

Colorectal cancers

Objectives:

- Definitions.
- Polyps.
- Basics of colorectal cancer.
- Surgery.
- Staging.

Resources:

- Davidson's.
- 436 doctors slides.
- Surgical recall.
- 435' team work

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COLOR INDEX:

NOTES, **IMPORTANT**, EXTRA, **DAVIDSON'S**<u>EDITING FILE</u>

FEEDBACK







Basic review:

The rectum has the intraperitoneal and extraperotineal part, Same as colon but has an only difference is that the rectum has no serosa.

- Colon wall from an inner layer to outer layer:
 - 1. Mucosa
 - 2. Submucosa
 - 3. muscularis propria
 - 4. Serosa
- Rectum wall:
 - 1. Mucosa
 - 2. Submucosa
 - 3. Muscularis propria
 - 4. mesorectal fat (no serosa so that's why in case of rectal cancer we give radiation because of the high recurrence which is within bony pelvis)

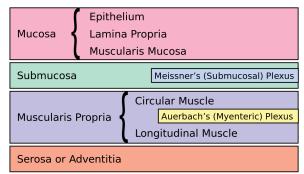
So, T3 of the colon = T4 of the rectum

*You need to differentiate between Muscularis Propria and Muscularis Mucosa.

"Any growth above the **basement membrane** is benign, whenever it reaches the basement membrane it's considered as malignant tumor and it can metastasize (through lymphatic+blood vessels)"

mucosa • submucosa • muscularis propria • serosa •

*serosa is not found on most of the rectum



Definitions:

- Colon = large bowel = large intestine. 150 cm in length
- **Rectum:** the terminal portion of the colon. 15 cm, most of the rectum is extra-peritoneal, has **no serosa** (that's how you can differentiate between rectum and colon)
- **Polyp:** benign growth (not invasive). (don't invade the basement membrane), benign polyps include: pseudopolyps, hyperplastic, Inflammatory, Adenomatous. All of them benign and will never develop cancer except **Adenomatous** with only 2% chance.

Polyp (benign) > invade basement membrane > malignant polyp (Carcinoma in situ) > invade submucosa > T1 cancer.

- Adenoma: type of polyp.
- Cancer: malignant growth (invasive¹).
- **Stage:** where the cancer is growing. (the fist step we do when we know the pt has cancer. في محلّه ولا راح مكان ثاني)
- **Primary:** the original tumour, where it started.
- Metastases: where the tumour has spread to.

Cancer:

A cancer cell is **immortal**, multiplies uncontrollably, can live on its own without neighbors, and can live in other parts of the body.

A brief intro to the lecture:

- % of all large bowel cancers occur in the rectum and sigmoid colon. the most clinical feature are change in bowel habit and blood per rectum.
- Diagnosis is done by colonoscopy and biopsy, CT colonography or barium enema "apple core sign"
- preoperative staging involves CT, MRI and PET CT scanning.

¹ invasion to basement membrane which located between Muscularis Mucosa and lamina propria contains lymphatics and Blood vessel



Colorectal Cancer

- Most cancers are acquired, some are inherited.
- Almost <u>all cancers begin as a benign polyp or adenoma</u>.
- Only a tiny percentage of adenomas become cancers.

بمعنى: كل الكانسرز تكون ادينوما في البداية، لكن مو شرط كل ادينوما تتحول لكانسر (فقط نسبة صغيرة منها).

Types of cancer: 1-sporadic 2-inherited 3-Familial

- If I see 100 pt., **95 of them will be Sporadic** (no genetics background). They will be aged around 50 60 y.o, while the rest will be: some will have IBD, Hereditary Nonpolyposis Colorectal Cancer (HNPCC) or Familial adenomatous polyposis (FAP)².
- What I want you to know about HNPCC that it's inherited colorectal cancer secondary to DNA mismatch gene repair. ابس مابيكم تضيعون

Polyp - Cancer Sequence:

- The process from benign polyp to cancer takes from **7 10 years**.
- The transformation into cancer is based on:
 - The type of polyp(Adenomatous, Villous(Worse), Sessile)
 - Size of polyp (>2cm)
 - Multiple polyps = greater risk of cancer.



- Villous type of adenomatous Polyp carries greater risk than tubular and tubulovillous.
- -Sessile polyp (directly adhere to the wall of bowel)> Pedunculated polyp(has a stick)
- Size greater than > 2cm.

