

Colorectal Cancer



https://www.uclahealth.org/physiciansupdate/campaign-aims-to-boost-rates-of-colon-cancer-screening



Overview

- History
- Anatomy and Function
- Colorectal Cancer Overview
 - Term Classification, Causes
- Signs and Symptoms
- Diagnosis
- Treatment
- Risk Factors and Prevention



Awareness Ribbon and Month

March

Dark blue



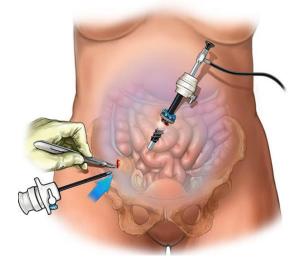


History

- 1739 Jean Faget
 - First rectal resection
 - Fatal outcome
- 1793 F. Duret
 - First colon stoma after rectal resection
 - Patient lived another 45 years
- 1879 "Hartmann Surgery"
 - No anal canal resection
 - Discovery: patients recovered much faster
- 1991 "Jacobs Laparoscopy"
 - Laparoscopical rectal resection



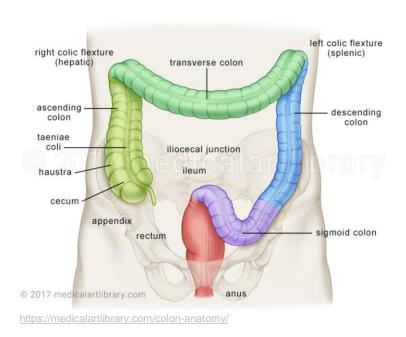
https://www.apotheken-umschau.de/Darm/Stoma---was-bedeutet-das-549821.html



https://www.healthdirect.gov.au/laparoscopy



Anatomy and Function



- Large intestine (~1.8m)
 - Cecum
 - Colon
 - Rectum
 - Anal canal

Peristalsis

- Bowel movement
- Get rid of food left over after nutrients are removed



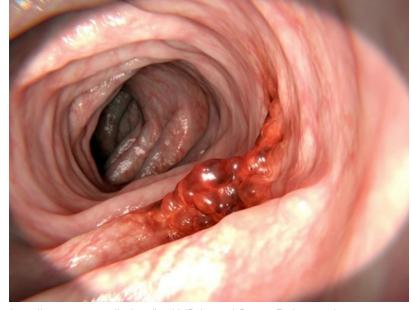
CRC – Overview

Localization frequency

- 50% rectum
- 30% sigmoid colon

Origin of CRC

- benign polyps (90%)
- Could take years
- Lifestyle factors



https://www.news-medical.net/health/Colorectal-Cancer-Pathogenesis.aspx



Term Classifications

Polyp

 small clump of cells that forms on the lining of the colon

Adenoma

glandular proliferation

Polyps Polyps O Mayo Foundation for Medical Education and Research. All rights reserved.

Small polyps



Large polyp

Tumor

too fast cell growth

Carcinoma

 cancer type developing from epithelial cells (malignant tumor)

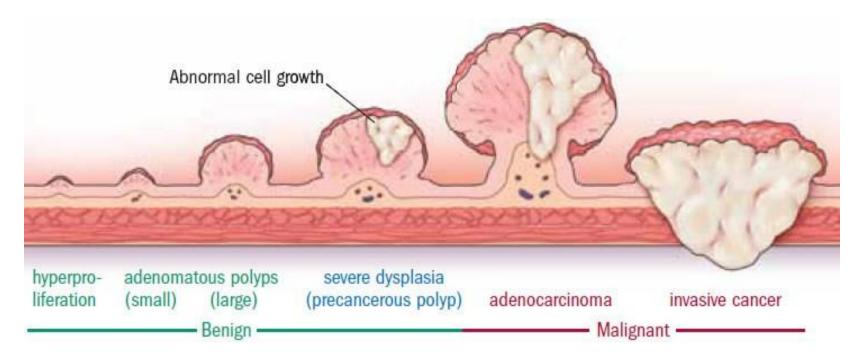


Colon cancer

https://www.mayoclinic.org/diseases-conditions/colon-polyps/symptoms-causes/syc-2035287



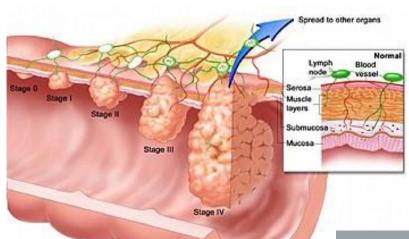
Adenoma-Carcinoma-Sequence



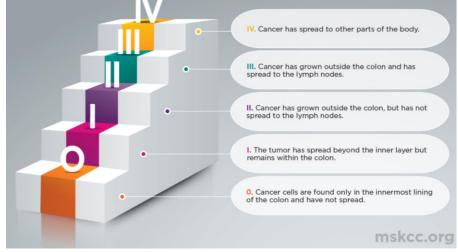
https://www.health.harvard.edu/diseases-and-conditions/they-found-colon-polyps-now-what



