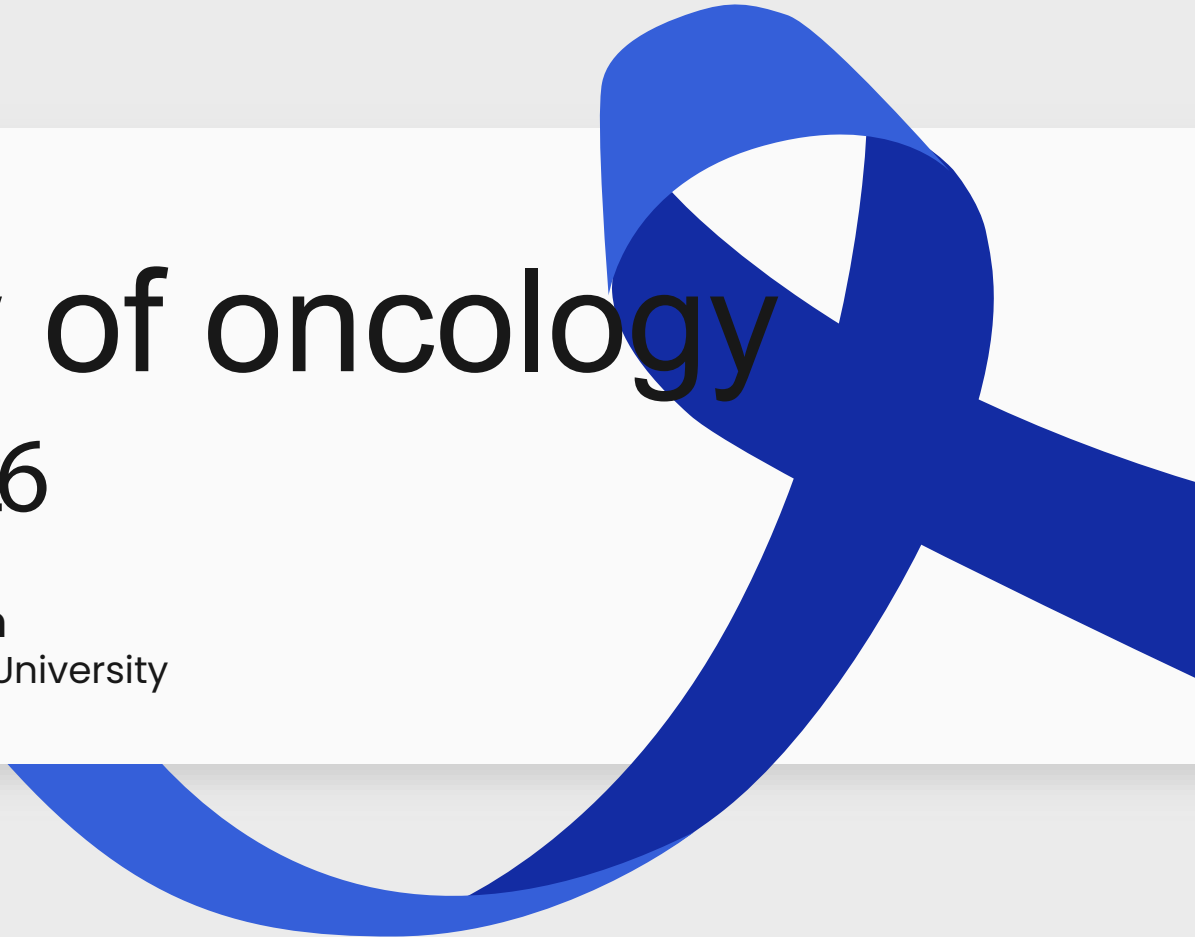




คณะแพทยศาสตร์ มหาวิทยาลัยขอนแก่น
FACULTY OF MEDICINE KHON KAEN UNIVERSITY

Overview of oncology for resident 2026

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Faculty of Medicine, Khon Kaen University



Disclosures

- This academic session is sponsored by Berlin pharmaceutical industry.
- This session is oversimplified framework to guide your learning and cannot replace your lecture and personal reading.
- The information is accurate at the time of presentation and independently from sponsor.

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Cancer screening

Hereditary cancer

Cancer screening

Cancer screening



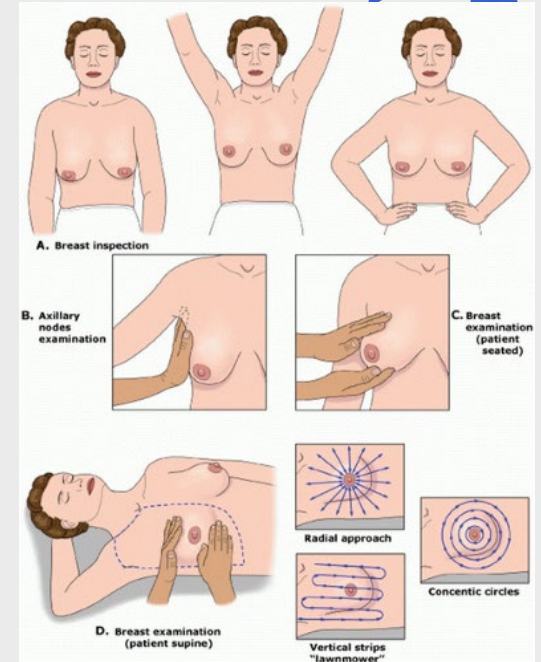
- **Part of secondary cancer prevention**
- **Cost-effectiveness and effective program**
 - : Breast cancer
 - : Colorectal cancer
 - : Cervical cancer
- **Lung cancer screening using low dose CT is still debatable.**
- **No role of tumor marker in cancer screening**

Breast Cancer screening



Average risk and asymptomatic

| Age | Tool |
|---------------|---|
| Age 25 - 39 | Clinical Breast Examination q 1-3 yr |
| Age \geq 40 | Clinical Breast Examination q 1-3 yr Mamogram q 1 yr |



NCCN has not established an upper age limit for screening as long as that individuals in overall good health (some guidelines limit to age of 75)

Breast Cancer screening

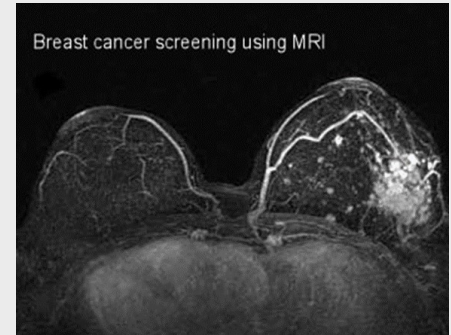


Germline BRCA Pathogenic/Likely pathogenic

| Tool | Starting Age |
|--|--------------|
| Cinical Breast Examination every 6-12 months | Age 25 |
| MRI Breast every 1 year | Age 25 |
| Mammogram every 1 year | Age 30 - 75 |



Breast cancer screening using MRI



colorectal Cancer scree

Starting at
age of 45-50



Average risk

- Age 45–75 years
- No personal history of adenoma or sessile serrated polyp/sessile serrated lesion or CRC
- No personal history of inflammatory bowel disease (IBD)
- No personal history of high-risk CRC genetic syndromes
- No personal history of cystic fibrosis
- No personal history of childhood cancer
- Negative family history for confirmed advanced adenoma
- Negative family history for CRC

| Tools | Frequency (normal finding) |
|--|-------------------------------|
| Colonoscopy | 10 years |
| Flexible sigmoidoscopy | 5 -10 years |
| CT colonography | 5 years |
| Stool occult blood test | 1 year |
| Multitargeted stool DNA-based testing | 3 years |

colorectal Cancer screening



Increased risk population

| FAMILY HISTORY CRITERIA | SCREENING ^{fff} | |
|--|---|---|
| ≥1 first-degree relative with CRC at any age | Colonoscopy beginning at age 40 y or 10 y before earliest diagnosis of CRC, whichever is first | Repeat every 5 y ^{ddd,fff,ggg,hhh} or if positive, repeat per colonoscopy findings |
| Second- and third-degree relatives with CRC at any age | Colonoscopy beginning at age 45 y ^{ddd} | Repeat every 10 y or if positive, repeat per colonoscopy findings |
| First-degree relative with confirmed advanced adenoma(s) (ie, high-grade dysplasia, ≥1 cm, villous or tubulovillous histology, TSA), or advanced SSPs/SSLs (≥1 cm, any dysplasia) at any age ^{eee,iii,jj} | Colonoscopy beginning at age 40 y or at age of onset of adenoma in relative, whichever is first | Repeat every 5–10 y ^{fff,ggg} or if positive, repeat per colonoscopy findings |

Cervical Cancer screening



| | การตรวจหาเชื้อเอชพีวีกลุ่มความเสี่ยงสูง (High-risk HPV Testing) | | การตรวจเซลล์วิทยา (Conventional หรือ Liquid-Based Cytology) |
|--|--|---|--|
| | ตรวจหาเชื้อเอชพีวี กลุ่มความเสี่ยงสูง (แนะนำมากกว่า) | ตรวจหาเชื้อเอชพีวีกลุ่มความเสี่ยงสูง ร่วมกับการตรวจเซลล์วิทยา (Co-testing) | |
| อายุที่เริ่มตรวจ | 25 ปี | 25 ปี | 25 ปี |
| ความถี่ | ทุก 5 ปี | ทุก 5 ปี | ทุก 2 ปี |
| อายุที่หยุดตรวจ | > 65 ปี ถ้าผลการ ตรวจปกติ ติดต่อกัน 2 ครั้ง | > 65 ปี ถ้าผลการตรวจปกติ ติดต่อกัน 2 ครั้ง | > 65 ปี ถ้าผลการ ตรวจปกติ ติดต่อกัน 5 ครั้ง |
| <ul style="list-style-type: none"> • สตรีที่อายุน้อยกว่า 25 ปี ไม่แนะนำให้ตรวจคัดกรอง ยกเว้นในสตรีที่มีความเสี่ยงสูง เช่น ติดเชื้อเอชไอวี มีคู่นอนหลายคน เป็นหูดหงอนไก่หรือเป็นโรคติดเชื้อทางเพศสัมพันธ์ เป็นต้น • สตรีที่ตัดมดลูกพร้อมกับปากมดลูกออกแล้วและไม่มีประวัติ CIN 2-3 หรือ AIS หรือมะเร็งปากมดลูก | | | |

Lung Cancer screening **only for High risk population**



National
Comprehensive
Cancer
Network®

NCCN Guidelines Version 1.2026 Lung Cancer Screening

[NCCN Guidelines Index](#)
[Table of Contents](#)
[Discussion](#)

RISK ASSESSMENT^{a,b,c,d}

- Cigarette smoking history^e
- Radon exposure^f
- Occupational exposure^g
- Cancer history^h
- Family history of lung cancer in first-degree relatives
- Disease history (chronic obstructive pulmonary disease [COPD] or pulmonary fibrosis)
- Cigarette smoking exposureⁱ (second-hand smoke)
- Risk calculator to enhance determination of risk status^{j,k}

Individuals not eligible for lung cancer screening:

