PATHOPHYSIOLOGY

Overview

What is low back pain?

- Pain below the costal margin and above the gluteal folds, with or without radiation to the lower extremity¹
- Acute vs. chronic low back is pain classified according to duration:
 - Acute: less than 3 months^{2,3}
 - Chronic: more than 3 months^{2,3}



1. Airaksinen O *et al. Eur Spine J* 2006; 15(Suppl 2):S192-300; 2. International Association for the Study of Pain. *Unrelieved Pain Is a Major Global Healthcare Problem.* Available at: <u>http://www.iasp-pain.org/AM/Template.cfm?Section=Press_Release&Template=/CM/ContentDisplay.cfm&ContentID=2908</u>. Accessed: July 22, 2013. 3. National Pain summit Initiative. *National Pain Strategy: Pain Management for All Australians*. Available at: http://www.iasp-pain.org/PainSummit/Australia_2010PainStrategy.pdf. Accessed: July 22, 2013.

The Pain Continuum

Time to resolution

Acute pain

nsult

Normal, time-limited response to 'noxious' experience (less than 3 months)

- Usually obvious tissue damage
- Serves a protective function
- Pain resolves upon healing

Chronic pain

Pain that has persisted beyond normal tissue healing time (usually more than 3 months)

- Usually has no protective function
- Degrades health and function

Acute pain may become chronic

Chapman CR, Stillman M. In: Kruger L (ed). *Pain and Touch*. Academic Press; New York, NY: 1996; Cole BE. *Hosp Physician* 2002; 38(6):23-30; International Association for the Study of Pain. *Unrelieved Pain Is a Major Global Healthcare Problem*. Available at: <u>http://www.iasp-pain.org/AM/Template.cfm?Section=Press</u> <u>Release&Template=/CM/ContentDisplay.cfm&ContentID=2908</u>. Accessed: July 24: 2013; National Pain Summit Initiative. *National Pain Strategy: Pain Management for All Australians*. Available at: <u>http://www.iasp-pain.org/PainSummit/Australia_2010PainStrategy.pdf</u>. Accessed: July 24, 2013; Turk DC, Okifuji A. In: Loeser D *et al* (eds.). *Bonica's Management of Pain*. 3rd ed. Lippincott Williams & Wilkins; Hagerstown, MD: 2001.

Natural History of Low Back Pain



Adapted from: Gunn CC et al. Spine 1980; 5(3):279-91.

Pathophysiology of Low Back Pain

Central sensitization/ dysfunctional pain May develop over time in some patients with chronic low back pain

Nociceptive pain Most patients with acute non-specific low back pain (85%) Chronic low back pain has been acknowledged to have multiple potential mechanisms. This is called "mixed pain."

Neuropathic pain Radiculopathy (7%)

Manusov EG. Prim Care 2012; 39(3):471-9; Neblett R et al. Pain 2013; 14(5):438-45; Vellucci R. Clin Drug Investig 2012; 32(Suppl 1):3-10; Woolf CJ, Salter MW. Science 2000; 288(5472):1765-9.

Low Back Pain

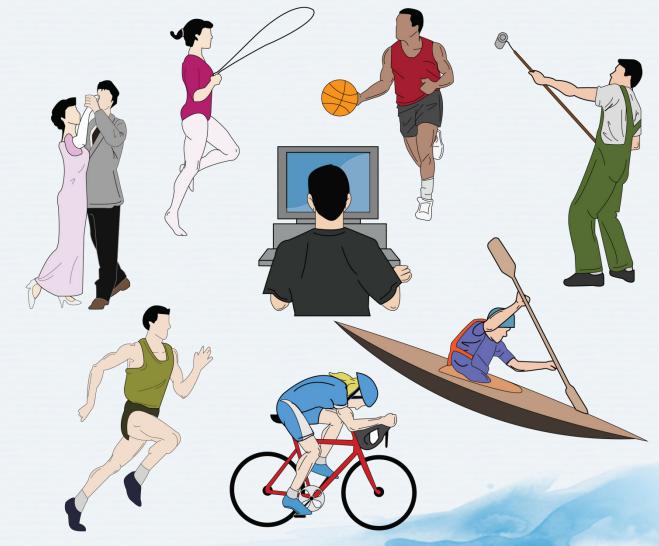
Simple classification of low back pain:

