

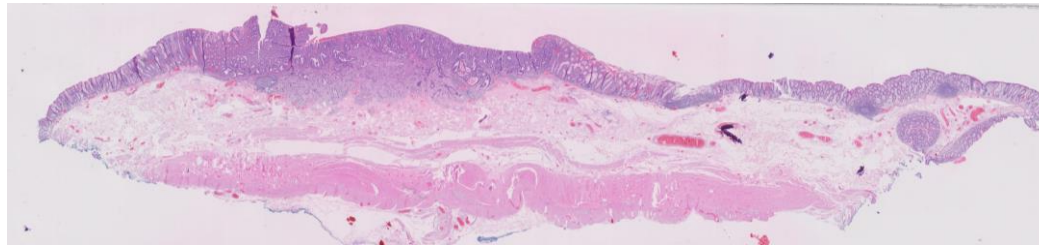
Webinar

Resection techniques in endoluminal surgery

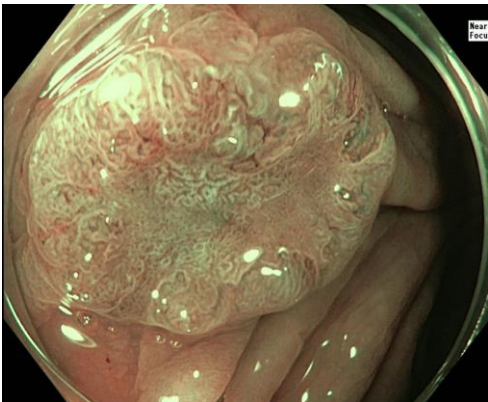


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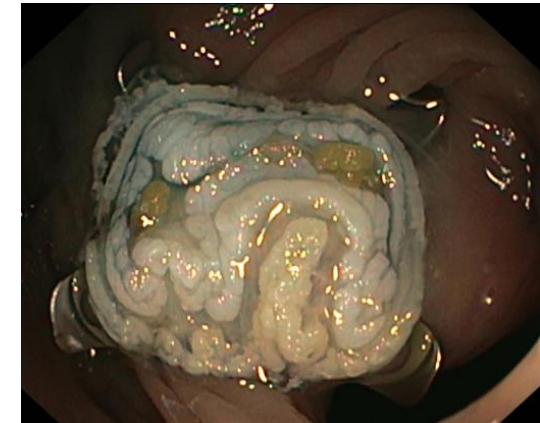
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eFTR in T1 cancers: Expanding the horizons for early colonic cancer



*Barbara Bastiaansen
Gastroenterologist
Amsterdam UMC*

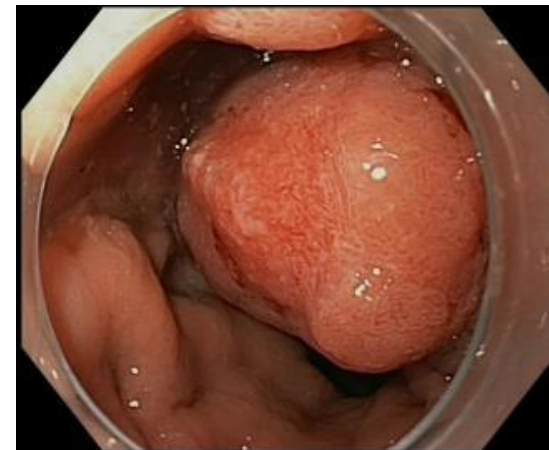
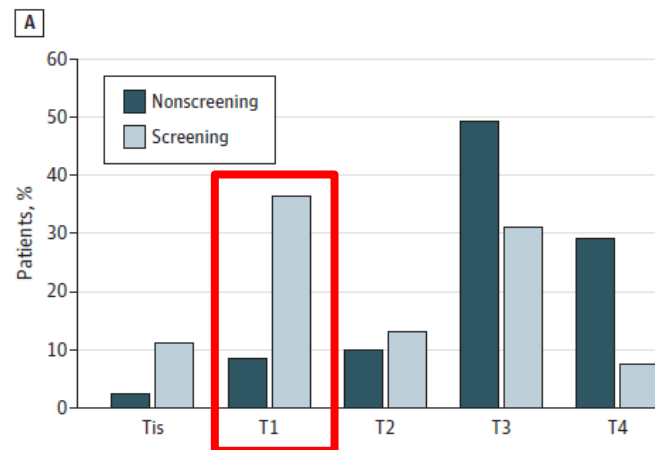




Early colorectal cancer

- Incidence is rising due to screening programmes
- 40 % of all screen detected cancers are T1
- Potential endoscopic cure!