- Symptoms of lung cancer (see [NCCN Guidelines for Non-Small Cell Lung Cancer](#))
- Previous lung cancer (see Surveillance in the [NCCN Guidelines for Non-Small Cell Lung Cancer](#))
- Functional status and/or comorbidity that would prohibit curative intent treatment^l (see Principles of Surgery and Principles of Radiation Therapy in the [NCCN Guidelines for Non-Small Cell Lung Cancer](#))
- Likely near-future competing cause of death

RISK STATUS

Higher risk^{i,m,n}

- Age ≥50 y (category 1) and
- ≥20 pack-year history of smoking cigarettes (category 1) or ≥20-year history of smoking cigarettes¹ (category 2B)

In candidates for screening, a discussion of benefits/risks is recommended^{c,k}

Low-dose CT (LDCT)^{o,p}
(category 1)

Screening Findings ([LCS-2](#))

* Use low-dose CT

* CXR is not recommended for lung cancer screening

Lower risk

- Age <50 y and/or
- <20 pack-year history of smoking cigarettes or <20-year history of smoking cigarettes¹ (category 2B)

Lung cancer screening not recommended

Hereditary cancer

Hereditary cancer



Characteristics suspicious for hereditary cancer

- Cancers occurring at **younger ages** than usual
- **More than one type of cancer** in a single person
- Cancers occurring in **both of a pair of organs**
- Cancer occurring in the **sex not usually affected** (e.g. breast cancer in a man)
- **Two or more family members with similar cancer** or cancer syndrome related cancer
- Clinical finding associated with hereditary cancer

Cancer syndrome needed to know

- BRCA-related cancer
- Familial colorectal cancer: FAP, HNPCC

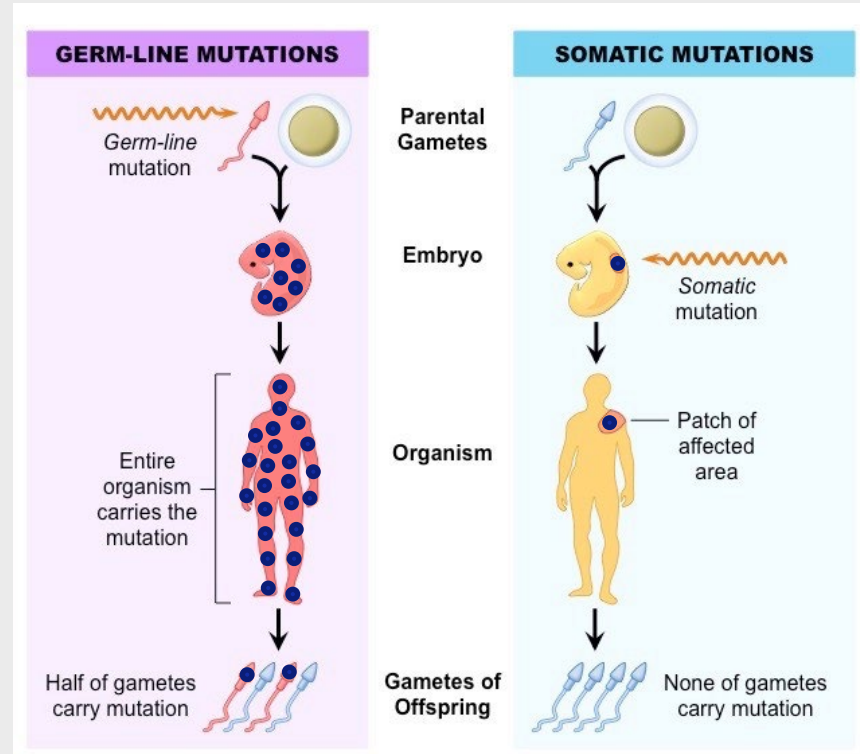
mutation



- **Somatic mutations**
 - : Occur in a single body cell and cannot be inherited
 - : Only tissues derived from mutated cell are affected

- **Germline mutations**
 - : Occur in gametes and can be passed onto offspring
 - : Every cell in the entire organism will be affected

Hereditary cancer

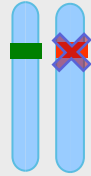


Knudson's Two Hit Hypothesis

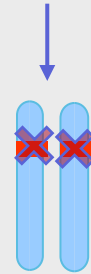


Inherited case

One normal allele
+
One mutant allele
(1st "hit")



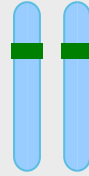
Two mutant alleles
(2nd "hit")



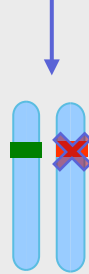
Tumorigenesis

Sporadic case

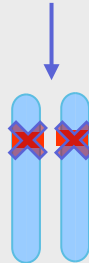
Two normal alleles



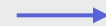
One normal allele
+
One mutant allele
(1st "hit")



Two mutant alleles
(2nd "hit")



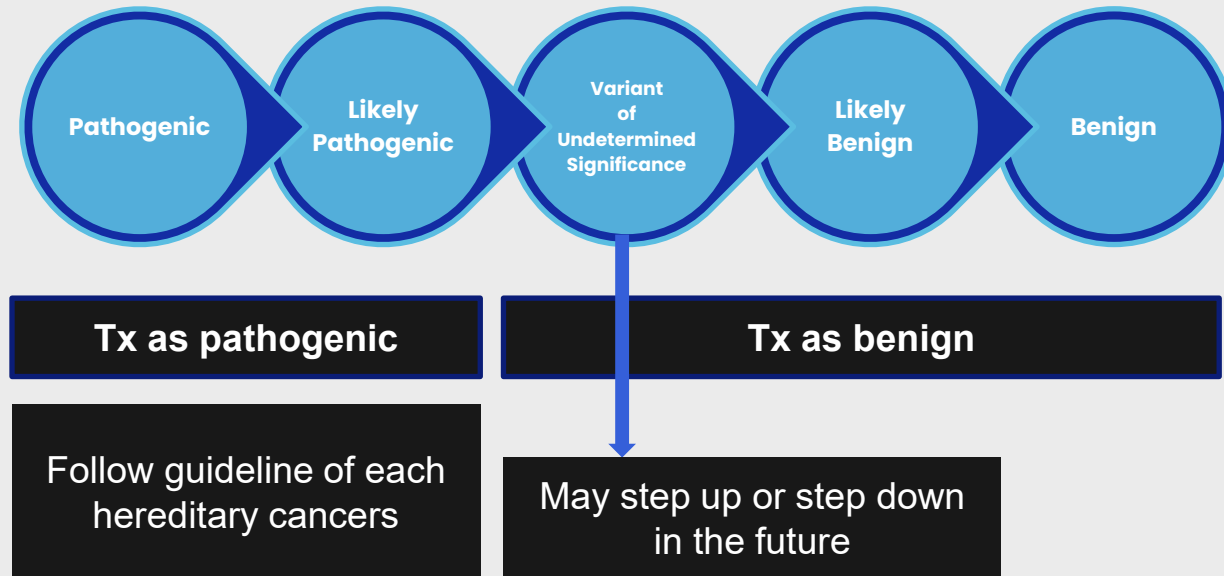
Tumorigenesis



Germline variants interpretation



Based on ACMG-AMP Guideline 2015

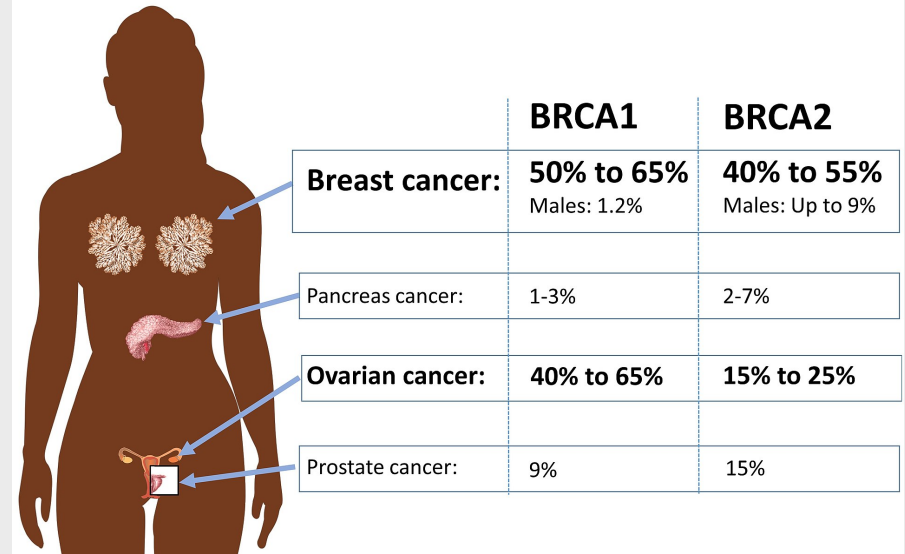
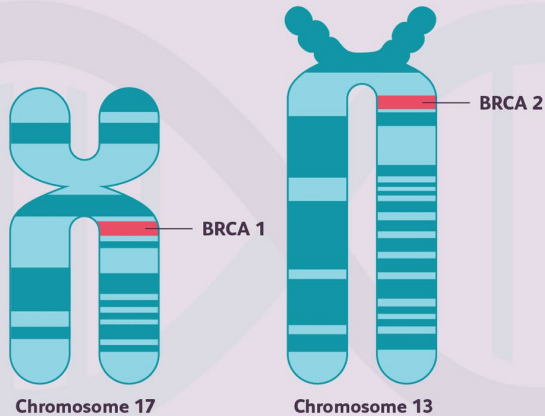


BRCA-RELATED CANCER



- BRCA1 and BRCA2** are tumor suppressor genes, involving in DNA repair pathway called homologous recombinant repair.

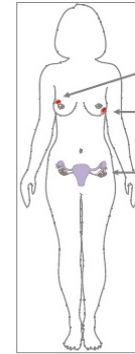
Where Are Your BRCA Genes?