- Specific spinal pathology
- Radiculopathy
- Non-specific low back pain



Etiology

Causes of Low Back Pain: Repetitive Trauma (Overuse)



Common Causes of Low Back Pain

Mechanical (80-90%)

(e.g., disc degeneration, fractured vertebrae, instability, unknown cause [most cases])

Neurogenic (5-15%)

(e.g., herniated disc, spinal stenosis, osteophyte damage to nerve root)

Non-mechanical spinal conditions (1-2%) (e.g., neoplasm, infections, inflammatory arthritis, Paget's disease)

Referred visceral pain (1-2%)

(e.g., gastrointestinal disease, kidney disease, abdominal aortic aneurism)

Other (2-4%)

(e.g., fibromyalgia, somatoform disorder, "faking" pain)

Cohen S. BMJ 2008; 337:a2718.

Etiology of Low Back Pain with Nerve Root Involvement

Cause	% of cases
Herniated disc	4
Spinal stenosis (narrow canal)	3
Other	-1%
 Fractures/crushing due to osteoporosis 	
 Joint diseases (e.g., degenerative osteoarthritis, 	
rheumatoid arthritis, ankylosing spondylitis)	
 Congenital (e.g., scoliosis or kyphosis) 	
Spondylolisthesis	
Overuse syndromes	
 latrogenic (e.g., post-chemotherapy fibrosis) 	
 Infection (e.g., vertebral [epidural]) 	
 Tumor (primary or metastatic) 	

Etiopathogenic Theories of Nerve Root Pain

1. Anatomic

- Deficient root protection
- Blood/cerebrospinal fluid mixed nutrition
- Vulnerability of capillary barrier

2. Compressive/edema

- Ease of inducing vascular stasis
- Situation in nerve root canal
- Time of onset of compression
- Situation of dorsal root ganglion

3. Congestive/edema

- Microvascular injury due to venous congestion
- Decreased fibrinolytic activity

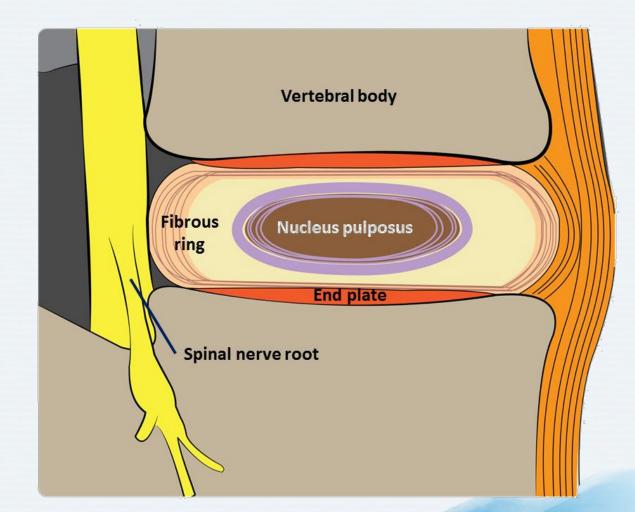
4. Immune

5. Inflammatory

6. Neural

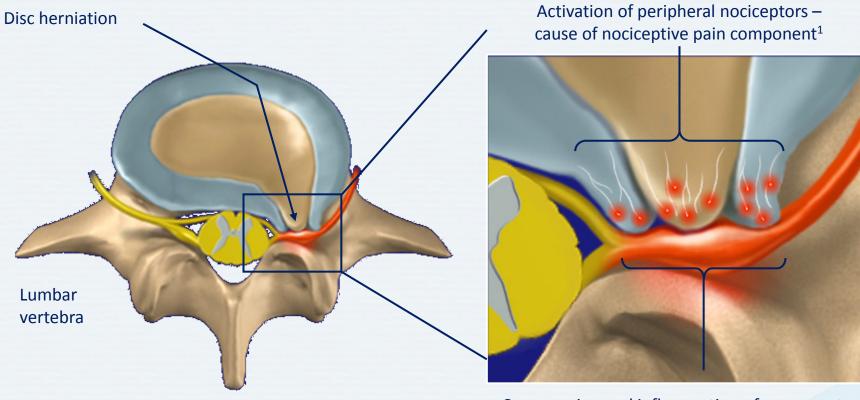
- Sensitivity of nociceptors
- Neurogenic inflammation
- Dorsal root ganglion disease
- Altered pool of endogenous neuropeptides
- Ectopic discharge from ischemic axons
- Centralization of pain

Relationship between Vertebral Body and Longitudinal Ligament*



*Sagittal section of vertebral body and intervertebral disc Shankar H *et al. Tech Reg Anesth Pain Manage* 2009; 13(2):67-75.