Figure 1. Distribution of Staging



Toes Zoutendijk et al Gut 2017



Curative endoscopic Tx for T1 CRC depends on

- Radicality → *en bloc* R0 resection
- Absence of high risk features:
 - ✓ Deep submucosal invasion (i.e. $\geq 1000\mu\text{m}$, meaning Sm2-3 or Haggitt 4)
 - ✓ Lymphovascular invasion

✓
✓ **Endoscopic recognition is important!**



Endoscopic recognition of T1 CRC in practice..

Suboptimal endoscopic cancer recognition in colorectal lesions in a national bowel screening programme

Vleugels JLA, et al. Gut 2019



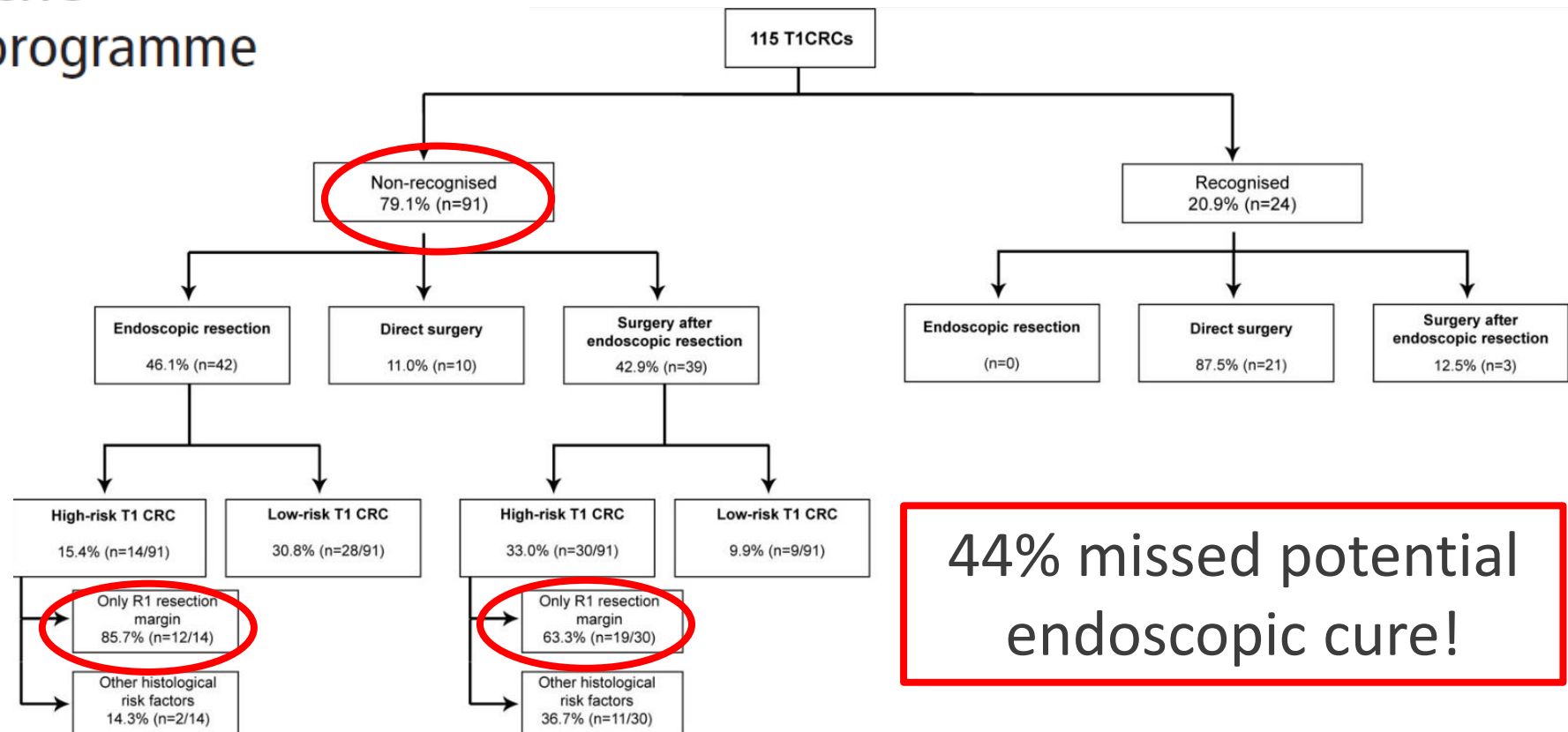
- 3622 screening colonoscopies, 274 CRC of which 91 T1 CRC
- 61% misdiagnosed as cancer
- Leading to overuse of surgery: 41 % vs 11% (in correct recognized cancers)