Stages of CRC



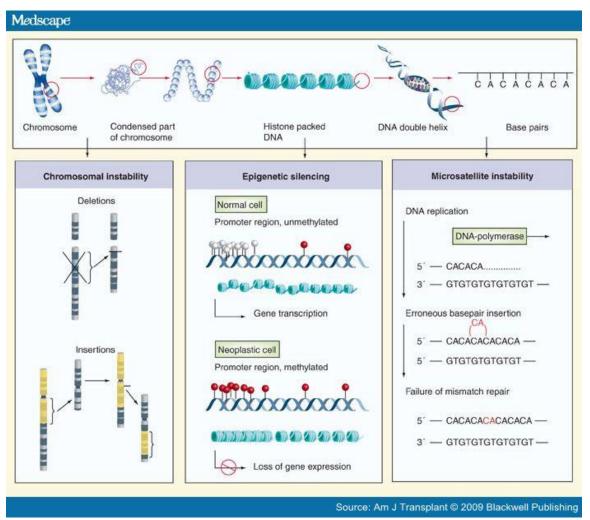
https://healthykcmag.com/colorectal-cancer-awareness-month/



https://www.mskcc.org/cancer-care/types/colon/diagnosis/stages



Cause: Gene Mutations



CIN (Chromosomal INstability)

 Mutations in genes causing duplication or deletion of entire chromosomes

Epigenetic Silencing

- "Hypermethylation"
- Epigenetic inactivation of genes (by methylating the genes' promoter region)

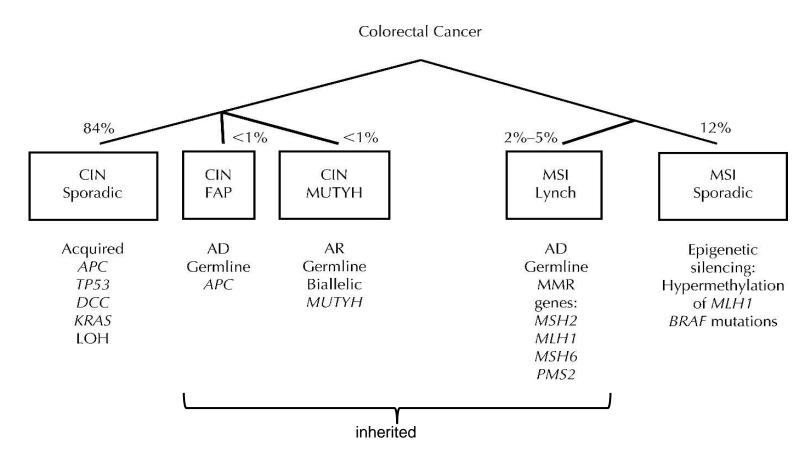
MSI (MicroSatellite Instability)

 Mutations in mismatch-repair (MMR-) genes leading to mutations in microsatellite DNA sequences

https://img.medscapestatic.com/article/703/573/703573-fig5.jp



Cause: Gene Mutations



https://www.archivesofpathology.org/doi/full/10.1043/2010-0613-RAIR.1



Example MSI: Sporadic vs. Lynch Syndrome

- Sporadic (12%)
 - Tendency: older patients
 - Epigentic silencing (MLH1)
 - BRAF↑

- Lynch Syndrome (2-5%)
 - Germline mutation in one of the mismatch repair (MMR) genes
 - 90%: MLH1 or MSH2
 - No BRAF mutation



Signs and Symptoms

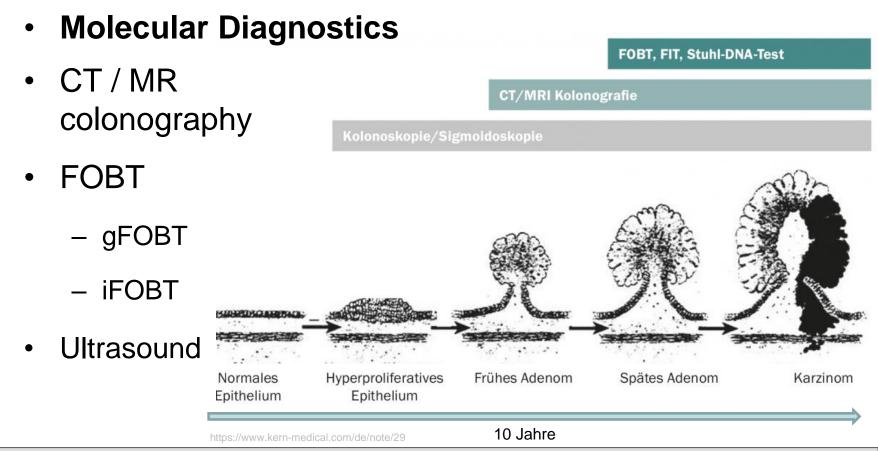
Problem: No obvious early symptoms → Screening!