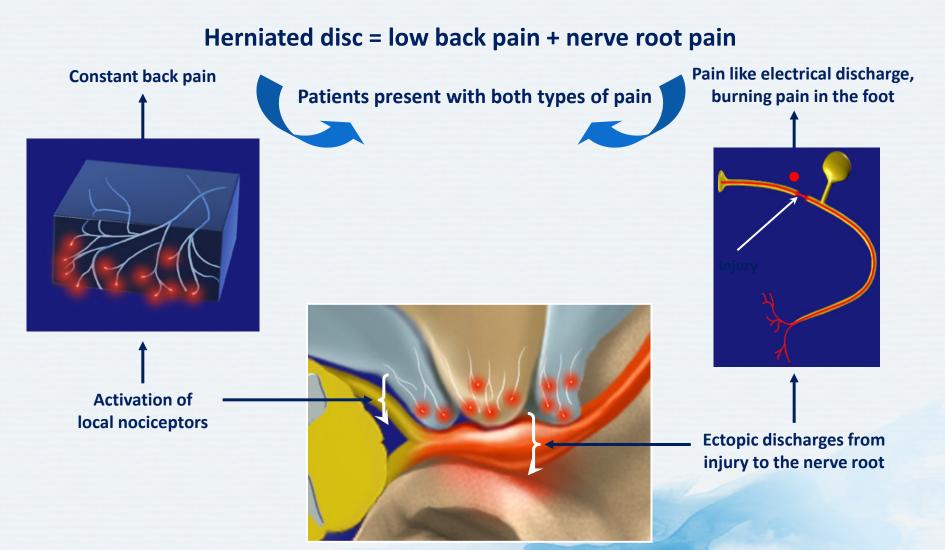
Example of Coexisting Pain: Herniated Disc Causing Low Back Pain and Lumbar Radicular Pain



Compression and inflammation of nerve root – cause of neuropathic pain component²

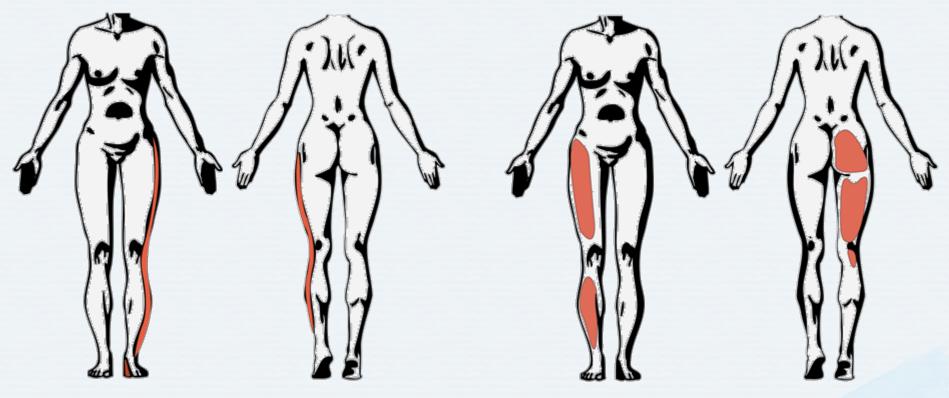
Brisby H. J Bone Joint Surg Am 2006; 88(Suppl 2):68-71.
 Freynhagen R, Baron R. Curr Pain Headache Rep 2009; 13(3):185-90.

Mixed Pain Example



Archival images, Pfizer, Inc.

Topographic Patterns of Pain Projection



Radiating Pain

Referred Pain

Vivian D. In: Lennard TA et al (eds). Pain Procedures in Clinical Practice. 3rd ed. Elsevier; New York, NY: 2011.