BRCA-RELATED CANCERS

- Multiple cases of early onset breast cancer
- Ovarian cancer (with family history of breast or ovarian cancer)
- Breast and ovarian cancer in the same woman
- Bilateral breast cancer
- Ashkenazi Jewish heritage
- Male breast cancer

BRCA1-Associated Cancers: Lifetime Risk



Breast cancer 50%–85% (often early age at onset)

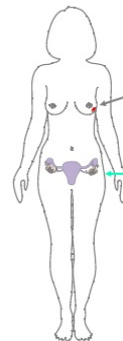
Second primary breast cancer 40%–60%

Ovarian cancer 15%–45%

Possible increased risk of other cancers (eg, prostate, colon)

ASCO

BRCA2-Associated Cancers: Lifetime Risk



breast cancer
(50%–85%)

ovarian cancer
(10%–20%)

male breast cancer
(6%)

Increased risk of prostate,
laryngeal, and pancreatic cancers
(magnitude unknown)



ASCO



Clinical use of CAR-RELATED CANCER



- **Early cancer screening**
- **Risk-reducing surgery**
 - : Prophylactic mastectomy – to reduce risk of breast cancer
 - : Prophylactic salpingo-oophorectomy – to reduce risk of ovarian cancer
- **Chemopreventive medication**
 - : Antiestrogens for breast cancer
- **Targeted therapy**
 - : Use of PARP inhibitor in BRCA-mutated cancer via synthetic lethality pathway
 - : Olaparib
 - : Rucaparib
 - : Niraparib
 - : Talazoparib

brCA-RELATED CANCER



Chemoprevention of Breast Cancer in *BRCA1/2* Carriers

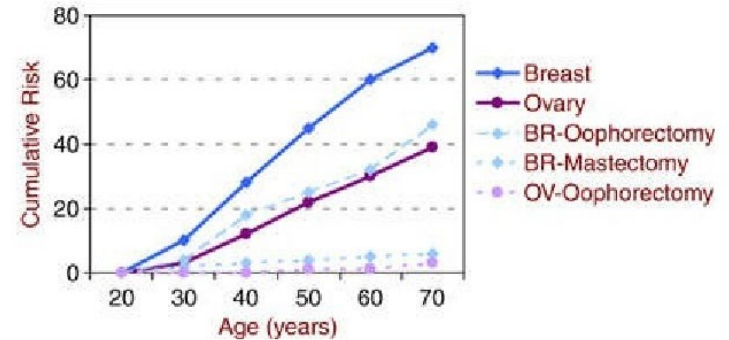
Tamoxifen



Risk reduction of 50% or more
in both *BRCA1* and *BRCA2* carriers

Gronwald J et al, Int J Cancer 2006;118(9):2281-4

Cancer risk reduction with prophylactic surgery



Domchek and Weber, Oncogene 2006; 25:5825-5831

TESTING CRITERIA FOR HIGH-PENETRANCE BREAST CANCER SUSCEPTIBILITY GENES (Genes such as **BRCA1**, **BRCA2**, **CDH1**, **PALB2**, **PTEN**, **STK11**, and **TP53**. See [GENE-A](#))^{a,g,h,i,j}

Testing is clinically indicated in the following scenarios:

- See General Testing Criteria on [CRIT-1](#).

- Personal history of breast cancer with specific features:

- ▶ ≤50 y

- ▶ Any age:

- ◊ Treatment indications

- To aid in systemic treatment decisions using PARP inhibitors for breast cancer in the metastatic setting^{k,l} (See [NCCN Guidelines for Breast Cancer](#))
 - To aid in adjuvant treatment decisions with olaparib for high-risk,^m HER2-negative breast cancer^l

- ◊ Pathology/histology

- Triple-negative breast cancer
 - Multiple primary breast cancers (synchronous or metachronous)ⁿ
 - Lobular breast cancer with personal or family history of diffuse gastric cancer (See [NCCN Guidelines for Genetic/Familial High-Risk Assessment: Colorectal, Endometrial, and Gastric](#))

- ◊ Male breast cancer

- ◊ Ancestry: Ashkenazi Jewish

- ▶ Any age (continued):

- ◊ Family history^o

- ≥1 close blood relative^p with ANY:

- breast cancer at age ≤50 y
 - male breast cancer
 - ovarian cancer
 - pancreatic cancer
 - prostate cancer with metastatic,^q or high- or very-high-risk group (Initial Risk Stratification and Staging Workup in [NCCN Guidelines for Prostate Cancer](#))

- ≥3 diagnoses of breast and/or prostate cancer (any grade) on the same side of the family including the patient with breast cancer

- Family history criteria: unaffected; or affected but does not meet above criteria

- ▶ Individual with a first- or second-degree blood relative meeting any of the criteria listed above (except unaffected individuals whose relatives meet criteria only for systemic therapy decision-making).^f

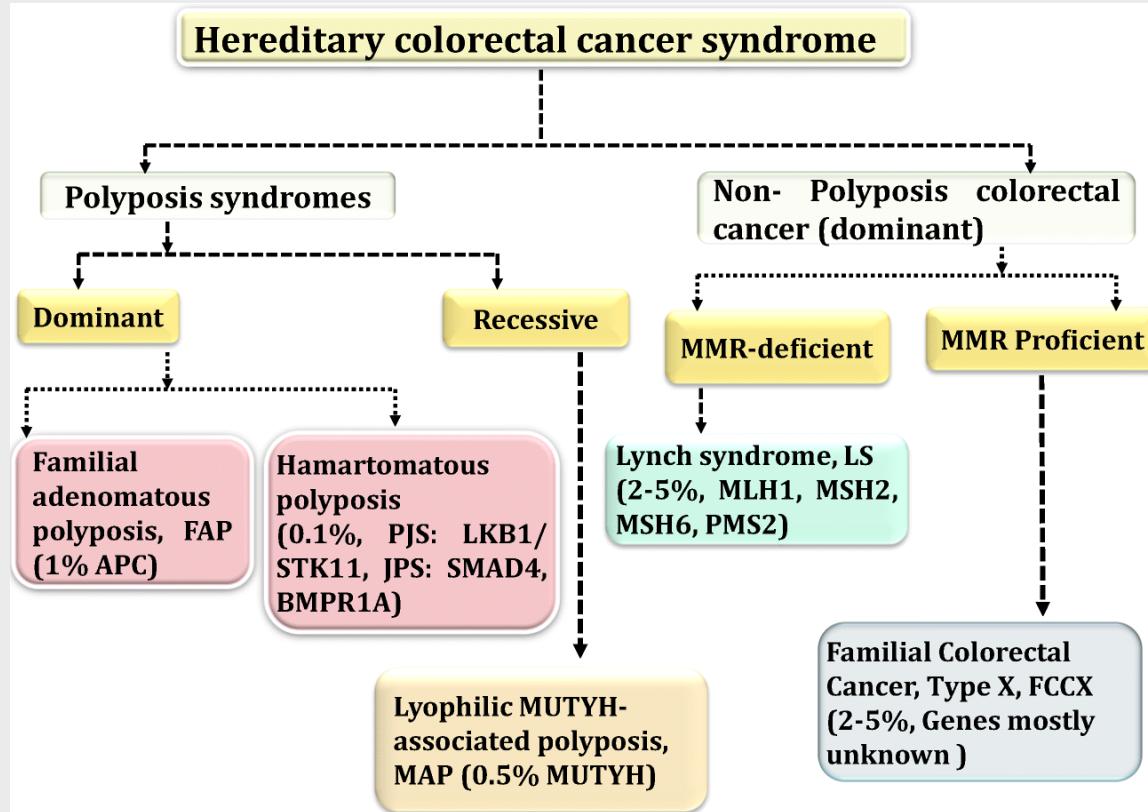
- ▶ Individuals who have a probability >5% of a **BRCA1/2** P/LP variant based on prior probability models (eg, Tyrer-Cuzick, BRCAPro, CanRisk).^s

Criteria met → [GENE-1](#)

If testing criteria not met, consider testing criteria for other hereditary syndromes

If criteria for other hereditary syndromes not met, then cancer screening as per [NCCN Screening Guidelines](#)

Colorectal CANCER syndrome



Colorectal CANCER syndrome



| Syndrome | Gene | Associated cancer |
|----------------------|--|--|
| Lynch (HNPCC) | MSH 2, MLH 1, MSH 6, PMS 2 (autosomal dominant) | Colorectal cancer (often right-sided), Endometrial cancer Stomach, Ovary, Bladder, Urinary tract, Kidney, Gall bladder, Brain, Breast, Small bowel |
| FAP | APC (autosomal dominant) | Colorectal cancer (>100 polyp) Medulloblastoma, Papillary CA thyroid, Hepatoblastoma Pancreatic cancer Gastric and Duodenal cancer |

clinical use of mmr gene



- Lynch syndrome
: MMR gene (MLH1, MSH2, MSH6, PMS2)
- Prognostic biomarker in stage II colon cancer
: can omit adjuvant chemotherapy
- Predictive biomarker for immunotherapy in advanced CRC



"Amsterdam" Criteria for HNPCC Diagnosis

- 3 relatives with colorectal cancer, where one is 1st degree relative of other two
- 2 generations of colorectal cancer
- 1 colorectal cancer before age 50
- FAP is excluded

02

Paraneoplastic syndrome

PARANEOPlastic syndrome



- Paraneoplastic syndromes are rare disorders with complex systemic clinical manifestations due to underlying malignancy.
- In paraneoplastic syndromes, the malignant cells do not directly cause symptoms related to metastasis; rather, they generate **autoantibodies, cytokines, hormones, or peptides** that affect multiple organ systems.
- **Symptoms can manifest before or after the diagnosis of cancer.**
- Prompt recognition of these syndromes is critical as it may reveal hidden malignancy, affecting clinical outcomes.

Common Paraneoplastic syndrome



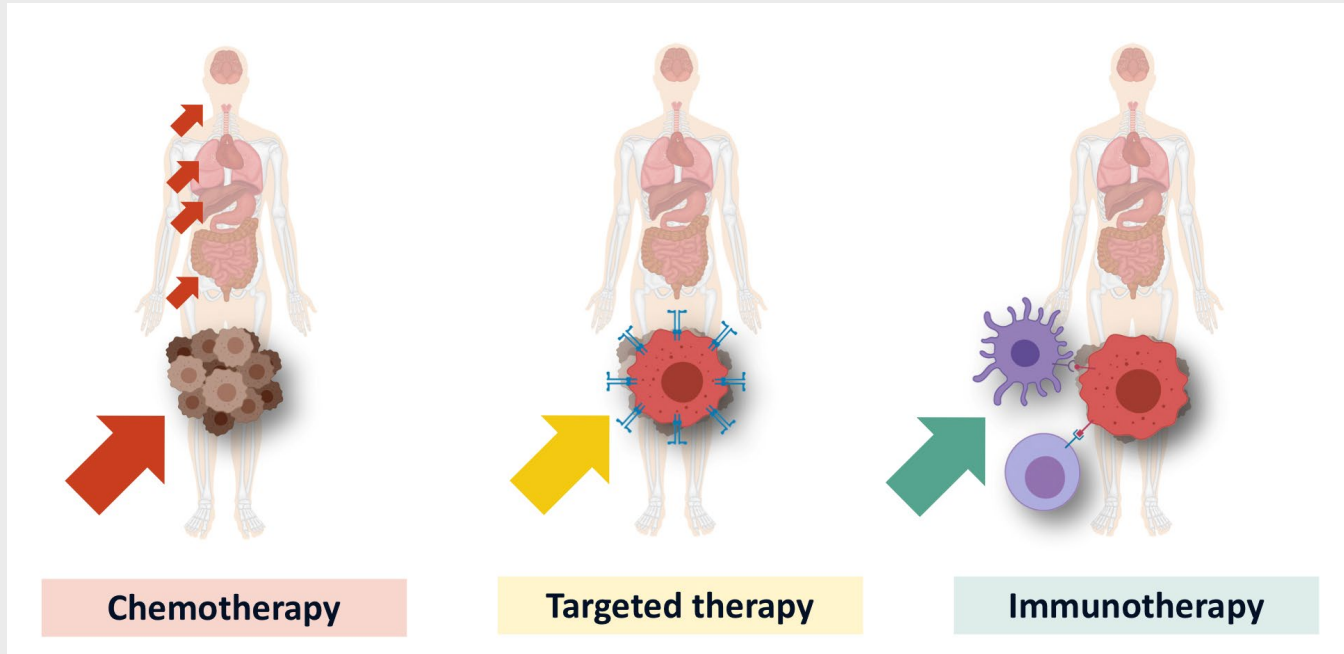
| Cancer | Common Paraneoplastic Syndrome |
|-------------------------|--|
| Small cell lung cancer | Ectopic ACTH SIADH Lambert-Eaton myasthenic syndrome Cerebellar degeneration Limbic encephalitis |
| Thymoma | Myasthenia gravis Pure red cell aplasia Hypogammaglobulinemia Paraneoplastic pemphigus |
| Adenocarcinoma of lung | Hypertrophic osteoarthropathy (HOA) |
| Squamous cell carcinoma | Hypercalcemia (PTHrP-related) |

