

BÖLÜM 7

Erken evre rektal kanserler için cerrahi uygulanmadan neoadjuvan tedavi?

Çevirenler: Uz. Dr. Cihan Ağalar, Doç. Dr. Aras Emre Canda
Dokuz Eylül Üniversitesi Tıp Fakültesi Genel Cerrahi Anabilim Dalı

ANAHTAR NOKTALAR

- Kolon kanseri taraması; rektum kanseri başvuru şikâyetlerini yüksek oranda erken evre tümörlerin tanınması nedeni ile değiştirecektir.
- Küçük boyutlu tümörlerin saptanmasında ilk başvuru zamanı farklılıkları nedeniyle biyolojik olarak yavaş büyüyen tümörler ile aynı evredeki semptomatik tümörlerin tanınmalarında farklılıklar olabilir.
- Erken tanı; rektumun korunabildiği minimal invaziv tekniklerin kullanımı için seçenek sağlayabilir, böylelikle hatırı sayılır oranda morbidite ve mortalitesi bulunan geleneksel cerrahi yöntemlerden kaçınılabılır ve hastanın yaşam kalitesi artırılabilir.
- Transanal endoskopik mikrocerrahi (TEM) kullanılarak yapılan lokal eksizyonlarla, erken evre rektum kanserlerinin büyük çoğunluğu tedavi edilebilir. Fakat onkolojik etkinliği radikal cerrahi uygulanan vakalarla uyuşmamaktadır.
- Alt rektum kanseri özel olarak değerlendirilmelidir. Lateral tümör yayılım riski yüksek vakalarda klasik total mezorektal eksizyonun (TME) yapıldığı cerrahi tek ve en mantıklı strateji olacaktır.
- Ameliyat öncesi radyoterapi ile transanal endoskopik mikrocerrahinin (TEM) birlikte kullanımı cazip bir seçenek olarak görünmektedir.
 - 1 Radyoterapi ile mezorektum ve pelvik yan duvardaki mikroskobik nodal metastazlar etkili şekilde tedavi edilebilir.
 - 2 Tümör boyutunun küçülmesiyle TEM ile temiz cerrahi sınırlar elde edilmesi mümkün olmaktadır.
 - 3 Tümör evresindeki gerileme ölçümü; klinik yanıtın değerlendirilmesine göre daha objektif ve güvenilirdir.
 - 4 Histopatolojik olarak yanıtız vakalar radikal cerrahiye yönlendirilebilir.

3. Hasta klinik tam yanıt sonrası ameliyatsız tedaviye karar verdiğinde aşağıdaki ifadelerden hangisi yanlıştır?
- A. Hastaya takibin ilk yılında 6 ayda bir anestezi altında muayene yapılmalı, tümör alanı görüntülenmeli, tümör alanındaki skar dokusundan biyopsi yapılmalıdır ve her üç ayda bir pelvik MR ile değerlendirilme yapılmalıdır
- B. Ameliyatsız yaklaşımın başarısızlık şansı 5 yılda yaklaşık %10-20'dir
- C. Tümörde nüks görülmesi halinde yapılacak kurtarıcı cerrahinin uzun dönem sonuçları başlangıçta yapılacak cerrahi rezeksiyondan daha kötü değildir
- D. Tümörde nüks görülecekse kanıtlar göstermektedir ki bu olay ilk 24-48 aylık sürede olacaktır