Potential Causes of Radiculopathy and Pseudoradiculopathy

Radiculopathy

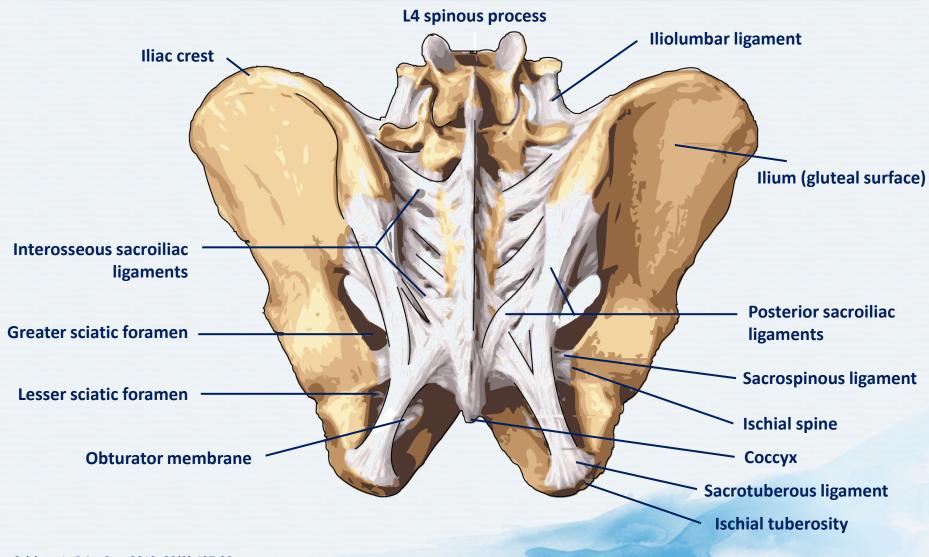
- Herniated disc with nerve compression
- Lumbar spinal stenosis
- Foraminal stenosis
- Diabetes mellitus
- Degenerative disc disease
- Residual adhesions from previous surgery
- Nerve root inflammation
- Spondylolisthesis

Pseudoradiculopathy

- Muscle contracture
- Facet pain
- Sacroiliac disease
- Osteochondrosis
- Trochanteric bursitis
- Local or systemic inflammation
- Infarction (can mimic root pain)

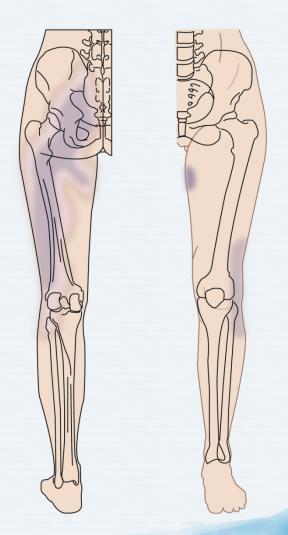
Freynhagen R, Baron R. Curr Pain Headache Rep 2009; 13(3):185-90.

Normal Anatomy of the Sacroiliac Joint



Salzberg L. Prim Care 2012; 39(3):487-98.

Typical Reference Pattern of Sacroiliac Pain



Soft Tissue Causes of Low Back Pain

Soft Tissue Condition	Clinical Features	Pain Pattern
Myofascial pain syndrome	Rope-like nodularity on physical examination	 Low back, buttocks, thighs (localized or regional)
Paraspinal muscle injury	 Muscle atrophy on MRI, ultrasound and CT 	Low back
Injury to quadratus Iumborum	 Decreased and painful lumbar flexion and rotation 	 Flank, low back, buttocks, lateral hip
Ischiatic bursitis	 Local tenderness at the ischial tuberosity 	• Buttocks
Cluneal nerve entrapment	 Resolution of pain with local nerve block 	Unilateral, iliac crest and buttocks
Sacroiliitis	 Inflammation of one or both sacroiliac joints 	 Pain in buttocks or low back and may extend to groin and one or both legs Often aggravated by prolonged standing or climbing stairs

CT = computed tomography; MRI = magnetic resonance imaging Borg-Stein J, Wilkins A. *Curr Pain Headache Rep* 2006; 10(50:339-44.

