

2 Mid

L7

genetic

Feature	Osteoarthritis (OA)	Rheumatoid Arthritis (RA) <i>Synovial</i>	Juvenile Idiopathic Arthritis (JIA) <i>genetic</i>	Suppurative Arthritis	Lyme Arthritis
Etiology	Degeneration of cartilage; aging or secondary to preexisting disease	Autoimmune; chronic inflammatory synovitis <i>Non-suppurative</i>	Unknown; autoimmune mechanism similar to RA	Bacterial infection (hematogenous spread)	Infection with <i>Borrelia burgdorferi</i>
Age Group	>50 years	4th-5th decade (F:M = 3:1)	<16 years <i>children</i>	Any age; children: <i>H. influenza</i> <2 years, <i>S. aureus</i> older children/adults, <i>N. gonorrhoeae</i> in young adults <i>sickle cell</i> <i>G salmonella</i>	Any age
Joints Affected	Few weight-bearing joints (knee, hip, spine, hands)	Symmetric polyarthritis (hands, feet, wrists, ankles, MCP, PIP joints) <i>Fibrous Ankylosis</i> <i>Bony Ankylosis</i> <i>destruc</i>	Large joints > small joints; oligoarthritis more common <i>1/2 Bone</i>	Mostly knee, but can affect other joints	Large joints (knees most common)
Pathogenesis	Cartilage degeneration > repair and proliferation	T-cell activation → IFN-γ, IL-17, TNF, IL-1 → synovial inflammation, cartilage destruction, bone resorption	Similar to RA but ANA (+) seropositivity is common, Rheumatoid Factor usually absent <i>RFX nodules X</i>	Direct bacterial infection causing purulent inflammation	Immune response to <i>Borrelia</i> infection
Clinical Features	Joint pain worsens with use, morning stiffness, crepitus, limited range of motion, osteophyte formation, radicular pain	Symmetric joint swelling, pain, warmth, stiffness in morning and after inactivity, ulnar deviation, systemic involvement (skin, heart, lungs, vessels) <i>2) Boyltoniere</i> <i>3) Swan neck</i>	Similar to RA, but systemic disease and large joint involvement more common <i>VV</i>	Acute onset of severe pain, swollen warm joint, fever, leukocytosis, ↑ ESR <i>Erythrocytes</i>	Intermittent arthritis attacks, often after initial Lyme disease rash
Serology & Markers	None specific	RF+ (80%), ACPA+ (70%)	ANA+ common, RF- usually	↑ WBC, ↑ ESR, positive culture/gram stain	Positive Lyme serology (+)
Treatment	Pain control (NSAIDs, intra-articular steroids), joint replacement for severe cases	Steroids, Methotrexate (MTX), Anti-TNF	Variable prognosis; treatment depends on severity	Joint aspiration + IV antibiotics	Antibiotics (e.g., doxycycline, amoxicillin)

genetic + environment



* A&A seronegative
 → Treat: Anti-IL-17
 → Absence RF
 → ligament
 → sacroiliac joint
 HLA-B27
 Bony Ankylosis (fusion)
 Ankylosing spondylitis
 Reiter syndrome
 Enteropathic salmonella, shigella
 psoriatic
 DIP joint
 ULA-B27 (sacroiliac)
 arthritis
 urethritis
 conjunctivitis
 Bacteria → Autoimmune