Future therapy: <u>block one of those steps by gene therapy</u>. (won't be asked about genes)

Removing polyps prevents cancer: **Colonoscopy**. Screen with colonoscopy at 50 y of age if normal repeat after 10 y because cancer will develop after 10 y.





Fig. 22-8 Sessile polyp.



Pedunculated on the Right

Colorectal Carcinoma:

Classification:

- Adenocarcinoma³ (the most common)
- Carcinoid⁴ (rare)
- Lymphoma (rare)
- Sarcoma
- Squamous cell carcinoma⁵

Sessile on Left

Epidemiology:

- 3th most common malignancy worldwide.
- 1st most common in Saudi males.
- 2nd to lung cancer as a cause of cancer death
- 21,500 new cases, 8900 will die (2008)

² FAP: Defect in APC gene > grow thousands of polyps > increase risk of developing cancer at the age of 30 y.o.

³ 95% of cases.

⁴ common in rectal can be found by DRE

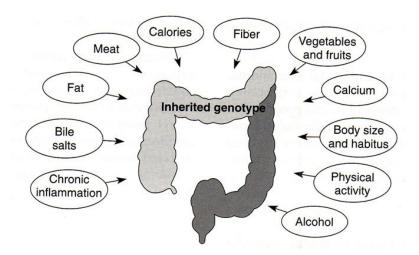
⁵ more common in long standing ulcerative colitis.



- risk of CRC women 1/16, men 1/14.
- peek incidence in 7th decade but it can occur at any age.

Etiology of Colorectal Cancer:

Causes are pure molecular genetics but those are predisposing factors.



Risk Factors:

- 1. Family history, Genetics: (main risk)
 - o Previous personal history increases the risk of recurrence, Increasing in age & being a Male.
 - o First degree family member history doubles the risk.
 - o Hereditary colorectal cancer syndromes.

Let's say a father diagnosed with CRC at 55 y.o, his son should be screened at 45 y.o for polyps since it proceeds CRC by 10 years, and keep screening every 5 years

- 2. Polyps
- 3. Inflammatory bowel disease
- 4. Others:
 - Diet and Nutrients "risk factor was high fat, red meat and low dietary ca and vit D deficiency" on the other side high fiber diet is a protective factor.
 - Smoking, ETOH alcohol
 - Lack of physical exercises.

Protective agents:

- Aspirin, NSAID
- Dietary Ca
- vit D supplement
- high fiber diet

This chart shows the effect of age on the incidence of colorectal cancer and colorectal Polyps

The incidence of colorectal cancer start peaking at 50 y.o, polyps proceed the incidence by 10 years. So when I get a young pt. (25 y.o) complaining of bleeding per rectal it's most likely hemorrhoids, or other

causes. We may get one case of colorectal of thousands in young

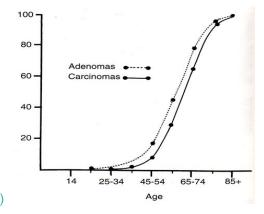
But when I get 50 y.o patient with per rectal bleeding you immediately think of colorectal cancer even if you see hemorrhoids, do colonoscopy!

Why Screening is important?

- Common
- Treatable
- Preventable
- Cost effective

For whom?

- Average risk group (every 10 years starting from age 50 y.o)
- High risk group (IBD, Polyps, 1st degree relative history of CRC)





Colorectal Cancer Risk Based on Family History:

General population	6%		
One 1st degree CRC	2-3X* (12-18%)		
Two 1st degree CRC	3-4X*		
One 1st degree CRC < 50 y	3-4 X*		
One 2nd or 3rd CRC	1.5X		
2 2nd degree CRC	2-3X*		
1 first degree with polyp	2X*		

Clinical presentation:

Sign & Symptoms	Presentation				
Bleeding	gross, occult, anemia (37%)(mainly in right colon cancer, because the widest portion of colon)				
Change in bowel habit pain, diarrhea, constipation, alternating pattern Most common presentation: abdominal pain, bleeding per rectal, change in bow habits (Diarrhea + Constipation), why? Narrowed lumen because of the tumor v cause constipation and the feces will build up pressure then diarrhea will result					
Obstruction more common with left sided lesions most common cause of bowel obstruction in the elderly "mostly Sigmoid cancer" (sigmoid is the narrowest portion of colon)					
Vague abdominal pain	"colicky"				
	Change in caliber of the stools				
	Weight loss				
Abdominal mass	especially in right sided colon cancer				
Tenesmus	over 50% occur with low rectal cancer				
Asymptomatic					

Investigations:

Colon:

Hx > PEx (mass, cachexia, LN adenopathy) > CBC + CEA > Radiology CT CAP if contrast contraindicated do CXR, U/S > only do MRI if you find something in the liver.