LETTER

Gut Month 2020 Vol 0 No 0

Optical diagnosis of T1 CRCs and treatment consequences in the Dutch CRC screening programme



Lonne W T Meulen, Gut 2020



Dilemma in T1 CRC..

Endoscopy

Vs

Surgery

- ✓ Locoregional recurrence
- ✓ Lymphatic spread
- ✓ Cancer related death



- ✓ Morbidity
- ✓ Mortality
- ✓ Functional loss

~ 90% overtreated with surgery!



Shared decision..

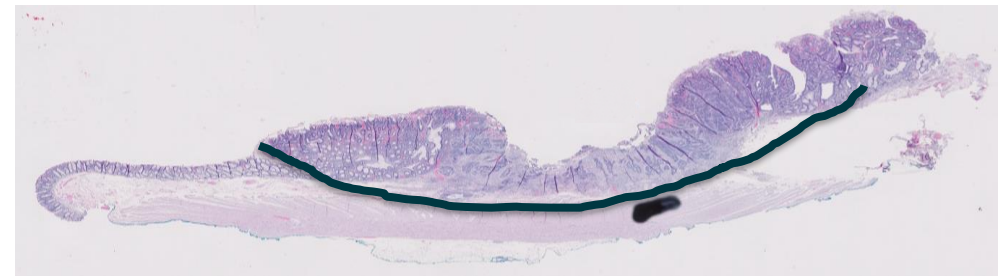
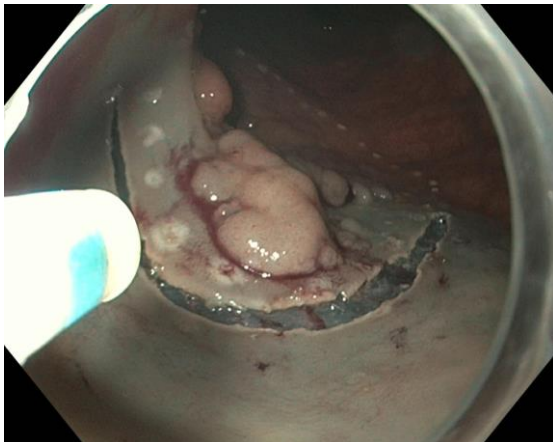




Endoscopic Submucosal Dissection

- ✓ Good histology
- ✓ R0 rate ~ 80%
- ✓ Potential cure for low risk T1CRC
- ✓ Unlimited size

- ✗ Time/resource consuming
- ✗ Difficult
- ✗ Higher complication rate
- ✗ Lack of ESD experts outside Asia
- ✗ Inappropriate for deep Sm invasive cancer





Deep submucosal invasion is NOT an independent predictor for lymphatic spread

- Risk of LNM if only deep invasion is present is 1 – 2 % ¹⁻⁵
- And the evidence is accumulating ...

| Study, year | N | Risk for LNM |
|--------------|-----|--------------|
| Suh 2012 | 118 | 2 (1,7 %) |
| Nakadoi 2012 | 249 | 3 (1,2 %) |
| Kim 2016 | 271 | 4 (1,5 %) |
| Shin 2018 | 164 | OR 0,88 |
| Yasue 2019 | 258 | 4 (1,6 %) |



Expanding the horizons..