Good

Tumor Type	Benign/Malignant	Common Location	Key Features	Genetic Mutation	Treatment
Joint Tumors	Mostly benign	Wrist (ganglion cyst), knee (Baker's cyst)	Ganglion cyst: not a true cyst, no synovial communication; Baker's cyst: true synovial cyst	No specific mutation	Surgical removal
Tenosynovial Giant Cell Tumor	Benign	Large joints, small hand tendons	Diffuse (PVNS) or localized	t(1;2)(p13q;37) (Type VI collagen α-3)	Surgical excision
Soft Tissue Sarcomas	Malignant	Extremities (thigh), deep soft tissues	Hematogenous metastasis, mostly sporadic, aggressive	NF1, Gardner syndrome, Li-Fraumeni syndrome, Osler-Weber-Rendu syndrome	Surgery, chemotherapy (CT), radiation therapy (RT)
Lipoma	Benign	Subcutaneous tissue	Well-encapsulated, mature fat cells	No specific mutation	Excision
Liposarcoma	Malignant	Extremities, retroperitoneum	Three subtypes: WD (less aggressive), myxoid, pleomorphic (most aggressive)	WD: MDM2 gene (chr 12), Myxoid: t(12;16)	Surgery, CT
Nodular Fasciitis	Benign	Subcutaneous soft tissues <i>Trauma</i>	Rapid growth, often mistaken for malignancy	t(17;22) MYH9-USP6 fusion gene	Self-limiting, excision if needed
Fibromas	Benign	Skin, subcutaneous tissue	Common, <u>fibroblastic</u> proliferation	No specific mutation	None required
Fibrosarcoma	Malignant	Soft tissues (<i>superficial cutaneous</i>)	<u>Cellular</u> , storiform pattern, high <u>mitotic rate</u>	No specific mutation	Surgery, CT
Superficial Fibromatoses	Benign	Palmar fascia, plantar fascia, penile region	Infiltrative, may impact function	No specific mutation	Surgery if needed
Deep Fibromatosis (Desmoid Tumor)	Locally aggressive	Abdominal wall, mesentery, limbs	Doesn't metastasize but recurs, infiltrative	CTNNB1 (β-catenin), APC (FAP syndrome)	Surgery, recurrence common
Rhabdomyoma	Benign	Associated with tuberous sclerosis	Rare	No specific mutation	None required
Rhabdomyosarcoma	Malignant	Children, skeletal muscle	Three types: embryonal, alveolar, pleomorphic	Specific mutations (not specified)	Surgery, CT, RT
Leiomyoma	Benign	Uterus (fibroids), skin, soft tissue	Well-circumscribed, hormonally responsive	Some cases: Fumarate hydratase (1q42.3) ✗	Surgery if symptomatic
Leiomyosarcoma	Malignant	Extremities, deep soft tissue, retroperitoneum, great vessels	Hemorrhage, necrosis, high mitotic rate	Complex genotypes	Surgery, CT
Synovial Sarcoma	Malignant	Deep soft tissues, extremities	Monophasic (spindle cells) or biphasic (spindle + glands)	t(X;18)(p11;q11) SS18 fusion gene	Surgery + CT
Undifferentiated Pleomorphic Sarcoma (UPS)	Malignant	Deep soft tissue, extremities	Highly pleomorphic, aggressive, formerly MFH	Complex genetic abnormalities	Surgery, CT, poor prognosis

L8

P. fibrous

L9 + 10

skeletal muscle

smooth muscle

Tumor Type	Correlated Mutation
Tenosynovial Giant Cell Tumor	t(1;2)(p13q;37) (Type VI collagen α -3)
Liposarcoma (Well-Differentiated)	MDM2 gene (chr 12)
Liposarcoma (Myxoid Type)	t(12;16)
Nodular Fasciitis	t(17;22) MYH9-USP6 fusion gene
Deep Fibromatosis (Desmoid Tumor)	CTNNB1 (β -catenin) or APC gene (FAP syndrome)
Leiomyoma (Subset Cases)	Fumarate hydratase (1q42.3)
Synovial Sarcoma	t(X;18)(p11;q11) SS18 fusion gene
Soft Tissue Sarcomas (Some Cases)	NF1, Gardner syndrome, Li-Fraumeni syndrome, Osler-Weber-Rendu syndrome
Undifferentiated Pleomorphic Sarcoma (UPS)	Complex genetic abnormalities

18

difficult diagnose

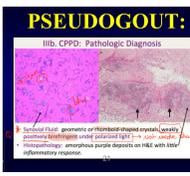
pseudogout
Idiopathic secondary

> 50 age
inflamm response
by deposit calcium phosphate

CPPD crystal
Risk factor:
DM, previous joint damage
HPTH, Hemochrom

Acute subacute chronic
Tr: supportive

Histologic:
Rhomboid-shape crystals
⊕ birefringent → not needle shape



Gout
earilage synovium

Big toe
inflamm response
by deposit urate

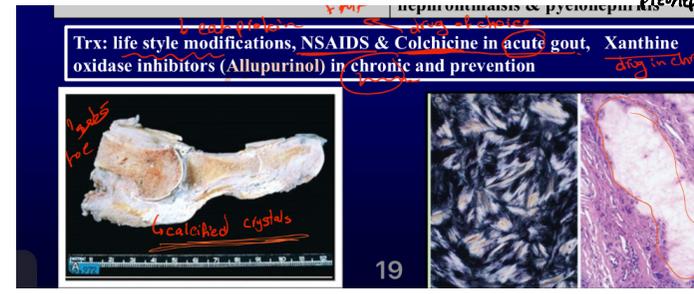
Deposition MSU crystals
↑ uric acid
Risk factors 20-30y, obesity
Alcohol, genetic predispos
Thiazide