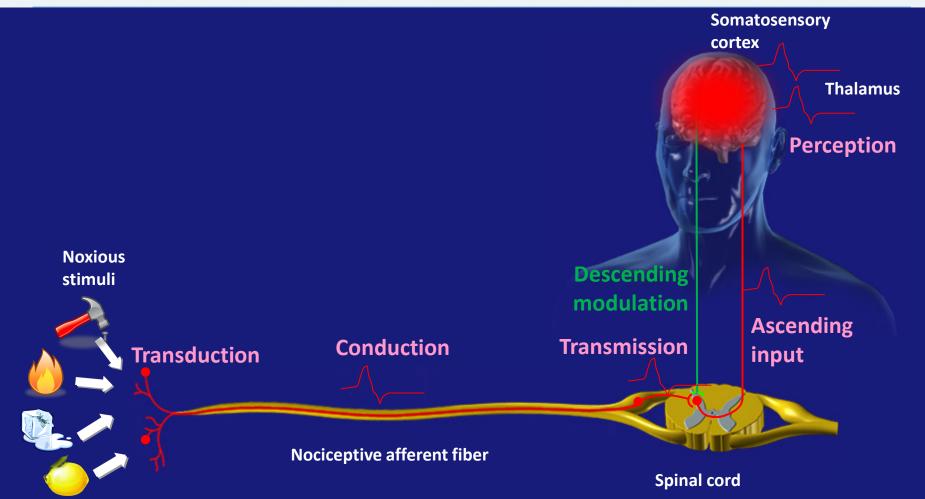
Soft Tissue Causes of Low Back Pain (cont'd)

Soft Tissue Condition	Clinical Features	Pain Pattern
Painful syndrome of hip abductor	 Tender gluteal muscles lateral to posterior-superior iliac spine Hip abductor muscle weakness Trendlenburg sign 	Buttocks, lateral aspect of thigh
Bursitis of the psoas	 Most painful movement is passive adduction in flexion Appearance on musculoskeletal ultrasound is consistent with inflammation 	• Groin, anterior thigh, knee, leg
Bursitis of the trochanter	 Positive "jump" sign secondary to thumb pressure over most prominent ridge of greater trochanter 	 Pseudoradiculopathy: pain does not extend distal to proximal tibia (insertion of iliotibial tract at Gerdy's tubercle)
Gluteal bursitis	 Pain on: Passive external rotation and passive abduction Passive abduction and either resisted external rotation or resisted abduction 	 Gluteal and trochanteric region, sometimes spreading to outer or posterior thigh and down to calf and lateral malleolus

Borg-Stein J, Wilkins A. Curr Pain Headache Rep 2006; 10(50:339-44.

Pathophysiology

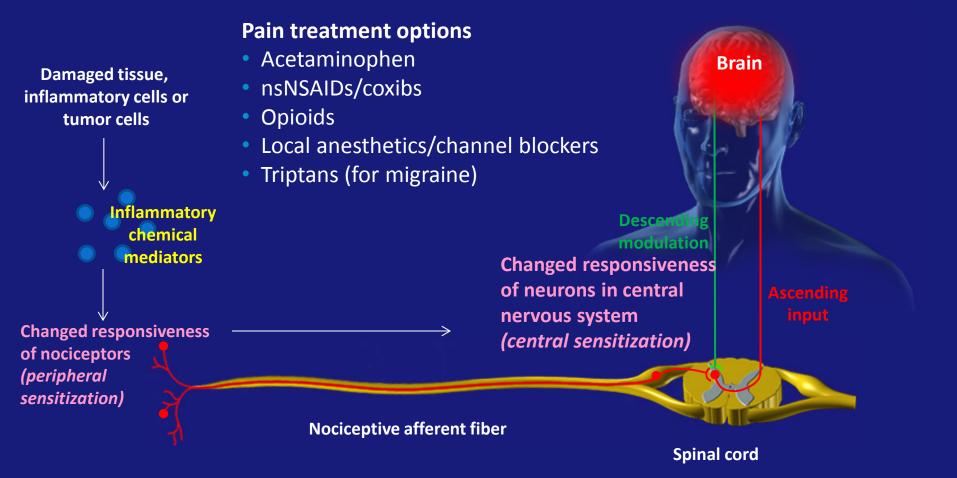
Nociception: Neural Process of Encoding Noxious Stimuli