Rectum:

Hx > PEx (mass, cachexia, LN adenopathy) > CBC + CEA > Radiology CT CAP if contrast contraindicated do CXR, U/S > must do MRI pelvis + endoscopy, why? To see invasion to give neoadjuvant before surgery to reduce local recurrence.

ليش ما اسوي MRI في الـ colon؟ لأتي شايله شايله ما راح أعطى neoadjuvant اساسا!



1. General

- Complete history and physical examination (DRE)
 - -Look for signs of advanced disease such as cachexia, ascites or jaundice.
 - -Digital rectal exam > used to detect low cancer.

2. Endoscopic

- To identify primary & synchronous lesions by:
 - o Flexible sigmoidoscopy
 - Oclonoscopy used to: 1. Diagnose, 2. Take biopsy, 3. Exclude synchronous lesions.

3. Staging (more details below)

- Endorectal ultrasound (rectal cancer).
- Chest x-ray (metastases).
- Liver ultrasound (metastases).
- Abdominal CT scan (metastases).
- CT CAP (chest, abdomen, pelvis) scan is the investigation of choice.

CT colonography used to:

- 1. Assess the local extent of the disease,
- 2. Roll out Metastasis.

(Colon or rectal cancer should undergo CT of the chest and abdomen and pelvis.)

4. Bloodwork

- CBC electrolytes.
- CEA Carcinoembryonic antigen (tumour marker). Don't use CEA as screening as well as diagnostic. Use it
 only for monitoring the pt . CEA give us two things:
- 1. Idea about the prognosis (response to treatment). 2. Follow up every 3 months (check for recurrence)





Apple core lesion *AKA shouldering sign By constriction of the lumen of the colon by a stenosing colorectal carcinoma

Colorectal Cancer Pathology

- Macroscopic: 2/3 ulcerating, other types are: polypoidal and stenosing
- Microscopic (differentiation): For grading
 - 1-Well differentiated.
 - 2-Moderately differentiated.
 - 3-Poorly differentiated tumors have mucinous histology has a poor prognosis



• Lymph node involvement: For staging, via the portal blood to the liver or by transperitoneal seeding, low rectal tumours may involve the inquinal node

Treatment:

1st: Surgical therapy

- Surgery is the most important variable in the treatment of colorectal cancer.
- Radiation and chemotherapy alone cannot cure any stage of colorectal cancer.
- The site of tumour dictates the basic procedure.

Principles of Surgery

- Examine the entire abdomen. To make sure there is no metastasis.
- Remove the appropriate segment of the colon with adequate margins (5cm to the left of the tumor & another 5cm to the right in the draining Blood supply+adequate number of LN)
- Remove the corresponding lymph nodes

(Before surgery you can't determine LN infiltration. Thus you must resect 12 lymph nodes to make sure there is no infiltration and metastasis)⁶

- Open vs laparoscopic approach⁷.
 - In resection we depend on blood supply
 - Tumor in Right colon > Right hemicolectomy (all the ascending colon and part of transverse).
 - Tumor in Left colon > Left hemicolectomy (all the Descending colon and part of transverse).
 - Tumor in Transverse Colon > Extended Right or Left hemicolectomy.
 - Tumor in sigmoid > Anterior resection
 - Tumor in the Rectum > lower anterior resection (Sigmoidectomy).
 - Tumor in Lower Rectum > Abdominoperineal resection (APR).

A B C D A B F

Figure 19–1. Resection of colon cancer. Right colectomy (A), right hemicolectomy with division of middle colic pedicle (B), transverse colectomy (C), resection of splenic flexure sparing left colic artery (D), left hemicolectomy (E), sigmoid colectomy sparing left colic artery (F). (Reproduced, with permission, from Niederhuber JE, ed. Fundamentals of Surgery. Stamford, CT: Appleton & Lange; 1998:322 as modified from Schwartz SI, Ellis H. Maingot's Abdominal Operations. 10th ed. Norwalk, CT: Appleton & Lange; 1988:1053.)