Acute: NSAIDs colchicine
chronic: xanthine (Allupurinol)

Acute Arthritis
Chronic Tophaceous Arthritis → thick synovium pannus
Tophi → cartilage, ligament bursa, tendon
Gouty nephropathy → deposition crystals in kidney

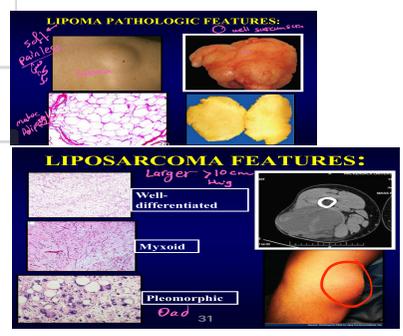
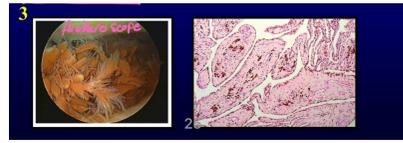
True cyst: baker cyst around knee (herniation process)

Joint tumor
Rare
common: Ganglion cyst, tenosynovial gainth
Not true
close to joint wrist



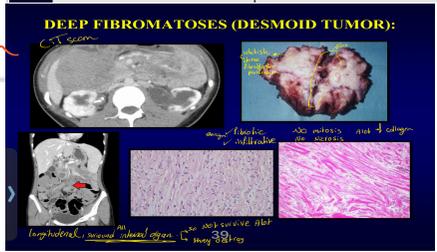
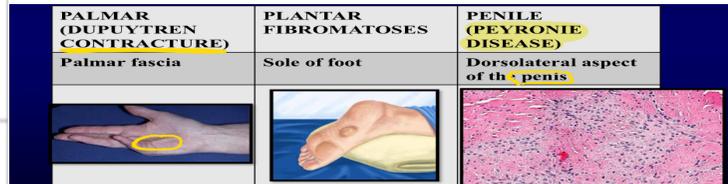
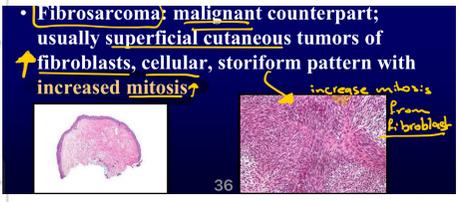
كيس زلالي (تورانيا) نسيجية

Tenosynovial Giant Cell Tumor	Benign	Large joints, small hand tendons	Diffuse (PVNS) or localized Pigmented (brown), villonodular synovitis.	t(12)(p13q37) (Type VI collagen α-3)	Surgical excision
Soft Tissue Sarcomas Ex: Ewing sarcoma	Malignant	Extremities (thigh), deep soft tissues	Hematogenous metastasis, mostly sporadic, aggressive Not genetic	NF1, Gardner syndrome, Li-Fraumeni syndrome, Osler-Weber-Rendu syndrome grade stages	Surgery, chemotherapy (CT), radiation therapy (RT)
Lipoma common	Benign	Subcutaneous tissue	Well-encapsulated, mature fat cells	No specific mutation	Excision
Liposarcoma	Malignant Large	Extremities, retroperitoneum Common in adult >50	Three subtypes: WD (less aggressive), myxoid, pleomorphic (most aggressive)	WD: MDM2 gene (chr 12), Myxoid: t(12;16)	Surgery, CT



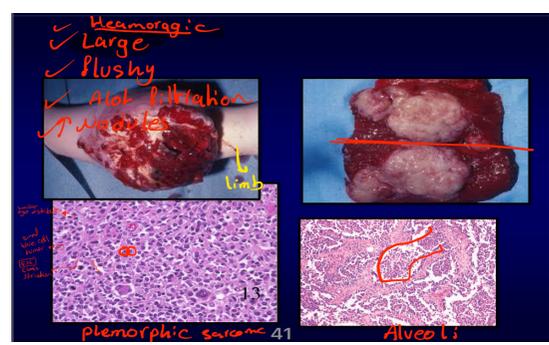
L9 L10 Fibrous Tumor:

Nodular Fasciitis	Benign	Subcutaneous soft tissues <i>Trauma</i>	Rapid growth, often mistaken for malignancy	t(17;22) MYH9-USP6 fusion gene	Self-limiting, excision if needed
Fibromas	Benign	Skin, subcutaneous tissue	Common, fibroblastic proliferation	No specific mutation	None required
Fibrosarcoma	Malignant	Soft tissues (<i>superficial cutaneous</i>)	Cellular, storiform pattern, high mitotic rate	No specific mutation	Surgery, CT
Superficial Fibromatoses	Benign	Dupuytren's contracture Palmar fascia, plantar fascia, penile region (<i>Peyronie disease</i>)	Infiltrative, fibroblast <i>exception</i>	No specific mutation	Surgery if needed
Deep Fibromatosis (Desmoid Tumor)	Locally aggressive	Abdominal wall, mesentery, limbs	Doesn't metastasize but recurs, infiltrative, <i>female > male</i>	CTNNB1 (β-catenin), APC (FAP syndrome)	Surgery, recurrence common <i>complete excision</i>

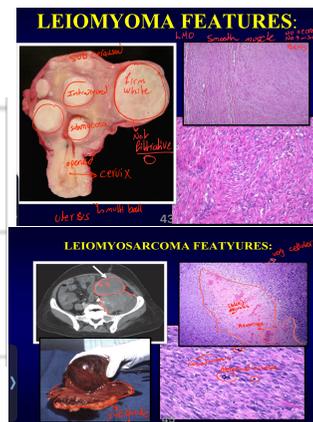


skeletal muscle Tumor:

Rhabdomyoma	Benign <i>exception</i>	Associated with tuberous sclerosis	Rare	syndrome	No specific mutation	None required
Rhabdomyosarcoma RMS	Malignant	Children, skeletal muscle <i>Common</i>	Three types: embryonal, <i>Common</i> alveolar, pleomorphic	Specific mutations (not specified)	Surgery, CT, RT <i>Radiotherapy</i>	↑ high grade



Smooth muscle Tumor



Leiomyoma	Benign	Uterus (fibroids), skin, soft tissue	Well-circumscribed, hormonally responsive	Some cases: Fumarate hydratase (1q42.3) X	Surgery if symptomatic
Leiomyosarcoma	Malignant	Extremities, deep soft tissue, retroperitoneum, great vessels <i>Adult, female</i>	Hemorrhage, necrosis, high mitotic rate	Complex genotypes	Surgery, CT <i>بتميز على location, size, grade</i>

uncertain origins-

<u>Synovial Sarcoma</u>	Malignant metastasis lung + lymph <i>exception</i>	<i>→ Around knee</i> Deep soft tissues, extremities <i>Adult (20-40)</i>	Monophasic (spindle cells) or biphasic (spindle + glands)	<u>t(X:18)(p11;q11) SS18 fusion gene</u> <i>confirm diagnose</i>	Surgery + CT <i>→ chemo therapy</i> <i>(excision) stages</i>
Undifferentiated Pleomorphic Sarcoma (UPS) <i>سkeletal</i> <i>صغير</i> <i>صغير</i>	Malignant <u>high grade mesenchymal</u>	Deep soft tissue, extremities	Highly pleomorphic, aggressive, formerly MFH <i>سماييل</i> <i>Tumor: ugly (anaplastic)</i> <i>pleomorphic</i> <i>Abnormal mitosis</i> <i>necrosis</i>	Complex genetic abnormalities	Surgery, CT, poor prognosis <i>RT</i>



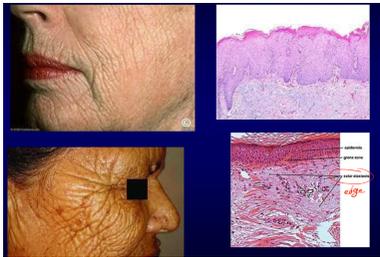
L11

Solar elastosis

sun damage

thick + yellow skin

Damage of collagen + elastic fiber



Actinic keratosis

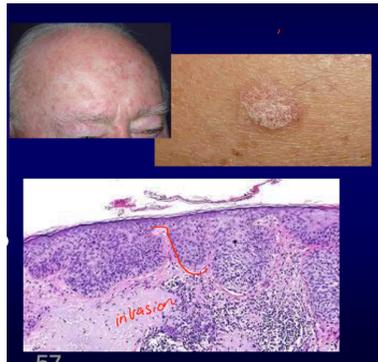
Pre-malignant

↳ progress: squamous cell carcinoma *in situ*

UV light damage

Tp53 mutation.