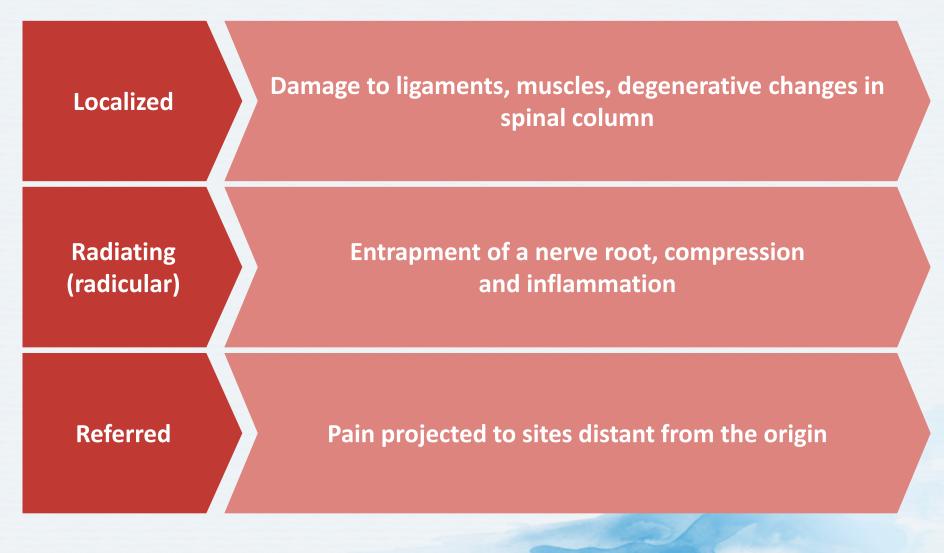
Consequences of encoding may be autonomic (e.g., elevated blood pressure) or behavioral (motor withdrawal reflex or more complex nocifensive behavior). Pain perception is not necessarily implied.

Scholz J, Woolf CJ. Nat Neurosci 2002; 5(Suppl):1062-7.

Treatment of Inflammatory Pain



Pain Types Related to Spinal Disorders

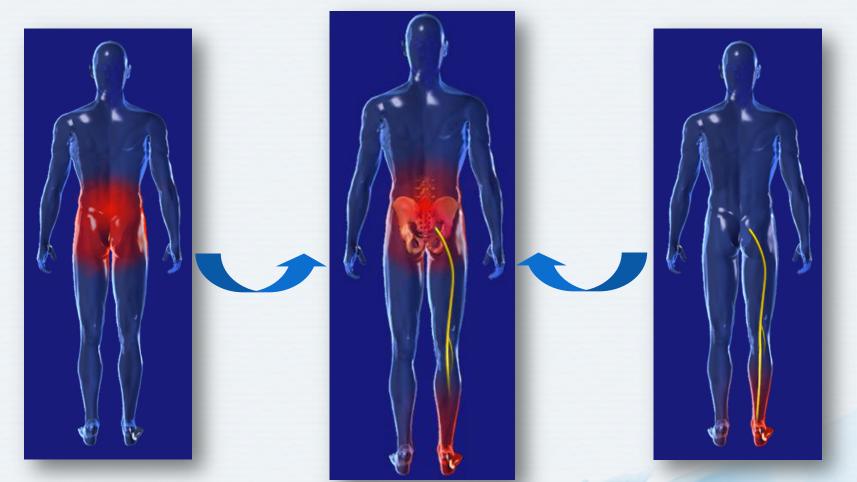


Low Back Pain Syndromes and Clinical Entities

Neuropathic (Nerve root)	Nociceptive (Musculoskeletal)
Lumbar nerve root syndrome	Internal disc tear
Herniated lumbar disc	Lumbar facet syndrome
Lumbar spinal stenosis	Dysfunction of the sacroiliac joint

Malik K, Benzon HT. In: Benzon HT et al (eds). Raj's Practical Management of Pain. 4th ed. Mosby; Maryland Heights, MO: 2008.