Kaynaklar

- 1 UK Flexible Sigmoidoscopy Screening Trial Investigators. Single flexible sigmoidoscopy screening to prevent colorectal cancer: baseline findings of a UK multicentre randomised trial. *Lancet* 2002; 359(9314): 1291-300.
- 2 UK Colorectal Cancer Screening Pilot Group. Results of the first round of a demonstration pilot of screening for colorectal cancer in the United Kingdom. *BMJ* 2004; 329(7458): 133.
- 3 Endreseth BH, Myrvold HE, Romundstad P, Hestvik UE, Bjerkeset T, Wibe A. Transanal excision vs. major surgery for T1 rectal cancer. *Dis Colon Rectum* 2005; 48(7): 1380-8.
- 4 Peeters KC, Marijnen CA, Nagtegaal ID, et al. The TME trial after a median follow-up of 6 years: increased local control but no survival benefit in irradiated patients with resectable rectal carcinoma. *Ann Surg* 2007; 246(5): 693-701.
- 5 Rutten HJ, den Dulk M, Lemmens VE, van de Velde CJ, Marijnen CA. Controversies of total mesorectal excision for rectal cancer in elderly patients. *Lancet Oncol* 2008; 9(5): 494-501.
- 6 Marijnen CAM, Kapiteijn E, van de Velde CJH, et al. Cooperative Investigators of the Dutch Colorectal Cancer Group. Acute side effects and complications after short-term preoperative radiotherapy combined with total mesorectal excision in primary rectal cancer: report of a multicenter randomized trial. *J Clin Oncol* 2002; 20(3): 817-25.
- 7 Hendren SK, O'Connor BI, Liu M, et al. Prevalence of male and female sexual dysfunction is high following surgery for rectal cancer. *Ann Surg* 2005; 242(2): 212-23.
- 8 Wallner C, Lange MM, Bonsing BA, et al. Causes of fecal and urinary incontinence after total mesorectal excision for rectal cancer based on cadaveric surgery: A study from the cooperative clinical investigators of the Dutch Total Mesorectal Excision trial. *J Clin Oncol* 2008; 26(27): 4466-72.
- 9 Temple LK, Bacik J, Savatta SG, et al. The development of a validated instrument to evaluate bowel function after sphincter preserving surgery for rectal cancer. *Dis Colon Rectum* 2005; 48(7): 1353-65.
- 10 Engel J, Kerr J, Schlesinger-Raab A, Eckel R, Sauer H, Holzner D. Quality of life in rectal cancer patients: a four-year prospective study. *Ann Surg* 2003; 238(2): 203-13.

- 11 Grumann MM, Noack EM, Hoffmann IA, Schlag PM. Comparison of quality of life in patients undergoing abdominoperineal extirpation or anterior resection for rectal cancer. *Ann Surg* 2001; 233(2): 149-56.
- 12 Wilson TR, Alexander DJ. Clinical and non-clinical factors influencing postoperative health-related quality of life in patients with colorectal cancer. *Br J Surg* 2008; 95(11): 1408-15.
- 13 Kobayashi H, Ueno H, Hashiguchi Y, Mochizuki H. Distribution of lymph node metastasis is a prognostic index in patients with stage III colon cancer. *Surgery* 2006; 139(4): 516-22.
- 14 Kobayashi H, Mochizuki H, Kato T, et al. Outcomes of surgery alone for lower rectal cancer with and without pelvic sidewall dissection. *Dis Colon Rectum* 2009; 52(4): 567-76.
- 15 MERCURY Study Group. Relevance of Magnetic Resonance Imaging-detected pelvic sidewall lymph node involvement in rectal cancer. *Br J Surg* 2011, 98(12): 1798-804.
- 16 <http://www.controlled-trials.com/isrctn14422743>
- 17 <http://clinicaltrials.gov/ct2/show/NCT00427375>
- 18 Bokkerink GMJ, de Graaf EJR, Punt CJA, et al. Study Protocol The CARTS study. Chemoradiation therapy for rectal cancer in the distal rectum followed by organ-sparing transanal endoscopic microsurgery. *BMC Surg* 2011, 11. 34.
- 19 Garcia-Aguilar J, Shi Q, Thomas CR, et al. A Phase II Trial of neoadjuvant chemoradiation and local excision for T2N0 rectal cancer: preliminary results of the ACOSOG Z6041 trial. *Ann Surg Oncol* 2012; 9(2): 384-91.
- 20 Janjan NA, Crane C, Feig BW et al. Improved overall survival among responders to preoperative chemoradiation for locally advanced rectal cancer. *Am J Clin Oncol* 2001, 24(2): 107-12.
- 21 Hiotis SP, Weber SM, Cohen AM, et al. Assessing the predictive value of clinical complete response to neoadjuvant therapy for rectal cancer: an analysis of 488 patients. *J Am Coll Surg* 2002; 194: 131-36.
- 22 Habr-Gama AP, Perez RO, Nadalin W. Operative versus non-operative treatment for stage 0 distal rectal cancer following chemoradiation therapy. *Ann Surg* 2004; 240: 711-8.
- 23 Habr-Gama A. Assessment and management of the complete clinical response of rectal cancer to chemoradiotherapy. *Colorectal Dis* 2006; 8(suppl.1 3): 21-4.
- 24 Dalton R, Velineni R, Osborne M, et al. A single-centre experience of chemoradiotherapy for rectal cancer: is there potential for non-operative management? *Colorectal Dis* 2012; 14: 567-71.
- 25 Maas M, Beets-Tan RG, Lambregts DM, et al. Wait-and-see policy for clinical complete responders after chemoradiation for rectal cancer. *J Clin Oncol* 2011; 29: 4633-40.
- 26 Glynne-Jones R, Hughes R. Critical appraisal of the 'wait and see' approach in rectal cancer for clinical complete responders after chemoradiation. *Br J Surg* 2012; 99: 897-909.
- 27 [Online]: <http://public.ukcrn.org.uk/Search/StudyDetail.aspx?StudyID=8565>
- 28 Maas M, Nelemans PJ, Valentini V et al. Long-term outcome in patients with a pathological complete response after chemoradiation for rectal cancer. *Lancet Oncol* 2010; 11. 835-44.
- 29 Rodel C, Martus P, Papadopoulos T, et al. Prognostic significance of tumor regression after preoperative chemoradiotherapy for rectal cancer. *J Clin Oncol* 2005; 23: 8688-96.