↳ localised superficial.



Seborrheic keratosis

middle / older

Benign tumor.

* Very common

• FGFR3 mutations

clinical,

↳ coin like

↳ pigmented [↑]↑

↳ elevated (stuck on)

Seborrheic keratosis:



(Epidermal) Epithelial inclusion cyst

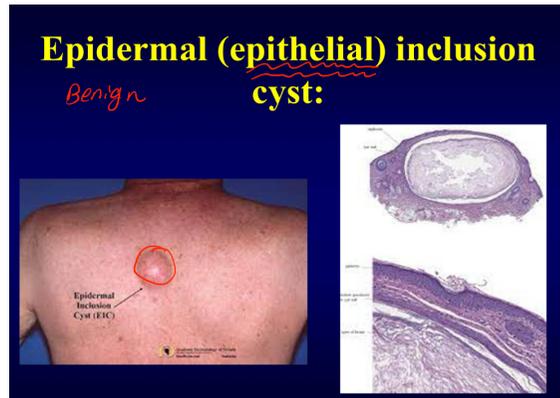
Skin
DUMP
Benign
(Remove excision)

Dermoid cyst

- (EIC)
- Benign
 - fill with keratin debris
 - Rupture / not

- (mature) element ^{Wiz}
(bone, hair, muscle, collagen)
- greasy yellow material
- (immature) (malignant) ^{ناجراً}
ovary/testis ^{بيوت} Metast

لازيم سا عسر
ف، لفتح

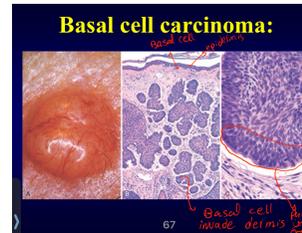


Squamous cell carcinoma

* common neoplasm
sun damage, localised

* Invasive

- Risk factors
- Immunosuppression (HPV)
 - sun exposure prolong
 - Radiation
 - old burns
 - tars, oils



Basal cell carcinoma

Small sun ^{تقرح} large nucleus ^{on epidermis invade to dermis}

Multiply ^{Wiz}

- PTCH1, TP53
- papules + pigment
- localise
- Gorlin syndrome
- Nevus

From melanocyte ← **Melanocytic neoplasm**

congenital **NEVUS**

Absence Atypical

- Benign
- BRAF, RAS (mutation)

- clinical :-

- ↳ sharp demarcated
- * ↳ elevated pigmented

- excision surgically

melanoma تتحول مع الوقت

- no change with time.

Histology:-

↳ symmetry

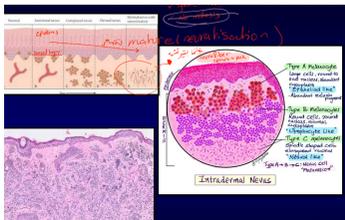
↳ maturation as move deep

As neurofiber pink

بتغير بمرور الوقت

stages:

- 1- Junctional [Between dermis-epidermis]
- 2- compound [Junction + dermis]
- 3- Intra dermal



Dysplastic Nevus

- Atypical.
- sporadic
- Familial → multiple → High life Risk

= with / without sun

* Microscopic diagnosed not Clinical

Histology:-

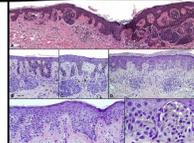
↳ Asymmetry

↳ Fusion Junctional

↳ superficial dermal fibrosis

* lymphocytic infiltration.

↳ Melanin incontinence



Melanoma

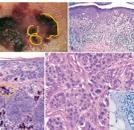
- malignant (Fatal)
- surgical cured (excision)

mutation	Early transition	late mutation transition
	BRAF RAS	P53 PTEM
		Vertical growth then led to Metastasis

bad prognosis

Features:

- ✓ Irregular Border
- ✓ pigmentation
- ✓ ↑* single cells
- ✓ ↑ thickness (called Breslow thick)
- ✓ Deep invasion → stage
- ✓ Large Atypical. nuclei
- ✓ cherry red nucleoli



Treat:

- Anti BRAF + KIT
- Immuno therapy T-cell (Immune check point inhibitor)

بيلانين مع تحلل

شوا الظلام قبل انها melanoma:

- ✓ Rapid enlarge
- ✓ Itch + pain
- ✓ New pigment