03

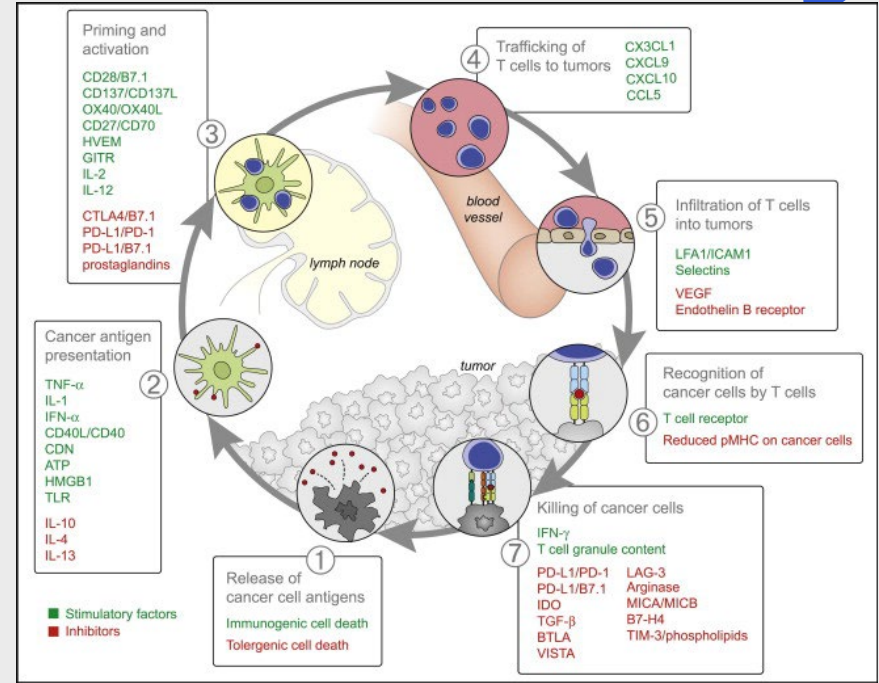
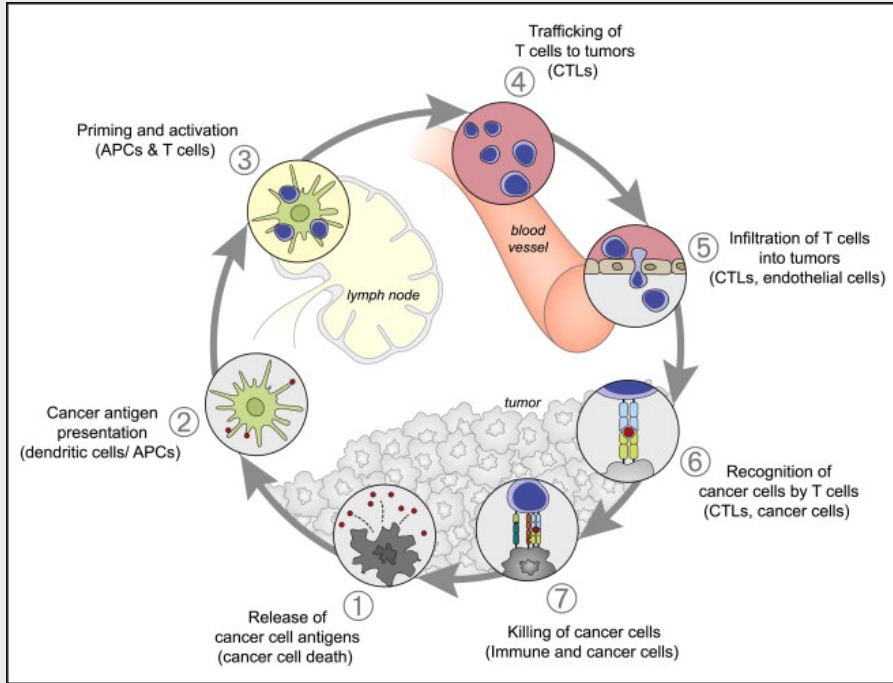
Common cancer management

Cancer treatment

CANCER treatment



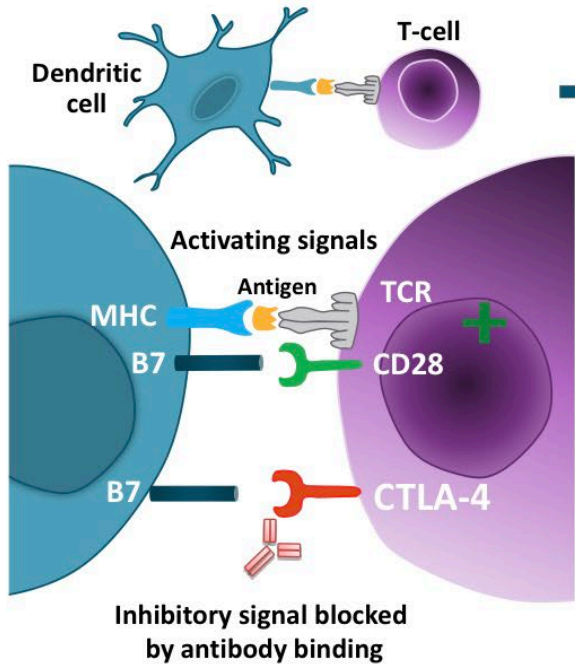
immunotherapy



immunotherapy

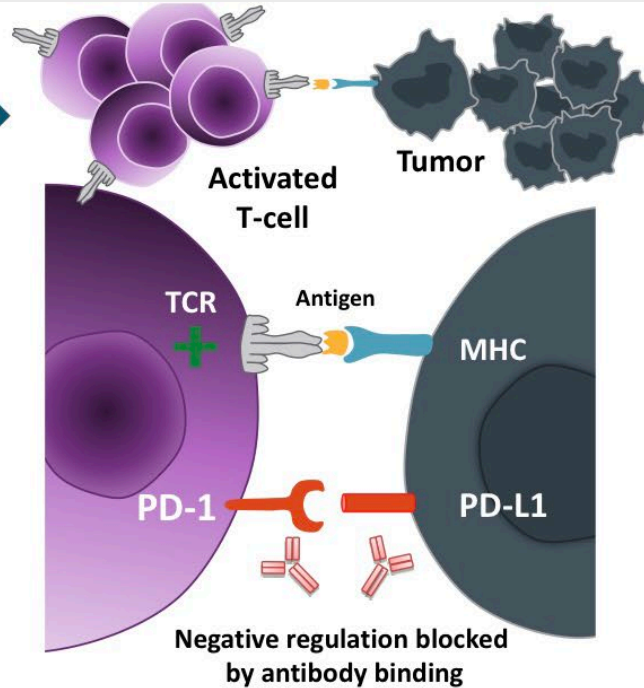


Priming Phase



Activation →

Effector Phase



Anti-CTLA-4
Ipilimumab
Tremelimumab

Anti-PD-1
Avelumab
Cemiplimab
Dostarlimab
Nivolumab
Pembrolizumab
Retifanlimab

Anti-PD-L1
Atezolizumab
Durvalumab

breast cancer

Breast cancer



Surgery

Breast conserving surgery
(Wide excision, Lumpectomy)

Mastectomy



CMT

Anthracycline-based
(Doxorubicin, Cyclophosphamide)

Taxane-based
(Docetaxel, Paclitaxel)

Others



RT

Post-op RT

Palliative RT

Breast cancer



Hormonal

SERM: Tamoxifen
(Pre-menopause)

Aromatase inhibitor
: Letrozole, Exemestane
Anastrozole

SERD: Fulvestrant



Anti-HER2

Trastuzumab
Pertuzumab
Lapatinib
TDM-1
Trastuzumab-deruxitan



Others

CDK4/6 inhibitors:
Ribociclib, Palbociclib,
Abemaciclib

EARLY Breast CANCER



Adjuvant treatment after surgery (in brief)

| | Hormone | CMF | Anti-her 2 | RT |
|---|---------|-----|------------|--|
| HR+HER2- Luminal A: ER+ PR+ Ki67 <20 Luminal B: ER+ PR +/- Ki67 > 20 | ✓ | ✓ | | BCS* LN ≥4 T > 5 cm Margin < 1 mm |
| HR+HER2+ | ✓ | ✓ | ✓ | |
| HR-HER2+ | | ✓ | ✓ | |
| HR-HER2- (triple negative) | | ✓ | | |

*BCS: Breast-conserving surgery

ADVANCED Breast CANCER



HR+ HER2-

- Visceral crisis : CMT
- No visceral crisis : Hormone + CDK4/6 inhibitor

HR- HER2+

- CMT + Anti-HER2 (Pertuzumab/Trastuzumab)

Triple negative

- CMT+/-Immunotherapy: Pembrolizumab
- PARP inhibitor in germline BRCA mutation

Visceral crisis

- Severe organ dysfunction/rapid disease progression
- Liver: TB > 1.5x ULN
(in the absence of biliary tract obstruction)
- Lung: dyspnea at rest
(after drainage of pleural effusion)

Note:

No definite role of surgery and RT in metastatic setting except for palliative symptom control