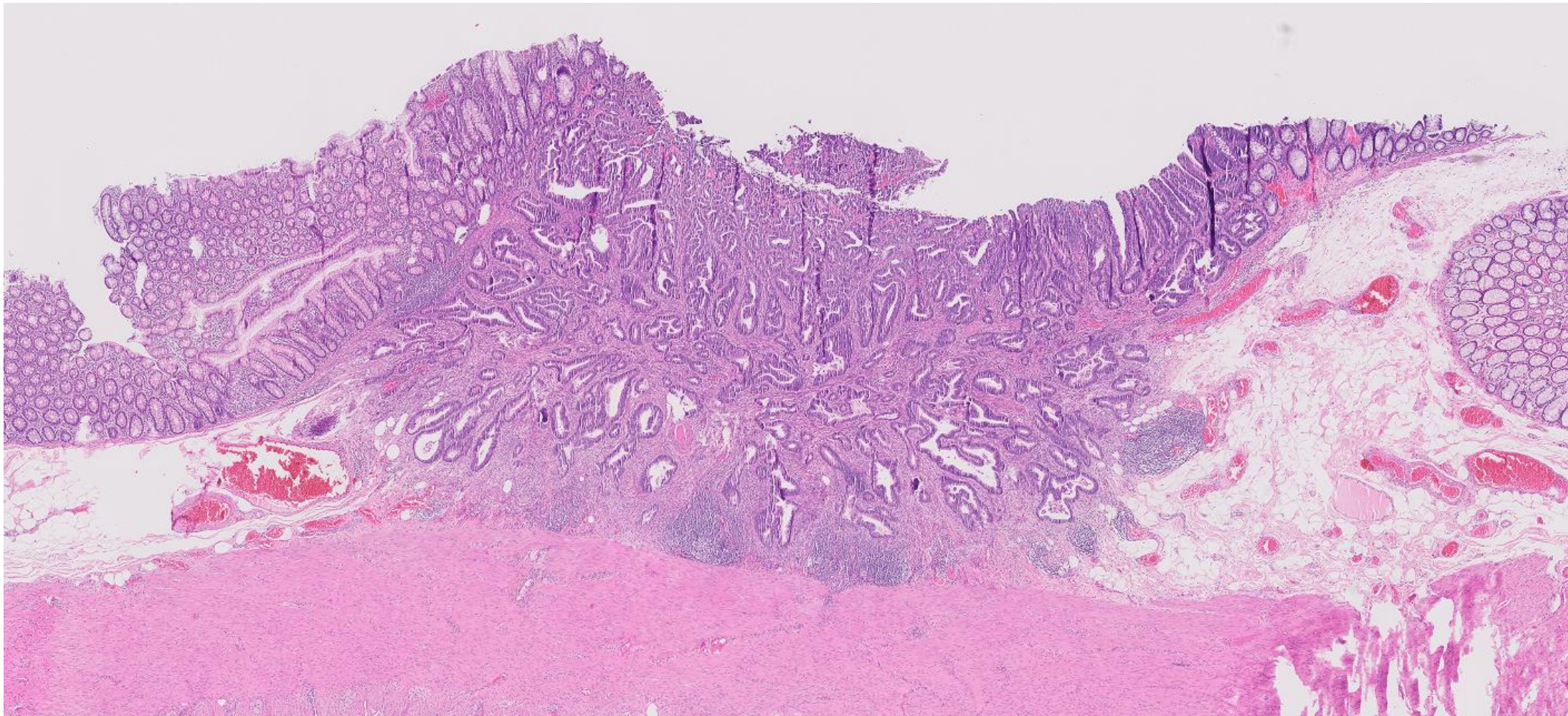
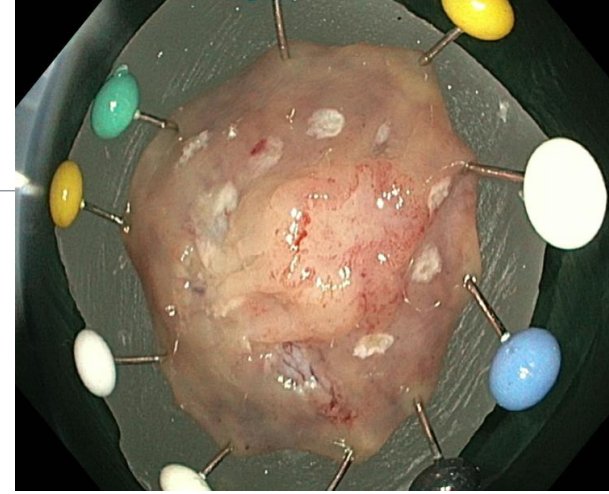




