = 516)	R (<i>n</i> = 211)
Age (years)	68.8 ± 11.2	68.7 ± 11.5	68.6 ± 11.9	68.6 ± 12.9	64.9 ± 11.2	63.9 ± 11.5
Sex (male)	32 (52.5)	108 (64.3)	92 (67.2)	35 (71.4)	322 (62.4)	134 (63.5)
Tumor size (mm)	26.3 ± 11.4	22.8 ± 14.0	20.6 ± 11.0	20.0 ± 9.8	19.0 ± 10.8	26.2 ± 19.8
Morphology						
Flat type	32 (52.5)	88 (52.4)	74 (54.0)	26 (53.1)	108 (20.9)	80 (37.9)
Protruded type	17 (27.9)	34 (20.2)	24 (17.5)	18 (36.7)	302 (58.5)	77 (36.5)
Depressed type	12 (19.7)	46 (27.4)	39 (28.5)	5 (10.2)	106 (20.5)	54 (25.6)
Initial treatment (endoscopic resection)	38 (62.3)	101 (60.1)	86 (62.8)	34 (69.4)	352 (68.2)	124 (58.8)
Treatment (surgical resection ^a)	38 (62.3)	104 (61.9)	76 (55.5)	23 (46.9)	363 (70.3)	141 (66.8)
Depth of invasion (T1b)	43 (70.5)	119 (70.8)	88 (64.2)	29 (59.2)	401 (77.7)	178 (84.4)
Histological grade (Por or Muc ^b)	12 (19.7)	22 (13.1)	25 (18.2)	2 (4.1)	63 (12.2)	25 (11.8)
Vascular invasion (+)	13 (21.3)	43 (25.6)	33 (24.1)	5 (10.2)	129 (25.0)	82 (38.9)
Lymphatic invasion (+)	14 (23.0)	38 (22.6)	33 (24.1)	8 (16.3)	169 (32.8)	77 (36.5)
Tumor budding (BD 2 or 3)	10 (16.4)	32 (19.0)	23 (16.8)	7 (14.3)	114 (22.1)	50 (23.7)
Lymph node metastasis (+) ^c	2/38 (5.3)	4/104 (3.8)	6/76 (7.9)	1/23 (4.3)	45/363 (12.4)	17/141 (12.1)
Local lymph node recurrence (+) ^d	0/23 (0)	0/64 (0)	0/61 (0)	0/26 (0)	1/153 (0.7)	4/70 (5.7)

Lymphovascular Infiltration, Not Depth of Invasion, is the Critical Risk Factor of Metastases in Early Colorectal Cancer

Retrospective Population-based Cohort Study on Prospectively Collected Data, Including Validation

Carl-Fredrik Rönnow, MD, PhD,* Victoria Arthursson, MD,* Ervin Toth, MD, PhD,†
Peter-Martin Krarup, MD, PhD,‡ Ingvar Syk, MD, PhD,* and Henrik Thorlacius, MD, PhD*✉

TABLE 4. Risk of Lymph Node Metastases in the Absence of Any Statistically Significant Risk Factors According to the Multivariate Analysis*

	Lymph Node Metastases
No significant risk factors*	77/1053 (7.3%)
Adjusted for age	
<50 yr	8/48 (16.7%)
50–59 yr	18/123 (14.6%)
60–69 yr	15/264 (5.7%)
70–79 yr	22/391 (5.6%)
≥80 yr	14/227 (6.2%)

*Lymphovascular invasion, perineural invasion, mucinous subtype, and age.

Clinical Implications of Microsatellite Instability in T1 Colorectal Cancer

Jeonghyun Kang, Hak Woo Lee, Im-kyung Kim, Nam Kyu Kim,
Seung-Kook Sohn, and Kang Young Lee

**STATUT MSI/MSS
4-6% des CCR T1
DU RECTUM**

Table 3. Factors Associated with Lymph Node Metastasis in T1 Colorectal Carcinoma

	Univariate	
	(%)	<i>p</i> value
Microsatellite status		
MSI-H	0 /10 (0)	0.213*
MSI-L & MSS	22/123 (17.9)	

Local recurrence after local excision of early rectal cancer: a meta-analysis of completion TME, adjuvant (chemo)radiation, or no additional treatment

S. E. van Oostendorp¹, L. J. H. Smits¹ , Y. Vroom¹, R. Detering² , M. W. Heymans³, L. M. G. Moons⁴, P. J. Tanis², E. J. R. de Graaf⁵, C. Cunningham⁶, Q. Denost⁷ , M. Kusters¹ and J. B. Tuynman¹