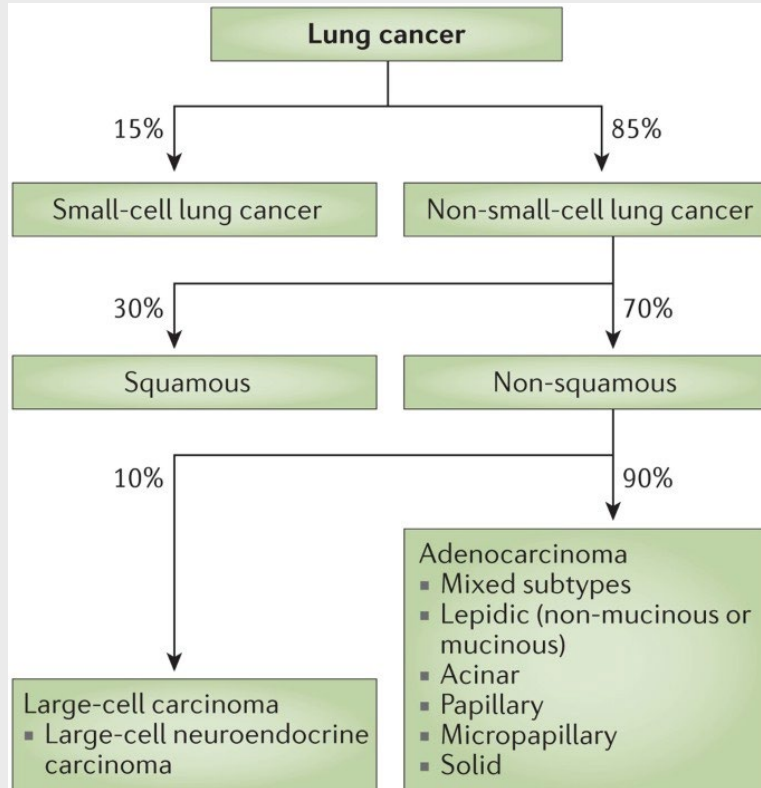
Treatment toxicity



| | Anthracycline | Anti-HER 2 |
|-------------------|--|---|
| Mechanism | Type I Oxidative stress and free radicles | Type II HER2 signaling blockage |
| Myocardial effect | Irreversible | Reversible |
| Dose-related | Dose-related | No cumulative dose-related |
| Risk | Hypertension | Prior/concurrent anthracycline, obesity |
| Prevention | Minimized dose of anthracycline Use of cardioprotective agent | F/U echocardiography Use of cardioprotective agent |

lung cancer

lung CANCER



Histologic subtypes associated with smoking

- 1. Squamous cell carcinoma**
- 2. Large cell carcinoma**
- 3. Small cell lung cancer**

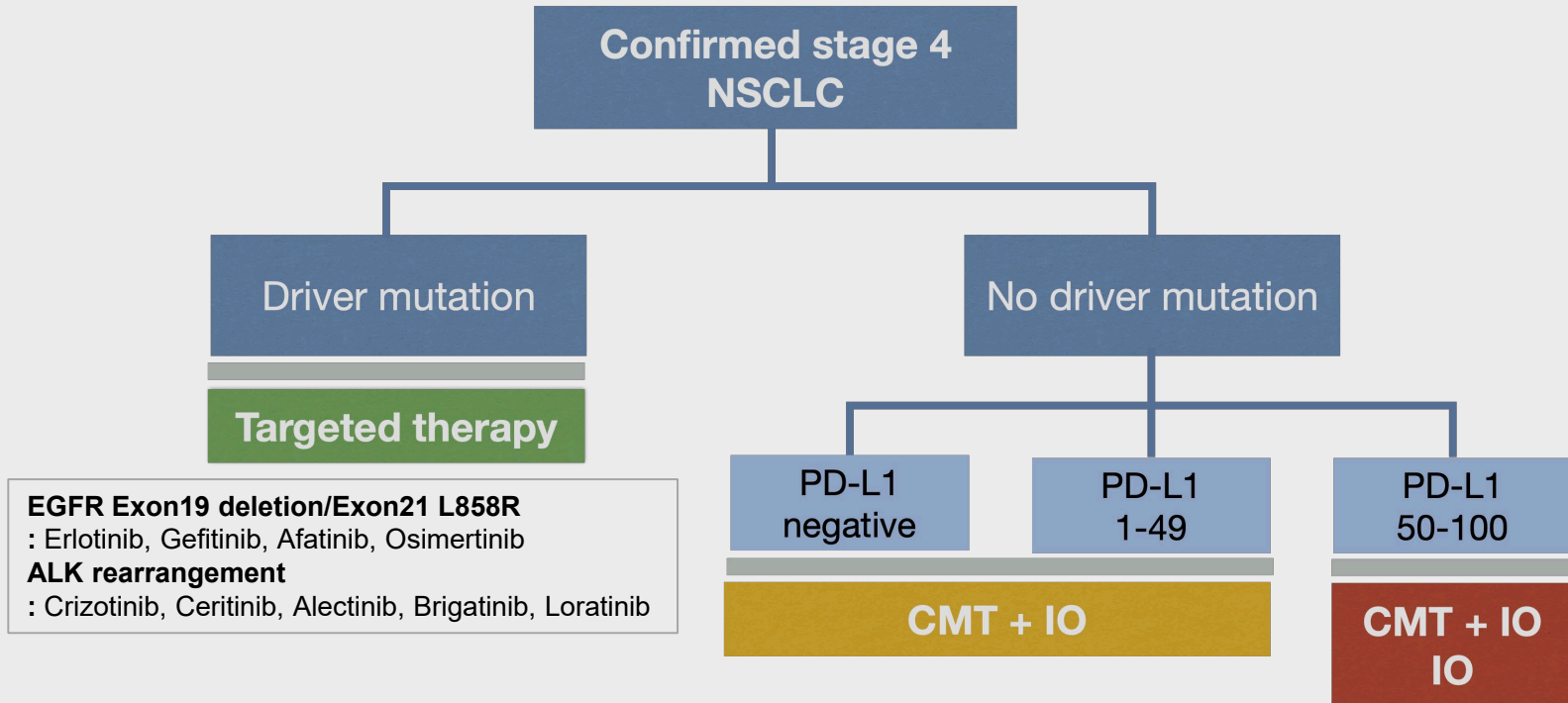
Non small cell lung CANCER (Concept)



| Stage | Treatment |
|-------|---|
| I | Surgery |
| II | Surgery + Adjuvant CMT (or pre/peri operative strategies) |
| III | Multidisciplinary team approach CCRT followed by Durvalumab (if EGFR/ALK neg) or Osimertinib (if EGFR mutant) |
| IV | Systemic treatment |

- Adjuvant chemotherapy recommended in node positive disease or tumor size ≥ 4 cm
- Adjuvant osimertinib recommended in resected stage IB – III with EGFR exon 19 del or exon 21 L858R mutation
- Adjuvant immunotherapy recommend in resected stage II-III without EGFR/ALK mutation and with PD-L1 positive
- Peri-operative or neoadjuvant chemotherapy and immunotherapy recommend in stage II-III without EGFR/ALK mutation

Non small cell lung CANCER stage IV



NSCLC with Brain metastasis



- Surgery may be offered for patients with brain metastases, considering the following factors:
 - Suspected brain metastases without a primary cancer diagnosis.
 - Large tumors with mass effect.
- Patients with symptomatic brain metastases should be offered local therapy (radiosurgery and/or radiation therapy and/or surgery)
- For patients with asymptomatic brain metastases, local therapy should not be deferred unless patients who received brain-penetrating drug
 - EGFR mutant: Osimertinib
 - ALK-rearrangement: Lorlatinib, Alectinib, Brigatinib

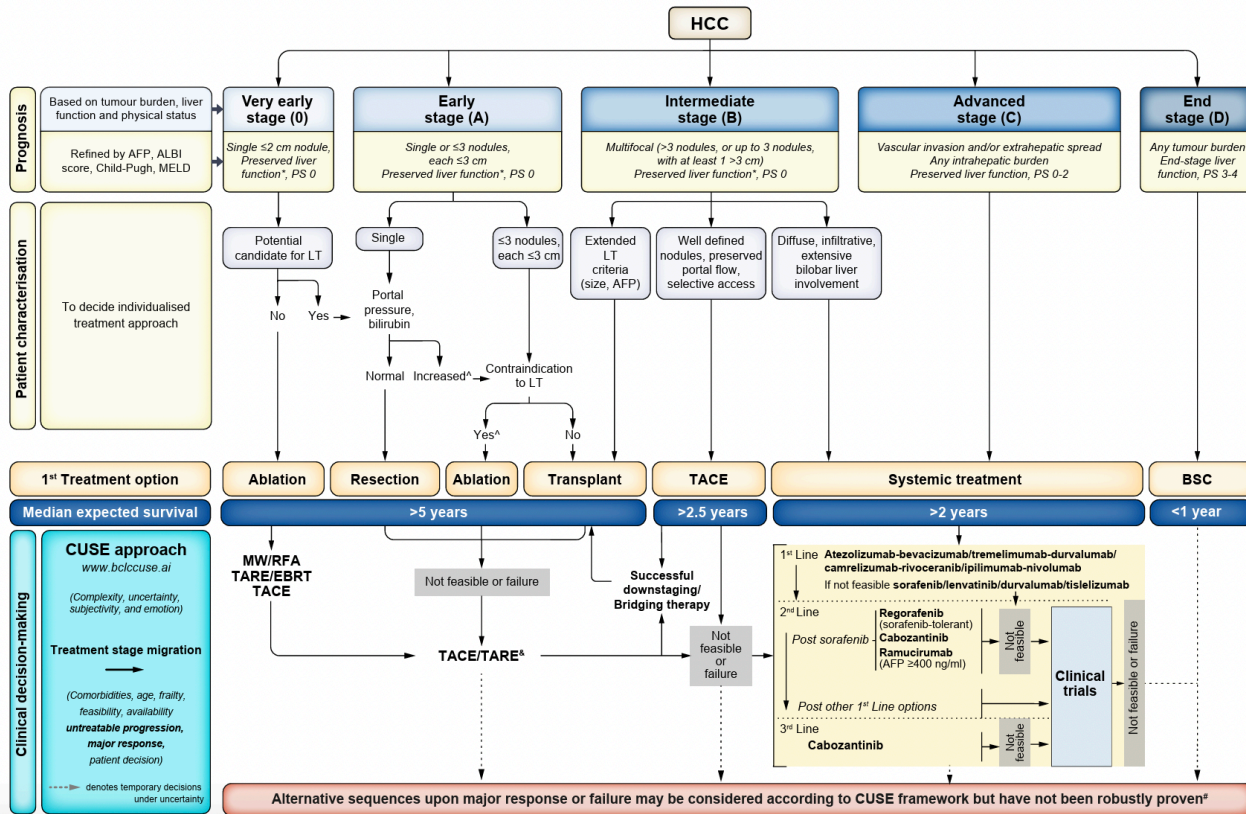
Small cell lung cancer



- Originated from neuroendocrine cell, accounting for 10–15% of lung cancer
- Rapid growth, tendency to metastasize
- Smoking-related disease, Highly systemic disease
- Paraneoplastic syndrome: SIADH, Ectopic ACTH, LEMG
- Staging: Limited vs. Extensive stage
- Poor prognosis
- **Treatment: Limited stage: CCRT with Cisplatin/Etoposide**
Extensive stage: Carboplatin/Etoposide + Immunotherapy

Hepatocellular carcinoma

Hepatocellular carcinoma



Barcelona Clinic Liver cancer (BCLC 2026) guideline

Role of systemic treatment

BCLC-B: Diffuse, infiltrative, extensive bilobar liver involvement

BCLC-C: Portal invasion and/or extrahepatic spread, preserved liver function, Performance 1-2