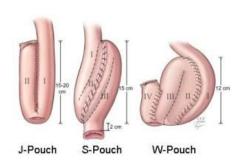
Surgery for rectal cancer:

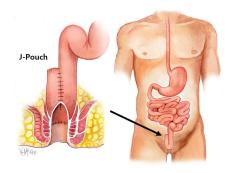
is excision of the entire mesorectum by using transanal endoscopic microsurgery (TEM) colonic J pouch 'ileoanal' may formed to improve defecation functions. - surgical procedure called a total proctocolectomy and ileal pouch anal anastomosis (IPAA), the entire colon and rectum are removed. A reservoir is created from the distal small bowel (called the ileum), which is then joined to the anal canal.

⁶ Studies showed that if you get 12 LN and all of them were negative the chance of positive LN are less than 1%. If Some are positive or if you didn't get 12 LN you should give chemotherapy.

⁷ Laparoscopic resections has shown some short-term benefits, less pain & shorter hospital stay (rapid recovery). However, there's no evidence of improved long-term outcomes over open surgery.







- Preparation for surgery :
 - 1. Patient: should be fasted prior to surgery.
 - 2. Comorbidity: should be addressed.
 - 3. Pre-operative: Give broad spectrum antibiotic.
 - 4. Thromboprophylaxis: "LMWH" to reduce risk of DVT and PE.
- Emergency colorectal resection:

In case of **perforation** or **obstruction** of colorectal cancer.









2nd: Follow Up

- Office visit every 3 months for two years then every 6 months for 3 years (total 5 years).
- Regular blood work (CEA). (when CEA starts to increase we should do colonoscopy & CT to check for recurrence)
- Colonoscopy: to check for recurrence: after 1 year of surgery, then after 3 years (4th year), then every 5 years.
- CT scan every year.

Who Gets Additional Treatment?

- 1. COLON:
 - All stage 3 patients (positive nodes) chemotherapy.
 - High risk stage 2 patients⁸
- 2. RECTUM:
 - All stage 2 and stage 3 patients should get radiation Before surgery and chemo (to decrease risk of recurrent).

⁸ who are high risk stage 2?

- poorly differentiated
- perineural/ perivascular invasion,
- inadequate retrieval of LN during resection
- obstruction/ perforation



-Difference between staging and grading:

Grading -> under <u>microscopy</u> is it poor or well differentiated.

staging-> position of the tumor is it metastasize to the liver or lymph node ..etc.

TNM staging system: Important

T stage: How far into the wall has it grown?						
Tis ⁹	invasion of mucosa only					
T1	Invasion of submucosa					
T2	Invasion of muscularis propria					
Full thickness (Serosa) in case of colon cancer. Perirectal fat and adjacent organs (as T4) in rectal cancer						
T4 Invasion into adjacent organs (Bladder)						
N stage: How many lymph nodes have been involved?						
N0	No lymph nodes involvement					
N1	1-3 lymph nodes					
N2	>3 lymph nodes					
N3	distant lymph nodes					
M stage: Are there distant organ metastasis?						
MO	No distant organ mets					
M1	Distant organ (liver, lung) (Peritoneum)					

^{*}Very important (don't mix Tumor stage with TNM)

very important (don't mix rumor stage with rivin)						
Based on the	TNM classification, we have 4 stages of Colorectal Cancer:	5 Year Survival chance				
Stage 0	Tis Tumors					
Stage 1	T1 and T2 tumors (No nodes nor mets)	90%				
Stage 2	T3 and T4 tumors (No nodes nor mets) / ^T3N0 tumors	80%^				
Stage 3	Any lymph node involvement (+ve node/s with any T) * (depends on number of nodes involved)	27-69%*				
Stage 4	Distant metastases (+ve mets with any T)	8% د التريكي قال إنها 50%				

In colon cancer we should use:

- Chemotherapy in Stage 3 & 4.
- Chemotherapy in Stage 2 if: poorly differentiated, perineural invasion, T2 Muscularis Propria perforated sigmoid cancer.

⁹ Carcinoma in situ

Keep in mind:

Tis - Invades BM

T1 - Submucosa

T3 - Serosa

T4 - Adjacent organs

* T4 in colon = T3 in rectum Rectum has no Serosa.