Endoscopic Full Thickness Resection (eFTR)



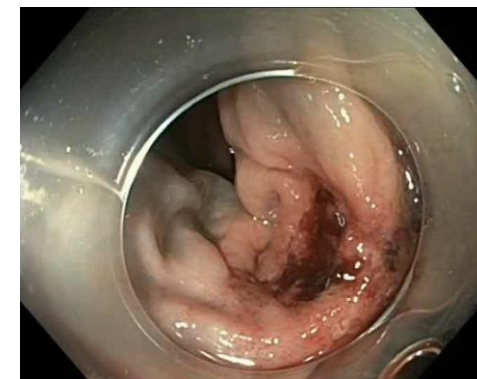
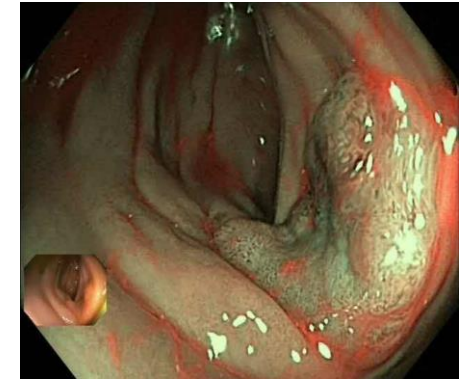
Histopathology





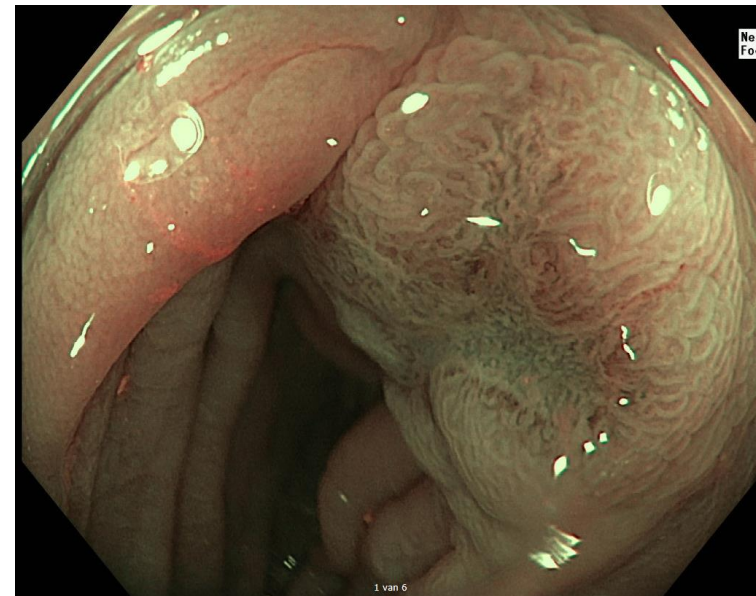
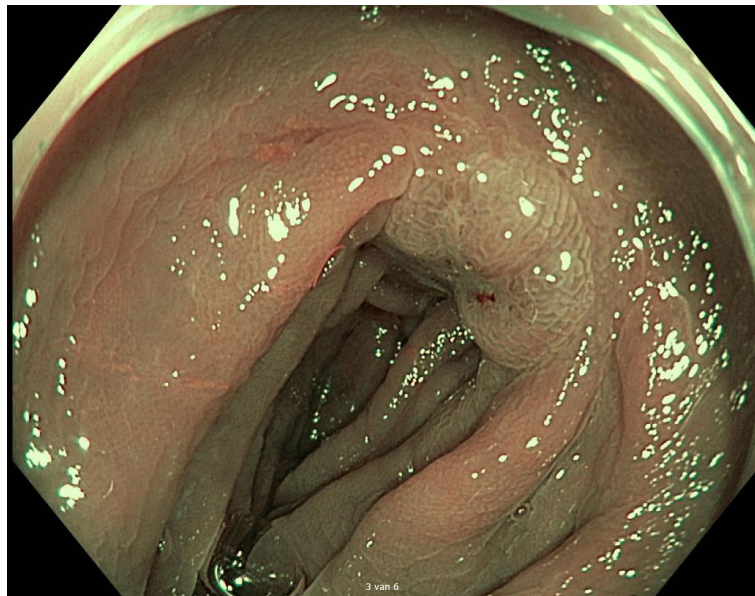
Case- 55 yo male patient