- 30 Wiltshire KL, Ward IG, Swallow C, et al. Preoperative radiation with concurrent chemotherapy for resectable rectal cancer: effect of dose escalation on pathologic complete response, local recurrence-free survival, disease-free survival, and overall survival. *Int J Radiat Oncol Biol Phys* 2006; 64(3): 709-16.
- 31 Bosset JF, Collette L, Calais G, et al. Chemotherapy with preoperative radiotherapy in rectal cancer. *N Engl J Med* 2006; 355: 1114-23.
- 32 [Online]: <http://www.controlled-trials.com/ISRCTN093514471>
- 33 [Online]: <http://clinicaltrials.gov/ct2/show/NCT01037049>
- 34 Collins S, Renehan A, Saunders M, Scott N, Susnerwala S, Sun Myint A. Rectal Cancer apparent complete response (aCR). Colorectal Clinical Subgroup, National Cancer Research Institute. Available from: http://www.gmccn.nhs.uk/hp/portal_repository/files/RectalCancerApparentCompleteResponseafterChemoradiotherapy.pdf
- 35 Di Fabio F, Pinto C, Fanti S, et al. Correlation between FDG-PET and pathologic response in patients with rectal cancer treated with neoadjuvant chemo-radiotherapy: First results of the Bologna Project. *Proc Am Soc Clin Oncol* 2005; 23: (abstr 3623).
- 36 Martoni AA, Di Fabio F, Pinto C, et al. Prospective study on the FDG-PET/CT predictive and prognostic values in patients treated with neoadjuvant chemoradiation therapy and radical surgery for locally advanced rectal cancer. *Ann Oncol* 2011, 22(3): 650-6.
- 37 Rutten H, Sebag-Montefiore D, Glynne-Jones R, et al. Capecitabine, oxaliplatin, radiotherapy, and excision (CORE) in patients with MRI-defined locally advanced rectal adenocarcinoma: Results of an international multicenter phase II study. *Proc Am Soc Clin Oncol* 2006; 24: (abstr 3528).
- 38 Whiteway J, Nicholls RJ, Morson BC. The role of surgical local excision in the treatment of rectal cancer. *Br J Surg* 1985 Sep; 72(9): 694-7
- 39 Garcia-Aguilar J, Mellgren A, Sirivongs P, et al. Local excision of rectal cancer without adjuvant therapy a word of caution. *Ann Surg* 2000; 231(3): 345-51.
- 40 Buess G, Hutterer F, Theiss J, Bobel M, Isselhard W, Pichlmaier H. A system for a transanal endoscopic rectum operation. *Chirurg* 1984; 55: 677-80.
- 41 Buess G, Theiss R, Hutterer F, et al. Transanal endoscopic surgery of the rectum - testing a new method in animal experiments. *Leber Magen Darm* 1983; 13: 73-7
- 42 Bach SP, Hill J, Monson JR, et al. A predictive model for local recurrence after transanal endoscopic microsurgery for rectal cancer. *Br J Surg* 2009; 96(3): 280-90.
- 43 Leong KJ, Wei W, Tannahill LA, et al. Methylation profiling of rectal cancer identifies novel markers of early-stage disease. *Br J Surg* 2011, 98: 724-34.
- 44 Lezoche E, Guerrieri M, Paganini AM, et al. Long-term results in patients with T2-3 NO distal renal cancer undergoing radiotherapy before transanal endoscopic microsurgery. *Br J Surg* 2005; 92(12): 1546-52.
- 45 Bujko K, Richter P, Kolodziejczyk M, et al. Preoperative radiotherapy and local excision of rectal cancer with immediate radical re-operation for poor responders. *Radiation Oncol* 2009; 92(2): 195-201.
- 46 Lezoche E, Guerrieri M, Paganini AM, et al. Transanal endoscopic versus total mesorectal laparoscopic resections of T2-N0 low rectal cancers after neoadjuvant treatment: a prospective randomized trial with a 3-years minimum follow-up period. *Surg Endosc* 2005; 19(6): 751-6.
- 47 Borschitz T, Wachtlin D, Mohler M, et al. Neoadjuvant chemoradiation and local excision for T2-3 rectal cancer. *Ann Surg Oncol* 2008; 15(3): 712-20.