Table 1 Weighted average local recurrence rates

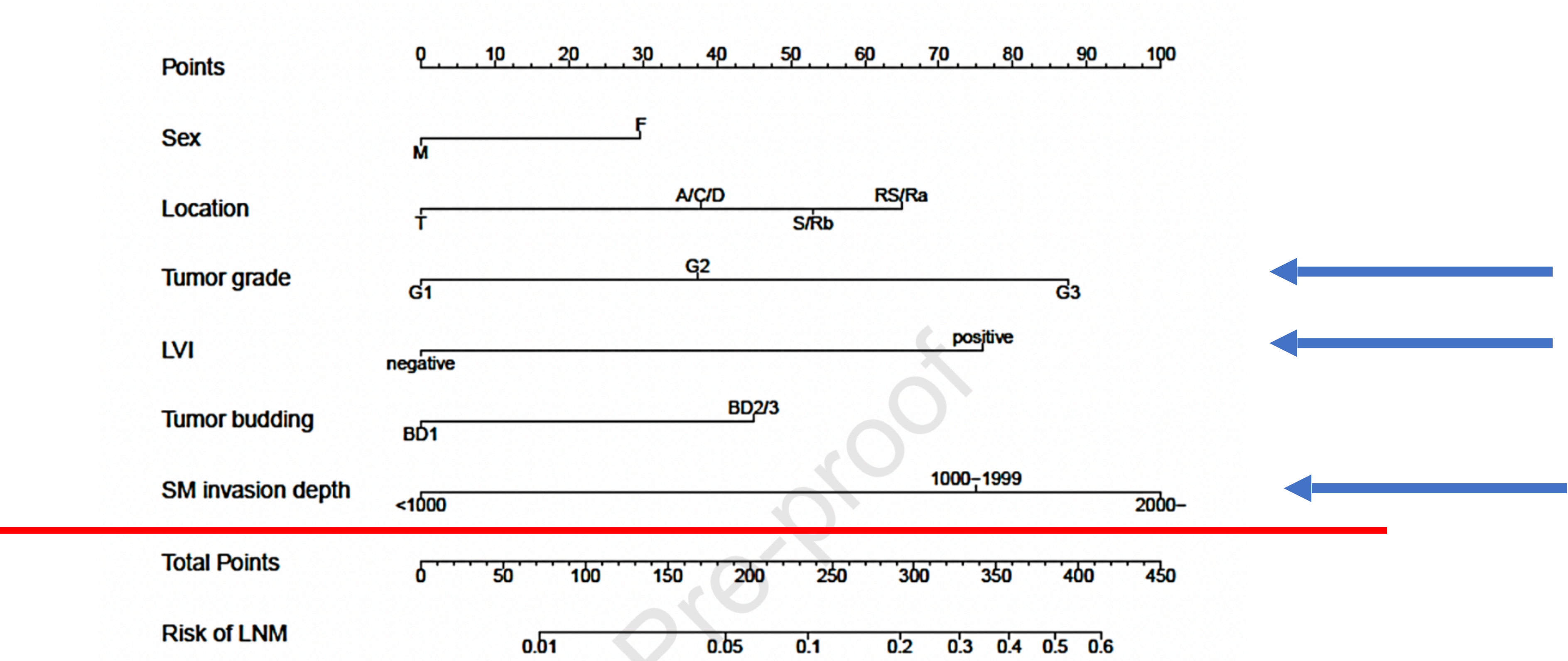
	Local recurrence					
	NAT		cTME		aCRT	
	Proportion of patients	Weighted average (%)	Proportion of patients	Weighted average (%)	Proportion of patients	Weighted average (%)
pT1	268 of 3050	8.1 (6.6, 9.9)	5 of 180	2.8 (1.2, 6.5)	24 of 385	4.8 (2.3, 9.8)
Low risk	75 of 1019	6.7 (4.8, 9.3)	0 of 28*	0	0 of 1*	0
High risk	44 of 282	13.6 (8.0, 22.0)	5 of 123	4.1 (1.7, 9.4)	10 of 254	3.9 (2.0, 7.5)
pT2	136 of 545	28.9 (22.3, 36.4)	3 of 70	4 (1, 13)	66 of 444	14.7 (11.2, 19.0)

Values in parentheses are 95 per cent confidence intervals. *Results from a single study. NAT, no additional treatment; cTME, completion total mesorectal excision; aCRT, adjuvant (chemo)radiotherapy.

CONCLUSION

- Augmentation de l'incidence des CCR T1
- Aucun examen fiable pour apprécier la profondeur de l'atteinte dans la sous muqueuse
- Nécessite d'une resection R0 avec des techniques endoscopiques avancées
- Importance de la qualité de l'examen histologique
- RCP Tumeurs superficielles

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M, male; F, female; C, cecum; A: ascending colon, T: transverse colon, D: descending colon, S: sigmoid colon, R: rectum (including rectosigmoid), RS, rectosigmoid; Ra, upper rectum; Rb, lower rectum; LVI: lymphovascular invasion; LNM, lymph node metastasis

Systematic review

Local recurrence after local excision of early rectal cancer: a meta-analysis of completion TME, adjuvant (chemo)radiation, or no additional treatment

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Table 2 Weighted average distant recurrence rates

	Distant recurrence					
	NAT		cTME		aCRT	
	Proportion of patients	Weighted average (%)	Proportion of patients	Weighted average (%)	Proportion of patients	Weighted average (%)
pT1	101 of 2658	3.4 (2.5, 4.6)	8 of 165	4.9 (2.4, 9.4)	14 of 280	5.0 (3.0, 8.3)
Low risk	25 of 783	3.2 (2.2, 4.7)	1 of 28*	4	0 of 1*	0
High risk	20 of 233	7.2 (3.6, 13.9)	6 of 108	5.6 (2.5, 11.8)	8 of 208	3.9 (1.9, 7.5)
pT2	28 of 398	6.2 (2.8, 13.0)	4 of 55	7 (3, 18)	17 of 254	5.8 (2.7, 11.9)

Values in parentheses are 95 per cent confidence intervals. *Results from a single study. NAT, no additional treatment; cTME, completion total mesorectal excision; aCRT, adjuvant (chemo)radiotherapy.