Hepatocellular carcinoma



Systemic treatment

First-line

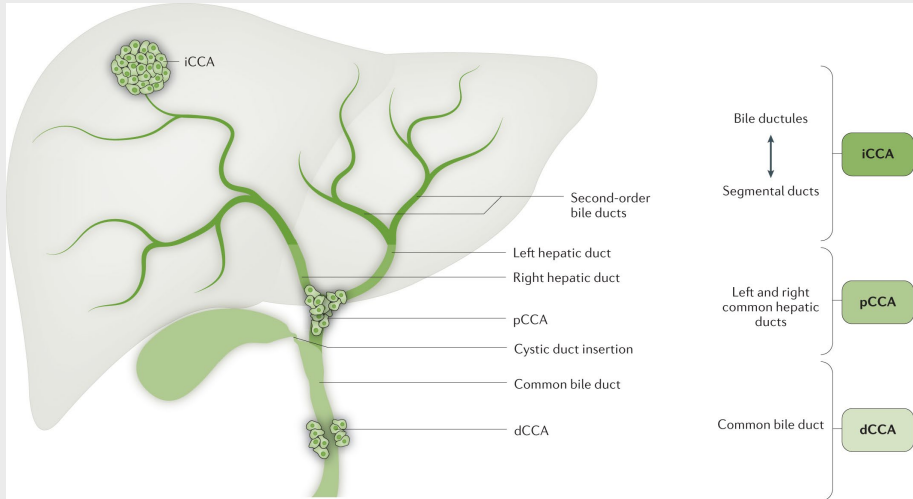
- Bevacizumab + Atezolizumab, Durvalumab + Tremelimumab, Ipilimumab + Nivolumab (preferred)
- Sorafenib, Lenvatinib
- Durvalumab

Second-line

- Targeted drugs: Regorafenib, Cabozantinib, Ramucirumab
- Immunotherapy: Nivolumab + Ipilimumab, Pembrolizumab

Cholangiocarcinom

Cholangiocarcinoma



- Malignant tumor of bile duct epithelium (cholangiocyte)
- Classified by anatomy into
 - Intrahepatic CCA
 - Peri-hilar CCA
 - Distal CCA
- Etiology
 - Fluke-related
 - : *Opisthorchis viverrine* (Southeast Asia)
 - : *Clonorchis sinensis* (Korea, China)
 - Non-fluke-related
 - : No identifiable-cause
- Classified by anatomy into
- Targetable mutation: IDH1 mutation, FGFR2 fusion, HER2 amplification

Clinical presentation



Clinical Presentation

- Obstructive jaundice (esp. in peri-hilar and distal CCA)
- Right upper quadrant pain
- Chronic dyspepsia
- Anorexia, weight loss

Investigation

- US: liver mass, IHD dilatation
- CT abdomen: Hypodense lesion, subcapsular retraction, perilesional bile duct dilatation



Treatment



Resectable (localized disease)

- Curative aim
- Surgery followed by adjuvant capecitabine

Unresectable (locally advanced, distant metastasis)

- Palliative aim
- Biliary drainage
- Systemic therapy (CMT +/- IO)
 - Cisplatin + Gemcitabine + Durvalumab/Pembrolizumab
 - Cisplatin + Gemcitabine
 - Targeted therapy
(second line setting if driver mutation detected: FGFR2, IDH1, HER2)

colorectal cancer

colon cancer



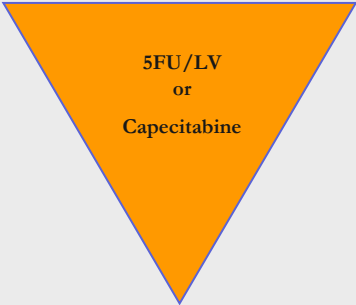
| Stage | Treatment |
|-------|--|
| I | Surgery |
| II | Surgery (+ Adjuvant 5FU in high risk patient) |
| III | Surgery + Adjuvant FOLFOX |
| IV | Surgery + Metastasectomy + Adjuvant CMT in oligometastatic setting Palliative CMT + targeted agent in unresectable case |

- **High risk stage 2**

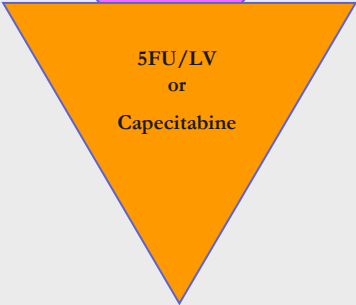
- T4, poorly differentiated histology, bowel perforation/obstruction, closed/positive margin
- Inadequate LN sampling (<12 LN)
- Lymphovascular invasion, Perineural invasion

- **Consider RT in rectal cancer**

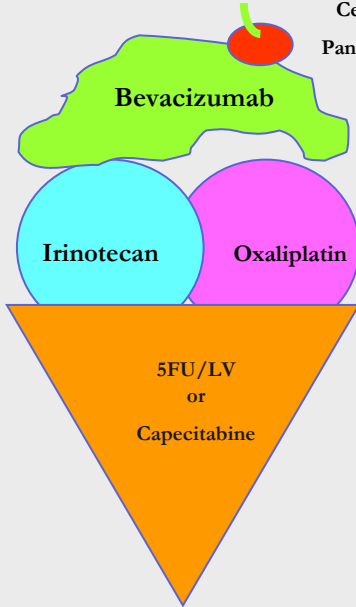
CMT in colon cancer



Stage 2



Stage 3



Stage 4

Cetuximab
Panitumumab

Palliative CMT in colon cancer



1. Single agent 5FU + LV (can be replaced by oral Capecitabine)

2. Better regimen (Doublet/Triplet Chemo)

2.1 Oxaliplatin + 5FU/LV : FOLFOX

2.2 Irinotecan+ 5FU/LV : FOLFIRI

2.3 Oxaliplatin + Irinotecan+ 5FU/LV : FOLFOXIRI

3. Best regimen (Targeted therapy + Chemotherapy)

3.1 For RAS/BRAF WT and left-sided tumor

: FOLFOX/FOLFIRI + Cetuximab/Panitumumab

3.2 Others

: FOLFOX/FOFIRI + Bevacizumab, FOLFOXIRI + Bevacizumab

: Cetuximab + Encorafenib in second line setting for BRAF V600E mutant

GASTRIC cancer

Gastric cancer



- **High incidence:** East Asia (Japan, Korea, China), Peak age: > 60 years
- **Key risk factors**
 - Helicobacter pylori infection (most important),
 - Chronic atrophic gastritis
 - Smoking
 - Diet: smoked, salted foods
 - Family history / hereditary syndromes
 - : CDH1 mutation → diffuse type
- **Pathology subtype (Lauren classification)**
 - Diffuse type (Linitis plastica): No clear mass, thickened rigid
 - Intestinal type: Gland-forming, H pylori-associated



Diffuse



Intestinal

Gastric cancer



- **Clinical Presentation**

- Epigastric pain, Weight loss, Early satiety, Nausea/vomiting, Anemia (iron deficiency), GI bleeding

- **Classic metastasis sites**

- **Virchow node** (left supraclavicular)
- **Sister Mary Joseph nodule** (umbilical)
- **Krukenberg tumor** (ovarian mets)
- **Blumer shelf** (rectal shelf)
- Peritoneal, Liver metastasis

- **Treatment for advanced stage**

- Chemotherapy +/- targeted therapy (HER2, Claudin)
+/- Immunotherapy



Krukenberg Tumor

germ cell tumor

Germ cell tumor



- **Malignancy of primordial germ cell**
- **Primary gonadal site (Testicular) 95%**
- **Relatively rare for extragonadal sites**
 - : Midline mass, mediastinum, retroperitoneum, pineal gland
- **Common in Male, Age : 20-40 yr**
- **Chemosensitive tumor**
- **Risk factors**
 - : Cryptorchidism, Hypospadias, Klinefelter's syndrome (mediastinal GCT), i(12p)

Clinical presentation



- Testicular mass
: Painless mass (almost always unilateral)
- Retroperitoneal mass
: Back pain, Palpable mass
- Mediastinal mass
: SVC syndrome, RS symptom
- Intracranial mass (Pineal/Pituitary)
: Parinaud syndrome, DI, Hydrocephalus, Hypopituitarism
- Gynecomastia
: high level of beta-hCG

Mnemonics

Characteristics of Parinaud's Dorsal Midbrain Syndrome with "CLUES"

CLUES

- C**onvergence retraction nystagmus
- L**ight Near Dissociation
- U**pgaze paralysis
- E**yelid retraction
- S**kew deviation



Tumor marker



| | | Beta-hCG | AFP |
|--------------|--------------------------|-------------------|--------------|
| Seminoma | | Mildly elevated ↑ | Never rising |
| Non-seminoma | Embryonal cell carcinoma | ↑ | ↑ |
| | Choriocarcinoma | ↑↑↑ | ↔ |
| | Yolk sac tumor | ↔ | ↑↑↑ |
| | Teratoma | ↔ | ↔ |
| | Mixed germ cell tumor | ↑ | ↑ |

False positive AFP: HCC, Hepatoid variant adenocarcinoma, Cirrhosis

False positive Beta-hCG: Marijauna, Gestational trophoblastic neoplasm, Malignancy, Marijauna

CMT in advanced Germ cell tumor



Standard therapy: BEP x 4 cycles

Cisplatin 20 mg/m² d 1-5
Etoposide 100 mg/m² d 1-5
Bleomycin 30 IU d 1,8,15
Q 3 weeks

Alternative therapy: VIP x 4 cycles

Cisplatin 20 mg/m² d 1-5
Etoposide 75 mg/m² d 1-5
Ifosfamide 1200 mg/m² d 1-5
Q 3 weeks

BEP regimen

- Bleomycin : pulmonary fibrosis
- Etoposide : hypersensitivity and cardiotoxicity
- Cisplatin : N/V, renal toxic, neuropathy

Prefer VIP to BEP if underlying lung disease or plan to RT chest

gist

Gastro intestinal stromal tumor (gist)



- Spindle cell neoplasm of the gastrointestinal tract
- Originated from interstitial cell of Cajal
- Presentation: accidental finding from CT/EGD, mass effect, GI bleeding
- Primary site : Stomach, small bowel, rectum, mesentery, retroperitoneum
- Common metastasis site: Liver
- IHC : Positive for CD117 (C-kit), DOG-1
- **Rx : Early stage** › **Surgery** (+ adjuvant imatinib for 3 years)
 : Advanced › **Tyrosine kinase inhibitor : Imatinib**
- GIST is chemoresistant tumor, no role of conventional chemotherapy !

Prostate cancer

Prostate cancer



Clinical

Locally advanced : LUTs, Bladder outlet obstruction, Hematuria

Advanced : Bone pain (Blastic bone metastasis), Cord compression,
Myelophthisis

Lab: PSA > 4 + Abnormal digital rectal exam

Risk stratification : PSA, Gleason and T-stage

Pathology : Adenocarcinoma, CK7-, CK20-, PSA+, NKX 3.1+

treatment



Local disease

- Low risk: radical prostatectomy or RT
- Intermediate risk : radical prostatectomy with pelvic LN dissection or RT
- High/Very high : RT with long-term ADT or radical prostatectomy with pelvic LN dissection

Locally advanced : Rx as high risk

Metastasis : ADT, Novel anti-androgen, CMT

* ADT: Androgen Deprivation treatment

treatment



Androgen deprivation therapy (ADT)

- Bilateral orchidectomy
- LHRH analogues : leuprolide, goserelin, buserelin

Novel anti-androgen treatment

- CYP-17 inhibitor: abiraterone
- New generation anti-androgen: enzalutamide, apalutamide, darolutamide

Side effects of



Less sexual desire

Impotence

Hot flush

**Breast tenderness and growth of
breast tissue**

Osteoporosis

Anemia

Decreased mental sharpness

Loss of muscle mass

Weight gain

Fatigue

Hypercholesterol

Depression

Bone metastasis in prostate cancer



Mostly osteoblastic lesion

Relate to level of PSA

Bone pain

Fracture

Malignant cord compression

Rx :

Local RT

Bisphosphonate or Denosumab

04

Cancer of unknown primary

Cancer of unknown primary



IHC marker for determining cell lineage

| Markers* | Most Likely Cell Lineage |
|------------------------------------|--------------------------|
| Pan-keratin (AE1/AE3 & CAM5.2) | Carcinoma |
| CK5/6, p63/p40 | Squamous cell carcinoma |
| S100, SOX10 | Melanoma |
| LCA± CD20± CD3± | Lymphoma |
| OCT3/4± SALL4± | Germ cell tumor |
| WT1, calretinin, mesothelin, D2-40 | Mesothelial tumor |

*These markers are not uniformly specific or sensitive and can be present on other tumors.