- Change in bowel habits
- Blood in stool or rectal bleeding
- Abdominal pain
- Fatigue
- Fever
- Ileus
- Unexplained weight loss
- Pallor and anemia



Diagnosis

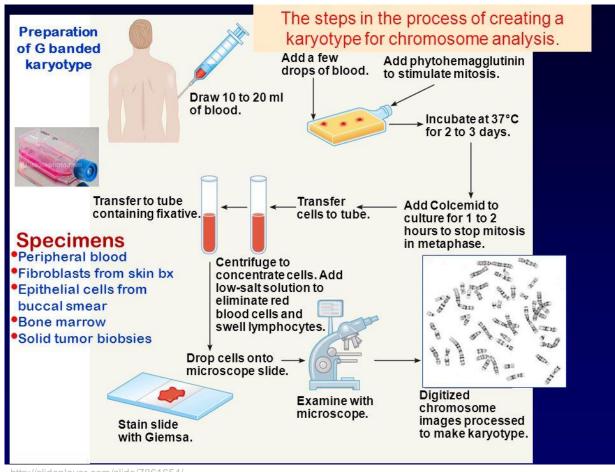
Colonoscopy + Biopsy (most sensitive and accurate)





Molecular Diagnostics

Chromosomal Instability (CIN):



Microsatellite Instability (MSI):

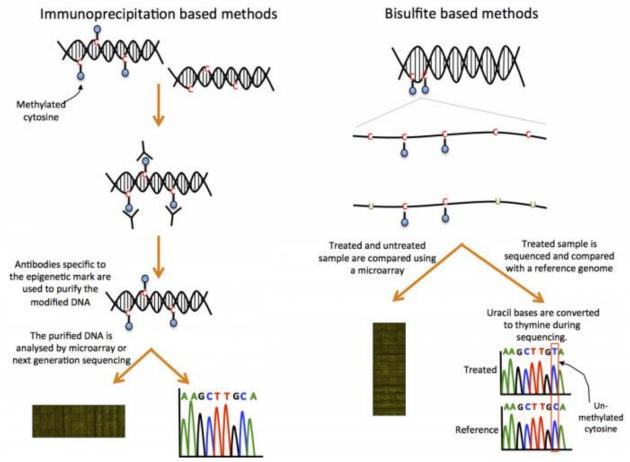
- PCR
- Sequencing **Techniques**

http://slideplayer.com/slide/7861654/



Molecular Diagnostics

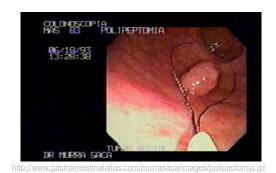
Epigenetic Silencing (DNA Methylation):



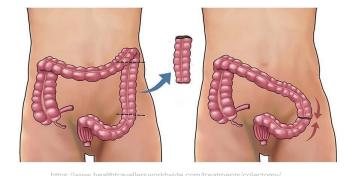
https://www.ebi.ac.uk/training/online/course/functional-genomics-ii-common-technologies-and-data-analysis-methods/epigenetic-modifications



Treatment



Loop resection



Colectomy



Chemotherapy



https://www.amazon.in/Mintkraf Yellow-Adhesive-Ionizing-Radiation/dp/R07RHGVFRF

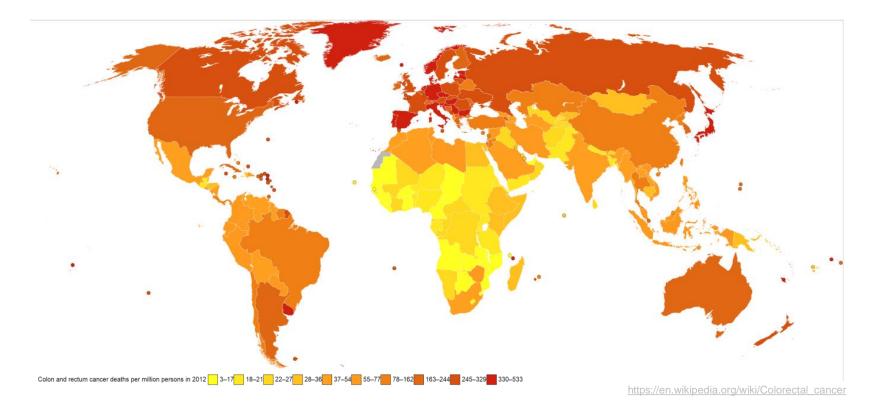
Radiation Therapy



Incidence and Mortality

Incidence men (Austria)	2.500
Incidence women (Austria)	1.800
Mortality (Austria)	2.150
Incidence (Germany)	60.000

Median age of disease: 75 years





Risk Factors



Unhealthy Food





IBD*



Age





Obesity



Inheritance



Diabetes



Alcohol



Smoking

*Inflammatory Bowel Desease



Prevention



Healthy Diet



Supplements



Water



Exercise



No red meat



Screening



No alcoho



No smoking



makeameme.o



Sources

- https://medicalartlibrary.com/colon-anatomy/
- https://www.darmkrebs-in-der-familie.de/node/25
- https://flexikon.doccheck.com/de/Kolorektales_Karzinom
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- https://www.awmf.org/uploads/tx_szleitlinien/021-007OLk_S3_Kolorektales-Karzinom-KRK_2019-01.pdf
- https://www.medizinische-genetik.de/index.php?id=crc-genetische-diagnostikc
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