- 2005: T3N0 sigmoid carcinoma →
(open) sigmoid resection with anastomosis at 15 cm
- 2015: p-EMR adenoma HGD descending colon,
no follow up...
- 2018: Surveillance colonoscopy
 - Polyp 15 mm descending colon, partial non lifting
 - Piecemeal resection attempt in referring center
 - Distal marking tattoo
 - Histology: “at least”HGD, strong suspicion submucosal cancer



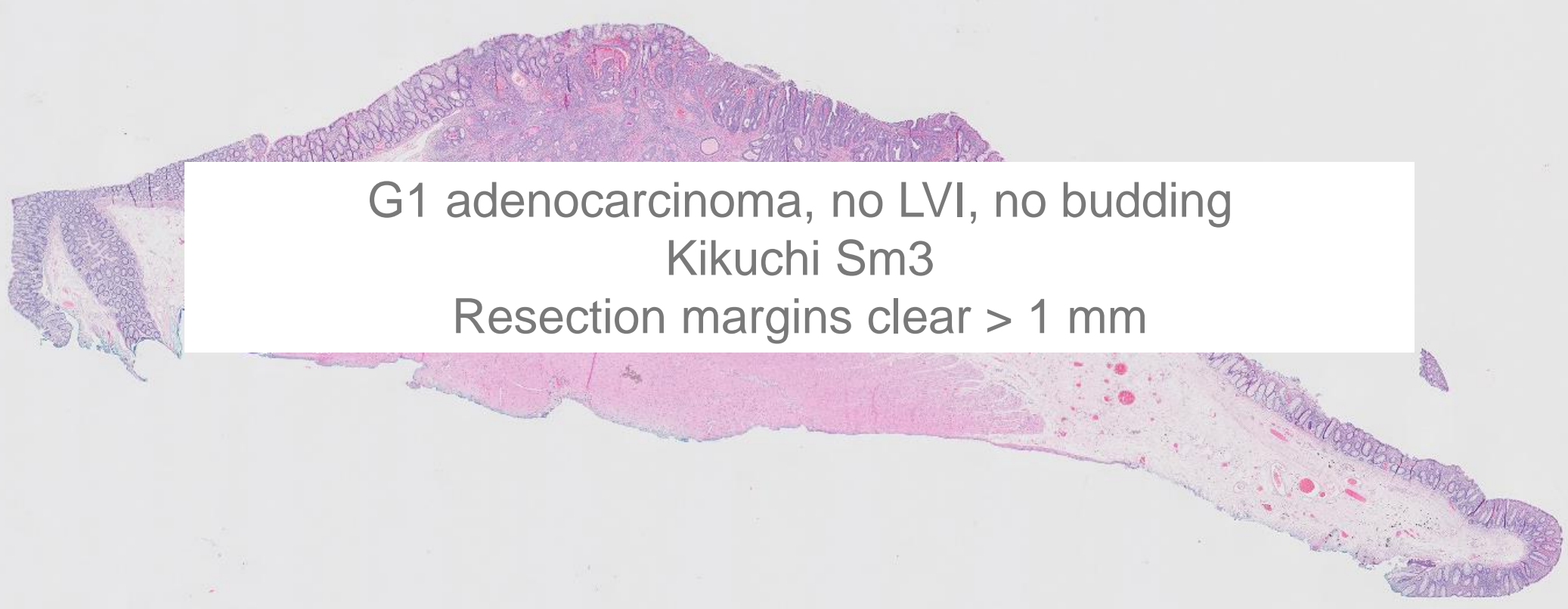


Case- 55 yo male patient





Case- 55 yo male patient



G1 adenocarcinoma, no LVI, no budding
Kikuchi Sm3
Resection margins clear > 1 mm

A long, thin, horizontal histological slide of a colon resection specimen, stained with hematoxylin and eosin (H&E). The slide shows the mucosal lining of the colon with a clear transition from normal mucosa to a tumor. A white text box is overlaid on the center of the slide, containing the following text: "G1 adenocarcinoma, no LVI, no budding", "Kikuchi Sm3", and "Resection margins clear > 1 mm".