Nociceptive and Neuropathic Components May Be Present in Low Back Pain



Nociceptive Component

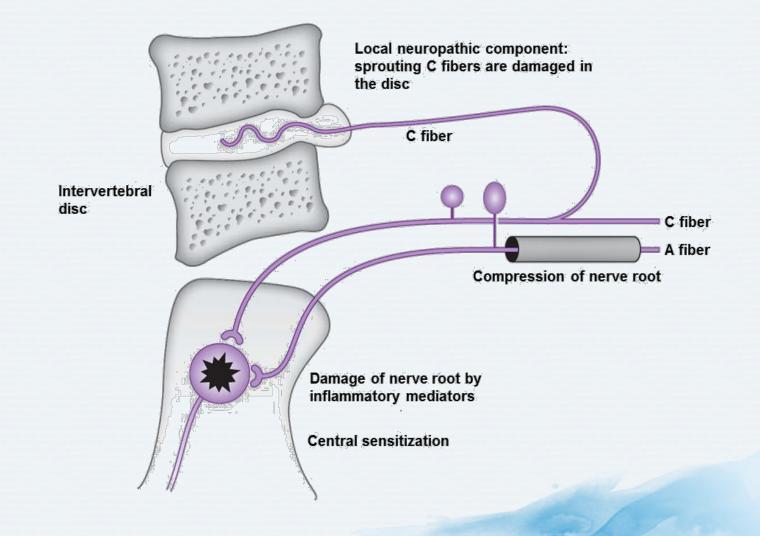
Neuropathic Component

Freynhagen R, Baron R. Curr Pain Headache Rep 2009; 13(3):185-90.

Neuropathic Component of Low Back Pain

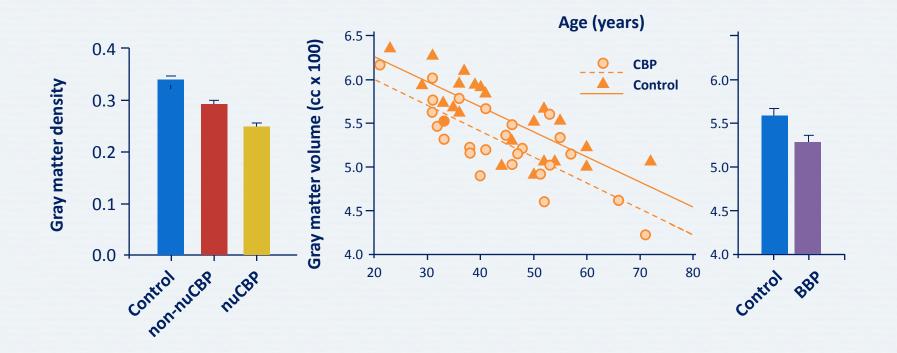
- Neuropathic component of low back pain may be caused by:
 - Mechanical compression of nerve root (mechanical neuropathic nerve root pain)
 - Damage to sprouting C-fibers within the degenerated disc (*localized neuropathic pain*)
 - Action of inflammatory mediators released from the degenerated disc (*inflammatory neuropathic nerve root pain*), even without mechanical compression

Pathophysiological Mechanisms in Neuropathic Low Back Pain



Freynhagen R, Baron R. Curr Pain Headache Rep 2009; 13(3):185-90.

Grey Matter Density and Volume Is Reduced in Chronic Back Pain



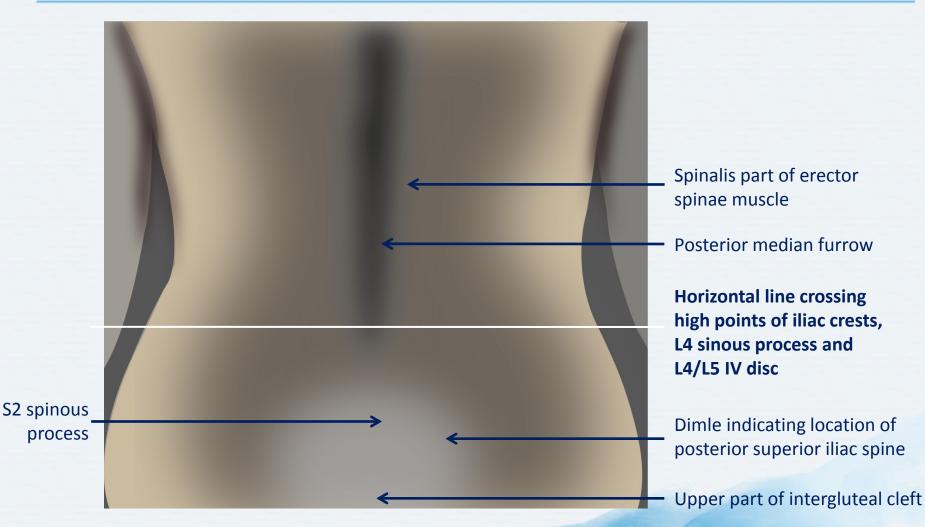
• 11% reduction in volume in patients with chronic back pain vs. controls

• Reduction may be reversible with proper treatment

Apkarian AV et al. J Neurosci 2004; 24(46):10410-5.