Cancer of unknown primary



IHC marker for Adenocarcinoma

CK7+ 20+ "CUP-O"

- **Cholangiocarcinoma**
- **Urothelial carcinoma**
- **Pancreatic carcinoma**
- **Ovarian cancer**

CK7+ 20- "Upper thorax"

- **Thyroid**
- **Breast**
- **Lung**
- **Pancreas, CCA, Ovary (non-mucinous)**

CK7- 20+ "GI malignancy"

- **Colorectal cancer**
- **Gastric cancer**
- **Merkel**

CK7- 20-

- **Prostate**
- **HCC**
- **RCC**
- **Squamous cell**

Cancer of unknown primary



IHC marker for Adenocarcinoma, CK7+ 20-

CK7+ 20- "Upper thorax"

- **Thyroid**
- **Breast**
- **Lung**
- **Pancreas, CCA, Ovary (non-mucinous)**

| Cancer | Additional positive IHC |
|---------|------------------------------|
| Thyroid | TTF-1, Thyroglobulin |
| Breast | GATA3, Mammaglobin, GDCFP-15 |
| Lung | TTF-1, Napsin A |

Cancer of unknown primary



| Primary markers | Cancer type | Additional markers to consider |
|--------------------|---|---|
| CK7+,CK20- | Lung [NSCLC (adenocarcinoma) and SCLC] | TTF1, SMARCA4, synaptophysin |
| | Thyroid | Thyroglobulin, TTF1, PAX8 |
| | Breast | GATA3, SOX10, ER, PgR, Mammaglobin, BRST1 |
| | Upper GI, pancreaticobiliary | CDX2, CK19, SMAD4, ARID1A, BAP1 |
| | Endometrial, endocervical, ovary (serous) | PAX8, ER, PgR, WT1, p53 |
| | Renal (papillary) | PAX8, PAX2, racemase, CD10 |
| | Salivary gland | GATA3, S100, SOX10, AR, HER2 |
| | Bladder | GATA3, p63 |
| CK7+, CK20+ | Bladder | GATA3, p63 |
| | Upper GI, pancreaticobiliary | CDX2, CK19, SMAD4, ARID1A, BAP1 |
| | Rectum | CDX2, SATB2 |
| CK7-, CK20+ | Colorectal, upper GI | CDX2, SATB2 |
| | Merkel cell | Synaptophysin |
| CK7-, CK20- | Renal | PAX8, PAX2, racemase, CD10 |
| | Hepatocellular | Arginase1, HepPar1 |
| | Germ cell | SALL4, PLAP |
| | Prostate | PSMA, NKX3.1 |
| | Gastric | CDX2 |
| | SCLC | TTF1, SMARCB1, synaptophysin |
| | Adrenal cortical | SF1, calretinin, inhibin |
| | Neuroendocrine | INSM1, synaptophysin |
| | Squamous cell | p40, p63, CK5/6 |

Favorable CUP

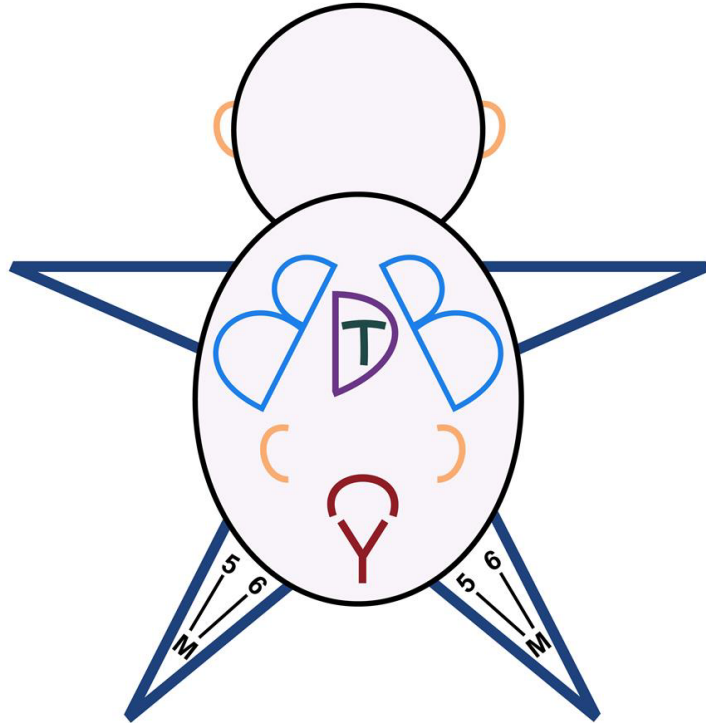


| Histologic type | Clinical feature | Treatment recommendation |
|---|--|---|
| Adenocarcinoma | Women with isolated axillary adenopathy | Treat as stage II breast cancer |
| | Women with peritoneal carcinomatosis | Treat as stage III ovarian cancer |
| | Men with elevated PSA or blastic bone metastases | Treat as advanced prostate cancer |
| Adenocarcinoma or PDC | Single metastatic lesion | Definitive local therapy (resection and/or radiation therapy) |
| Squamous cell | Cervical adenopathy | Treat as head and neck cancer with involved neck nodes |
| | Inguinal adenopathy | Inguinal node dissection Consider concurrent radiation therapy/chemotherapy (as in locally advanced cervical cancer) |
| Poorly differentiated carcinoma | Young men with midline tumor or elevated hCG/AFP | Treat as extragonadal germ cell tumor |
| | Other clinical presentations | Empiric platinum/paclitaxel chemotherapy |
| Poorly differentiated neuroendocrine carcinoma | Diverse clinical presentations | Treat with platinum/etoposide or paclitaxel/platinum/etoposide |

05

Side effects of systemic therapy

Side effect of chemotherapy



Cisplatin and Carboplatin

- ototoxicity
- nephrotoxicity

Vincristine

- peripheral neuropathy

Bleomycin and Busulfan

- pulmonary fibrosis

Trastuzumab and Doxorubicin

- cardiotoxicity

Cyclophosphamide

- hemorrhagic cystitis

Methotrexate, 5-FU, and 6-MP

- myelosuppression

Pulmonary toxicities



Bleomycin

- Induces reactive oxygen radicals
- Acute- Pneumonitis, ARDs
- Late – Lung fibrosis



Cardiotoxicities



5-FU

- Coronary spasm

Osimertinib

- QT prolongation

Anti-VEGF

- Hypertension, bleeding, thrombosis

Cardiotoxicity: Cardiomyopathy



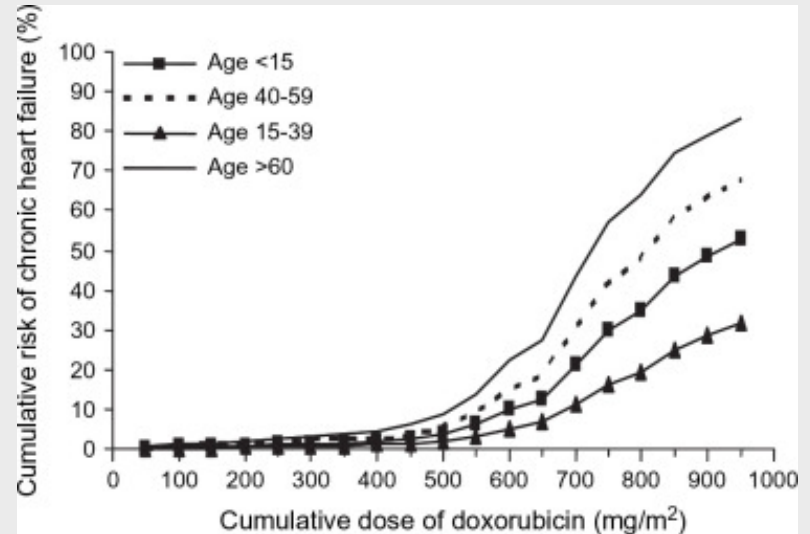
Doxorubicin

- Dose dependent, Irreversible
- Should not exceed cumulative dose of 350-450 mg/m²

Trastuzumab

- Dose independent, Reversible

Follow up echocardiography is mandated !



renal toxicities



Cisplatin

- Renal tubular damage (Fanconi syndrome)
- AKI
- SIADH

Cyclophosphamide

- Hemorrhagic cystitis from its active metabolite "acrolein"
- Prevention: mesna

Anti-VEGF

- Proteinuria

neuro toxicities



Oxaliplatin

- Acute – sensitive to cold
- Late – dose dependent peripheral neuropathy

Taxane (Paclitaxel, Docetaxel) / **Vincristine**
: Peripheral neuropathy

Ifosfamide, MTX : Encephalopathy

GI toxicities



5-FU, Capecitabine

- Mucositis, diarrhea

Oxaliplatin

: Veno-occlusive disease (blue liver)

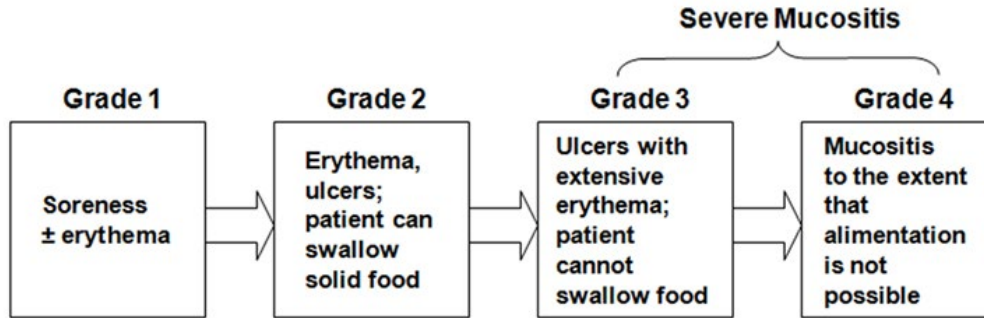
Irinotecan

- : Acute diarrhea (parasympathetic effect)
- : Late diarrhea from metabolite
- : Steatohepatitis (yellow liver)

mucositis



World Health Organization's Oral Toxicity Scale



Chemotherapy

MTX, 5-FU,
Cyclophosphamide,
Docetaxel, Doxorubicin,
Paclitaxel,

Targeted therapy

Everolimus, Sorafenib

Prevent

: Ice chip for 5-FU

Rx

: Systemic analgesic therapy
with narcotics

skin toxicities



Hand-foot syndrome and Hand foot skin reaction

- Capecitabine
- Tyrosine kinase inhibitor



Flagellate erythema

- Bleomycin



PRIDE syndrome EGFR Anti



PRIDE syndrome

- **P**apulopustular (acneiform) rash and/or paronychia
- **R**egulatory abnormalities of hair growth
- **I**tching
- **D**ryness (xerosis)
- **E**GFR inhibitors: Erlotinib, gefitinib, Osimertinib, Panitumumab, Cetuximab