eFTR for T1 CRC

Endoscopic full-thickness resection for early colorectal cancer

Armin Kuellmer, MD,^{1,*} Julius Mueller,^{1,*} Karel Caca, MD,² Patrick Aepli, MD,³ David Albers, MD,⁴ Brigitte Schumacher, MD,⁴ Anne Glitsch, MD,⁵ Claus Schäfer,⁶ Ingo Wallstabe, MD,⁷ Christopher Hofmann, MD,⁸ Andreas Erhardt,⁹ Benjamin Meier, MD,² Dominik Bettinger, MD,¹ Robert Thimme, MD,¹ Arthur Schmidt, MD,¹, the FTRD study group

Freiburg, Germany



Retrospective multicenter trial (96 hospitals)

1234 screened patients → n = 156

❖ group 1: re-resections after previous Rx/R1, n = 64

❖ group 2: primary non lifting, n = 92

- Endpoints: i.a. technical succes, R0, succesfull risk stratification

A. Kuellmer et al, GIE jan 2019



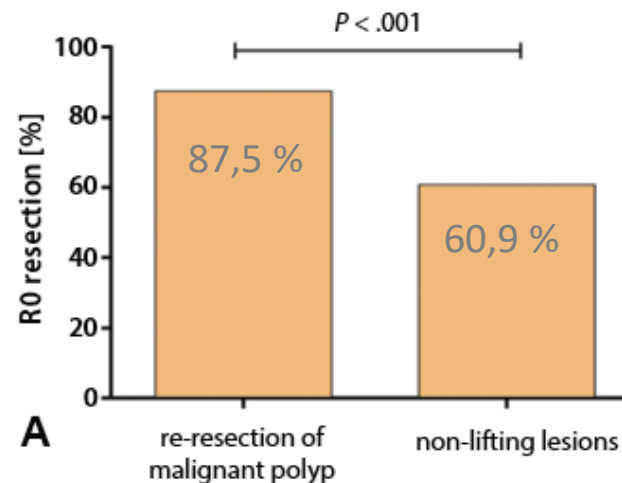
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Freiburg, Germany



- Overall R0 rate 71.8% (112 of 156)



- Exact histologic risk stratification in 99,3%!

A. Kuellmer et al, GIE jan 2019



Dutch prospective eFTR registry

- Prospective multicenter registry
- Started august 2015
- 23 participating hospitals
 - 39 certified endoscopists

- > 700 eFTR procedures
- ~350 T1 CRC related





| Procedures, total (%) | N = 331 (100) |
|------------------------------------|----------------|
| Male, n (%) | 212 (65.2) |
| Age (mean in years \pm sd) | 68.9 \pm 8.5 |
| Primary treatment | 133 (40.2) |
| Secondary treatment | 198 (59.8) |
| Median size, mm (IQR) | 15 (12 – 17) |
| Proximal (cecum – splenic flexure) | 101 (30.5) |
| Distal (descending colon – rectum) | 230 (69.5) |



| | Overall (n=331) | Primary treatment (n=133) | Secondary treatment (n=198) |
|--------------------------|----------------------------|--------------------------------------|--|
| Technical success, n (%) | 288 (87.0) | 119 (89.5) | 169 (85.4) |



| | Overall (n=321) | Primary treatment (n=129) | Secondary treatment (n=192) |
|---------------------------------|----------------------------|--------------------------------------|--|
| R0 resection, n (%) | 271 (84.4) | 103 (79.8) | 168 (87.5) |
| Full-thickness resection, n (%) | 261 (81.3) | 108 (83.7) | 153 (79.7) |



| | Overall (n=321) | Primary treatment (n=129) | Secondary treatment (n=192) |
|--------------------------------|--------------------|------------------------------|--------------------------------|
| T1 CRC, n (%) | 113 (35.1) | 98 (75.4) | 15 (7.8) |
| T2 CRC, n (%) | 23 (7.1) | 12 (9.2) | 11 (5.7) |
| Adenoma with LGD, n (%) | 15 (4.7) | 8 (6.2) | 7 (3.6) |
| Adenoma with HGD, n (%) | 10 (3.1) | 6 (4.6) | 4 (2.1) |
| Sessile serrated lesion, n (%) | 4 (1.2) | 2 (1.5) | 2 (1.0) |
| Normal scar tissue, n (%) | 151 (47.0) | 2 (1.6) | 149 (77.6) |
| Other, n (%) | 4 (1.2) | 1 (0.8) | 3 (1.6) |
| No pathology obtained, n (%) | 1 (0.3) | 0 (0) | 1 (0.5) |