Polyps: "Extra from Davidson"

Colorectal adenoma						
Туре	solitary neoplastic					
What is it?	classified as tubular, tubulovillous or villous adenomas					
Chance of Malignancy	Villous adenomas has 30% chance of malignancy. tubular adenoma is around 10%. Multiple adenomas are 24%.					
Diagnosis	The majority are asymptomatic, but symptoms include rectal bleeding or large bowel colic, polyp may prolapse through the anus, Patients with giant villous adenoma of the rectum may present with severe watery diarrhea which lead to hypokalemia					
Management	Colonoscopic polypectomy. Follow up: colonoscopy is recommended after 6–12 months and 2–3 year.					
	Familial adenomatous polyposis					
Туре	multiple neoplastic					
What is it?	inherited as an autosomal dominant trait.It's single gene disorders, the gene responsible is APC, which is located on the long arm of ch.5					
Chance of Malignancy	Adenomatous polyps usually develop during teenage years and early adulthood, with > 90% chance of colorectal cancer					
Diagnosis	Clinicopathological diagnosis: requires the presence of > 100 adenomatous polyps of the large bowel established by sigmoidoscopy biopsy. Gene analysis					
Management	Prophylactic surgical resection of the large bowel is indicated for familial adenomatous polyposis					
	Juvenile polyposis syndrome					
Туре	Multiple non-neoplastic					
What is it?	Autosomal dominant genetic disorder					
Diagnosis	Causal mutations of the SMAD4 or BMPR1A/ALK3 genes					
Metaplastic associated polyposis						
Туре	Multiple non-neoplastic					
What is it?	autosomal recessive					
Chance of Malignancy	very high risk					
Diagnosis	Histologically, has a sawtooth pattern					
Management	Prophylactic colectomy and ileorectal anastomosis					



Recall:

What is the lifetime risk of colorectal cancer?

6%

What is the incidence of rectal cancer?

Comprises 20% to 30% of all colorectal cancer

What Other risk factors for colorectal cancer?

Family history is important when taking history: FAP, Lynch's syndrome⇒ HNPCC Hereditary NonPolyposis Colon Cancer—autosomal dominant inheritance of high risk or development of colon cancer.

What signs/symptoms are associated with the following conditions:

Right sided lesions	Left sided lesions
Right side of bowel has a large luminal diameter, so a tumor may attain a large size before causing problems. Microcytic anemia, occult/melena more than hematochezia(left sided) PR, postprandial discomfort, fatigue	Left side of bowel has smaller lumen and semisolid contents. Change in bowel habits (small caliber stools), colicky pain, signs of obstruction, abdominal mass, heme(+) or gross red blood. Nausea, vomiting, constipation.

What unique diagnostic test is helpful in patients with rectal Cancer?

Endorectal ultrasound (probe is placed transanally and depth of invasion and nodes are evaluated).

What are the signs/ symptoms of rectal cancer?

Most common symptom is hematochezia (passage of red blood stool) or mucus; also tenesmus, feeling of incomplete evacuation of stool (because of the mass), and rectal mass.

What disease does microcytic anemia signify until proven otherwise in a man or postmenopausal woman? Colon cancer.

What are the current recommendations for colorectal cancer screening if there is a history of colorectal cancer in a first degree relative less than 60 years old?

Colonoscopy at age 40, or 10 years before the age at diagnosis of the youngest first degree relative, and every 5 years thereafter.

What are the white lines of Toldt?

Lateral peritoneal reflections of the ascending and descending colon.

What is the blood supply to the rectum:

-	What is the blood supply to the rectain.	-
	Arteries	Veins
	Proximal: Superior hemorrhoidal (or superior rectal) from the IMA. Middle: Middle hemorrhoidal (or middle rectal) from the hypogastric (internal iliac). Distal: Inferior hemorrhoidal (or inferior rectal) from the pudendal artery (a branch of the hypogastric artery).	Proximal: Via the IMV to the splenic vein, then to the portal vein. Middle: Via the iliac vein to the IVC. Distal: Via the iliac vein to the IVC.

What parts of the GI tract do not have a serosa?

Esophagus, middle and distal rectum.

How are they anatomically classified?

Sessile (flat), Pedunculated (on a stalk)

What are the histologic classifications of the following types:

Inflammatory (pseudopolyp): As in Crohn's disease or ulcerative colitis

Hamartomatous: Normal tissue in abnormal configuration **Hyperplastic:** Benign - normal cells - no malignant potential

Neoplastic: Proliferation of undifferentiated cells; premalignant or malignant cells.