Others

- Mucositis
- Photosensitivity

Secondary leukemia



Cyclophosphamide/Ifosfamide

- AML in 5–7 yrs
- MDS
- -5, -7 del
- AML: M1, M2

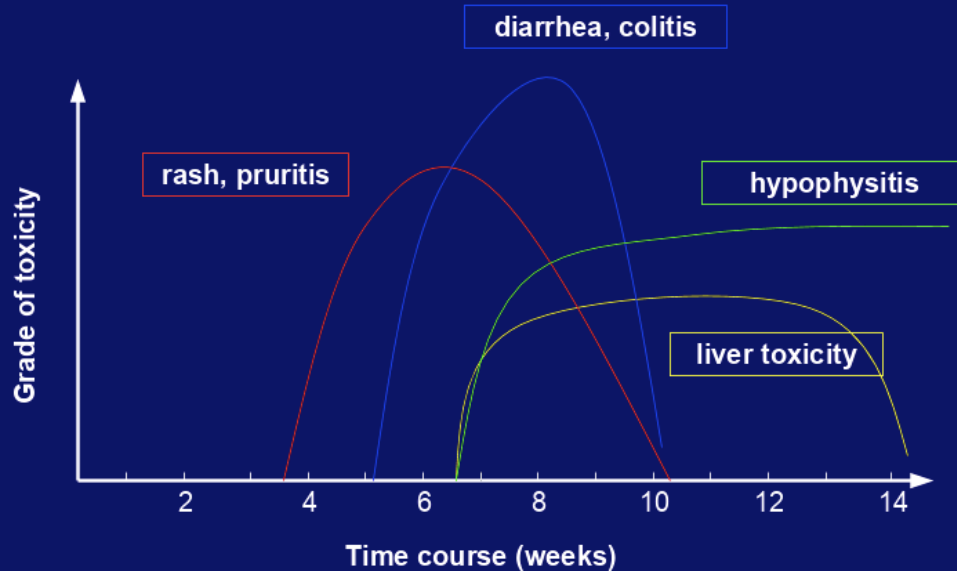
Etoposide

- AML in 1–3 yrs
- t(11q23)
- AML: M4, M5

immunotherapy



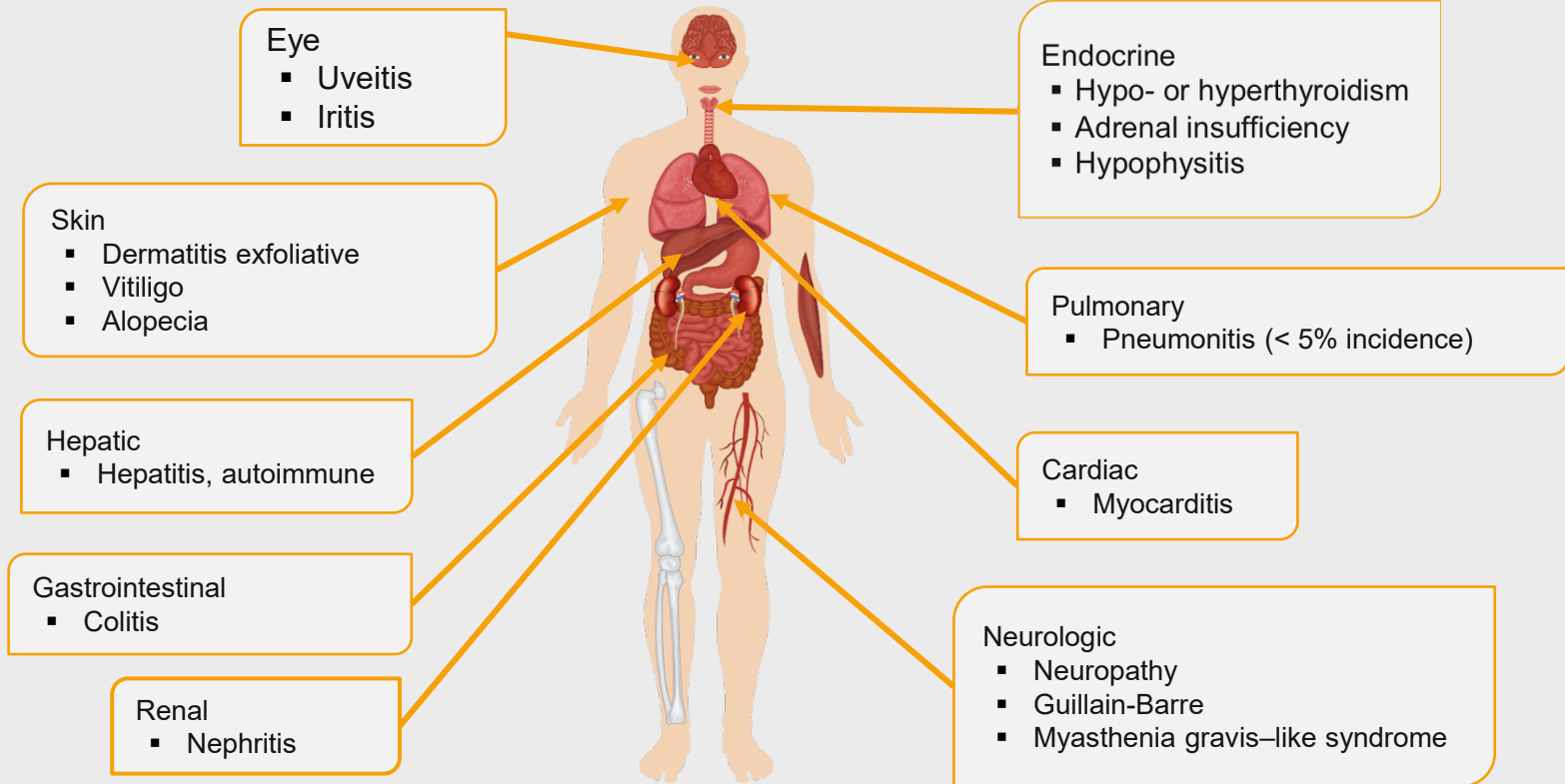
Kinetics of Appearance of irAEs¹



1. Weber JS, et al. *J Clin Oncol* 2012;30:2691-7.

- Immunotherapy functions by enhancing host immune system ability to clear cancer cell but it can also lead to immune mediated damage to healthy native tissue called Immune-related Adverse Events (irAEs)

immunotherapy



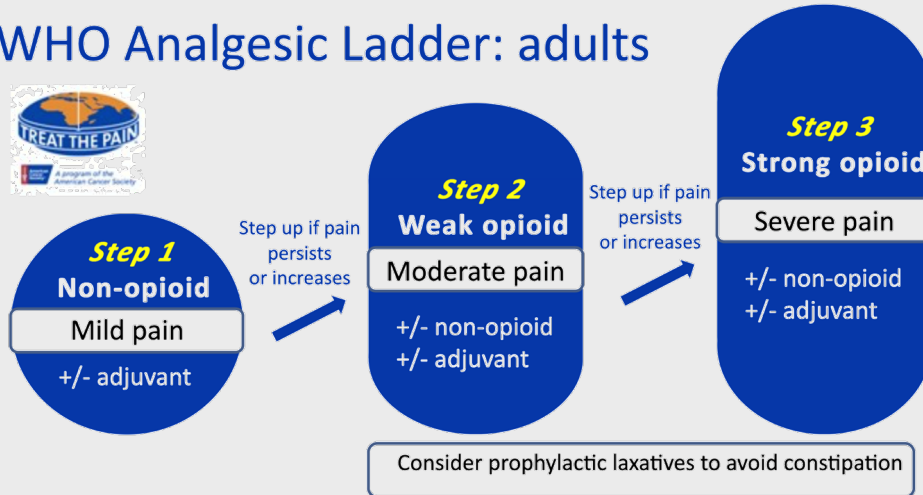
06

Palliative care

Cancer pain management



WHO Analgesic Ladder: adults



Numerical rating scale

| | |
|----------|----------|
| Mild | 1 2 3 |
| Moderate | 4 5 6 |
| Severe | 7 8 9 10 |

- Non-opioids** | ibuprofen or other NSAID, paracetamol (acetaminophen), or aspirin
- Weak opioids** | codeine, tramadol, or low-dose morphine
- Strong opioids** | morphine, fentanyl, oxycodone, hydromorphone, buprenorphine
- Adjuvants** | antidepressant, anticonvulsant, antispasmodic, muscle relaxant, bisphosphonate, or corticosteroid

Combining an opioid and non-opioid is effective, but do not combine drugs of the same class.

Time doses based on drug half-life ("dose by the clock"); do not wait for pain to recur

OPIOID Usage



- Starting with short-acting opioid, morphine IR/syrup 5-7.5 mg as needed
- If ≥ 4 dose needed per day, consider long-acting background opioid on total dose using previous 24 hr
- Consider pain adjunctive medication e.g. TCA/anti-convulsant in neuropathic pain

Breakthrough dose : 10-20% of background dose

If breakthrough dose using ≥ 4 dose per day,
reset new background dose and breakthrough dose

Pain crisis management



- Use intravenous bolus (peak effect 15 min) or subcutaneous route (peak effect 30 min)
- For opioid naïve: 2-5 mg of morphine
For opioid tolerant: 10-20% of total opioid taken in previous 24 hr

If pain unchanged, increased dose by 50-100%

If pain decreased but inadequately controlled, repeat same dose

If pain improved and adequately controlled, continue current dose as needed

OPIOID conversion



| Before | Conversion Factor | After |
|-----------------------------|--------------------------|------------------------------------|
| Morphine po 60 mg | $\div 3$ | Morphine IV 20 mg |
| Morphine po 60 mg | = | Kapanol 60 mg |
| Morphine po 60 mg | = | Fentanyl patch 25 mcg/hr |
| Fentanyl IV | = | Fentanyl patch |

dyspnea crisis



- **Use intravenous/subcutaneous route**
- **Non-specific measure**
 - ▶ Calm reassurance
 - ▶ Positioning
 - ▶ Oxygen
 - ▶ Opioids
 - ▶ Possibly sedative: midazolam, lorazepam

Death rattle



- **Positioning**
- **Antisecretory agents**
 - ▶ Atropine 0.4–0.8 mg SC q 1 hr PRN
 - ▶ Atropine eye drop 1% 4 drops SL q 4 hr PRN
- **Consider suctioning if secretions are**
 - ▶ Distressing
 - ▶ Proximal
 - ▶ Accessible

Overview of Oncology 2026 For Resident

THANK you

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