| | Primary treatment (n=110) | Secondary treatment (n=26) |
|-------------------------|------------------------------|-------------------------------|
| Low-risk, n (%) | 30 (27.3) | 3 (11.5) |
| R0 resection | 25 (83.3) | 0 (0) |
| R1/Rx resection | 5 (16.7) | 3 (100) |
| Missing, n (%) | 2 (1.8) | 0 (0) |
| High-risk, n (%) | 19 (17.3) | 7 (26.9) |
| R1/Rx resection | 19 (24.4) | 7 (30.4) |
| Missing, n (%) | 2 (0.9) | 0 (0) |

Successful risk stratification in 134/136 (98.5%)

High-risk features for LNM are: poor differentiation, lymphovascular invasion, deep submucosal invasion (Sm 2-3) or tumor budding if assessed



| | Curative resection |
|---|--------------------|
| Overall, n (%) | 196/321 (60.7) |
| Only adenocarcinomas at histology | 25/136 (18.4) |
| Excluding SM2-3 as risk factor | 67/136 (49.3) |
| Primary treatment overall, n (%) | 43/129 (33.2) |
| Only adenocarcinoma at histology | 25/110 (22.7) |
| Excluding SM2-3 as risk factor | 59/110 (53.6) |
| Secondary treatment overall, n (%) | 152/192 (79.2) |
| Only adenocarcinoma at histology | 0/26 (0) |
| Excluding SM2-3 as risk factor | 8/26 (30.8) |

A curative resection is defined as a histological R0 resection and in case of residual cancer without high-risk features for lymph node metastasis



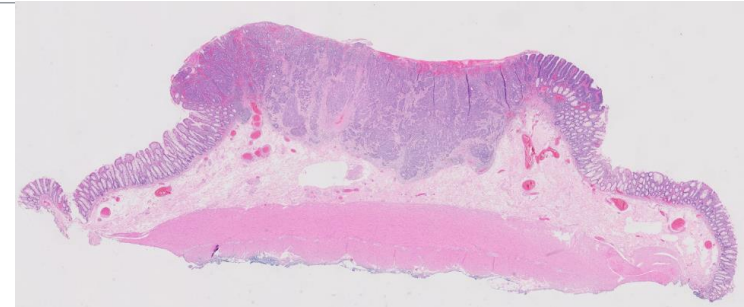
| | Overall (n=321) |
|---|----------------------------|
| Additional surgery, n (%) | 67 (20.9) |
| R1/Rx eFTR resection, n (%) | 22 (6.9) |
| One or more high-risk factors, n (%) | 29 (9.0) |
| Adverse events, n (%) | 6 (1.9) |
| Not performed (patient preference or comorbidity) | 1 (0.3) |
| Other reasons for surgery, n (%) | 9 (2.8) |
| Residual cancer | 4 (6.0) |



| | Overall (n=321) |
|--|--------------------|
| Overall | 26 (8.1) |
| <i>Mild adverse events, n (%)</i> | 13 (4.0) |
| Perforations (2 immediate / 2 delayed) | 4 (1.2) |
| Bleeding | 5 (1.6) |
| Abdominal pain | 2 (0.6) |
| Bladder retention | 8 (2.5) |
| <i>Moderate adverse events, n (%)</i> | 4 (6.1) |
| Bleeding | 6 (1.9) |
| <i>Severe adverse events, n (%)</i> | 7 (2.2) |
| Perforations (2 immediate / 5 delayed) | 7 (2.2) |



Conclusion/take home



- eFTR for T1 CRC is feasible and relative safe
 - ✓ Technical succes: 87%
 - ✓ R0 resection: 84% (80% for primary lesions)
- Delivers optimal histology and risk stratification in **>98%** cases
- eFTR could change traditional treatment paradigms and reduce the overuse of surgery:
 - ✓ R0 resection in deep invasive cancers
 - ✓ Completion treatment after previous Rx/R1 resection low risk T1 CRC

Long term oncological safety data needed !