Summary

Colorectal Cancer							
1.C	lassification	1.Adenocarcinoma	2.Carcinoid	3.Lymphoma 4.Sarco	oma 5.Squamous cell carcinoma		
2.1	Causes are pure molecular genetics but those are predisposing factors: 1.Meat 2.chronic inflammation 3.Bile salt 4.fat 5.calories 6.Fibers 7.vegetables & fruits 8.calcium 9.Alcohol 10.physical activity 11.body size and habitus						
3.1	1. Family history, Genetics 2. Polyps 3. Inflammatory bowel disease 4.smoking & alcohol 5. Diet and Nutrients						
	4.Clinical presentation 1. Bleeding 2.Change in bowel habit 3. Obstruction 4. Vague abdominal pain 5. Abdominal mass						
5.lr	1. General "history physical examination" 2. Endoscopic "To identify primary & synchronous lesions" 3. Staging 4. Blood work "CBC & CEA						
1st: Surgical therapy: A.Remove the appropriate segment of the colon with adequate margins B.Remove the corresponding lymph nodes C.In resection we depend on blood supply D.Open vs laparoscopic approach 2nd: Follow Up: 1.Office visit every 3 months for two years then every 6 months for 3 years (total 5 years) 2. Regular blood work (CEA) 3. Colonoscopy CT scan .							
			TNM	staging system:			
	(is)	(0)	(1)	(2)	(3)	(4)	
N	No lymph nodes involvement		1-3 lymph nodes	>3 lymph nodes	distant lymph nodes		
М	_	No distant organ mets	Distant organ (liver, lung)				
invasi on of mucos a only		Invasion of Invasion of muscularis submucosa propria Full thickness (Serosa) in case of colon cancer. Perirectal fat and adjacent organs (asT4) in rectal cancer					
Based on the TNM classification, we have 4 stages of Colorectal Cancer:							
Stage 0 Tis Tumors		Stage 1 T1 and T2 tumors (No nodes nor mets)	Stage 2 T3 and T4 tumors (No nodes nor mets) / ^T3N0 tumors	Stage 3 Any lymph node involvement (+ve node/s with any T) (depends on number of nodes involved)	Distant metastases (+ve mets with any T)		
	Apple core appearance is Adenomatous polyp						



Questions

- 1. Which of the following carries the highest risk of transforming to colon cancer?
 - A. Villous adenomatous polyp.
 - B. Tubular adenomatous polyp.
 - C. Tubulovillous adenomatous polyp.
 - D. Inflammatory polyp.
- 2. A 28-year-old male patient was diagnosed with ulcerative colitis 7 years ago. He was advised to do annual colonoscopy to detect any changes that might lead to colon cancer. Which of the following type of cancers may develop in this case?
 - A. Sarcomas.
 - B. Squamous cell carcinoma.
 - C. Carcinoid.
 - D. Juvenile polyposis syndrome.
- 3. A 45-year-old female came to screening clinic to check if she has colon cancer and she has a family history of colon cancer. Which of the following will increase her chance of developing colon cancer the most?
 - A. 1st degree relative with colon cancer at 67 years of age.
 - B. Two 2nd degree relatives with colon cancer.
 - C. 1st degree with adenomatous polyp.
 - D. 1st degree relative with colon cancer under the age of 50.
- 4. A 75-year-old male presented to the ER with acute abdominal pain resulted from perforated colon cancer. Which of the following is the investigation of choice to check for metastatic tumors?
 - A. Liver ultrasound.
 - B. Chest X-ray.
 - C. CT- CAP (Chest, Abdomen and pelvis).
 - D. CEA (Carcinoembryonic antigen).
- 5. A 67-year-old male had a right hemicolectomy one week ago. Which of the following investigation is the most important in following up this patient?
 - A. CEA (Carcinoembryonic antigen).
 - B. CBC with differential.
 - C. Abdominal X-ray.
 - D. Flexible sigmoidoscopy.
- 6. Regarding colon cancer staging, which of the following is correct?
 - A. N3 indicates more than 3 lymph nodes are positives.
 - B. T4 indicates metastasis to distant organs.
 - C. N0 indicates less than 3 lymph nodes are positives.
 - D. N3 indicates distant lymph node involvement.



- 7. A 58-year-old male was diagnosed to have colon cancer and went for colectomy. Unfortunately, the surgeon could not excise more than 9 lymph nodes. The Histopathologist reported that all lymph nodes are free of cancerous cells. Which of the following is your next step in managing this patient?
 - A. Assure the patient and ask him for annual checkup.
 - B. Ask the patient to comeback after 6 months to do CT.
 - C. Re-operate and try to excise more lymph nodes.
 - D. Give the patient prophylactic chemotherapy.
- 8. Regarding the histology of the rectum, which of the following is absent?
 - A. Mucosa.
 - B. Submucosa.
 - C. Muscularis propria.
 - D. Serosa.

Answers:										
1:A	2BD	3:D	4:C	5:A	6:D	7:D	8:D			