- 48 Bujko K, Nowacki MP, Nasierowska-Guttmejer A, Michalski W, Bebenek M, Kryj M. Long-term results of a randomized trial comparing preoperative short-course radiotherapy with preoperative conventionally fractionated chemoradiation for rectal cancer. *Br J Surg* 2006; 93(10): 1215-23.
- 49 Sauer R, Becker H, Hohenberger W, et al. Preoperative versus postoperative chemoradiotherapy for rectal cancer. *N Engl J Med* 2004; 351: 1731-40.
- 50 Pettersson D, Cedermark B, Holm T, Radu C, Pahlman L, Glimelius B, et al. Interim analysis of the Stockholm III trial of preoperative radiotherapy regimens for rectal cancer. *Br J Surg* 2010; 97(4): 580-7
- 51 Bujko K, Richter P, Kolodziejczyk M, et al. Preoperative radiotherapy and local excision of rectal cancer with immediate radical re-operation for poor responders. *Radiother Oncol* 2009; 16; 92(2): 195-201.
- 52 Graf W, Dahlberg M, Osman MM, Holmberg L, Pahlman L, Glimelius B. Short-term preoperative radiotherapy results in down-staging of rectal cancer: a study of 1316 patients. *Radiother Oncol* 1997; 43(2): 133-7
- 53 Kundel Y, Brenner R, Purim O, et al. Is local excision after complete pathological response to neoadjuvant chemoradiation for rectal cancer an acceptable treatment option? *Dis Colon Rectum* 2010; 53: 1624-31.
- 54 Lamarque PL, Gross CG. La radiothérapie de contact des cancer du rectum. *J Radiol Elec-trol* 1946; 27- 333-346.
- 55 Papillon J. Present status of radiation therapy in the conservative management of rectal cancer. *Radiother Oncol* 1990; 17- 275-83.
- 56 Sun Myint A, Grievey RJ, McDonaldz AC, et al. Combined modality treatment of early rectal cancer the UK experience. *Clinical Oncology* 2007- 19: 674-81.
- 57 Higgins KA, Willett CG, Czito BG. Non-operative management of rectal cancer: current perspectives. *Clin Colo Cancer* 2010; 9(2): 83-8.
- 58 Gerard J, Ayzac L, Coquard R, et al. Endocavitary irradiation for early rectal carcinomas T1 (T2). A series of 101 patients treated with the Papillon's technique. *Int J Radiat Oncol BiolPhys* 1996; 4: 775-83.
- 59 The Association of Coloproctology of Great Britain and Ireland. *Guidelines for the Management of Colorectal Cancer 3rd edn (2007)*. Available at: http://www.acpghi.org.uk/assets/documents/ICOLO_guides.pdf
- 60 Gerard J, Romestaing P, Chapet O. Radiotherapy alone in the curative treatment of rectal carcinoma. *Lancet Oncol* 2003; 4: 158-66.
- 61 Gerard JP, Chapet O, Nemoz C. Improved sphincter preservation in low rectal cancer with high-dose preoperative radiotherapy: the Lyon R96-02 randomized trial. *J Clin Oncol* 2004; 22: 2404-9.
- 62 Lindegaard J, Gerard JP, Sun Myint A, Myerson R, Thomsen H, Laurberg S. Whither papillon? Future directions for contact radiotherapy in rectal cancer. *Clin Oncol* 2007-19(9): 738-11.

ÇOKTAN SEÇMELİ SORULARIN CEVAPLARI

1. D
2. A,C
3. D