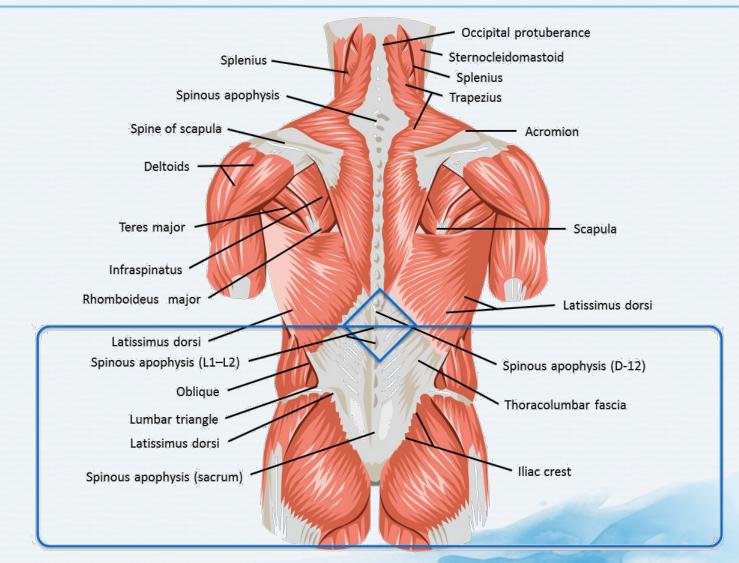
ANATOMY, IMAGING, AND SIMPLIFIED BIOMECHANICS

Superficial Anatomy of the Lumbar Spine



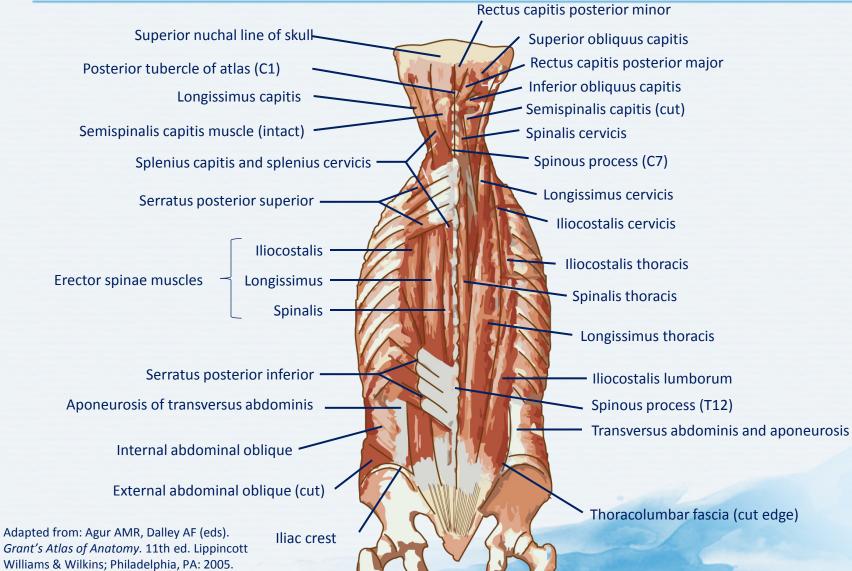
Adapted from: Agur AMR, Dalley AF (eds). Grant's Atlas of Anatomy. 11th ed. Lippincott Williams & Wilkins; Philadelphia, PA: 2005.

Superficial Muscles of the Back



Adapted from: Agur AMR, Dalley AF (eds). Grant's Atlas of Anatomy. 11th ed. Lippincott Williams & Wilkins; Philadelphia, PA: 2005.

Intermediate Layers of the Muscles of the Back

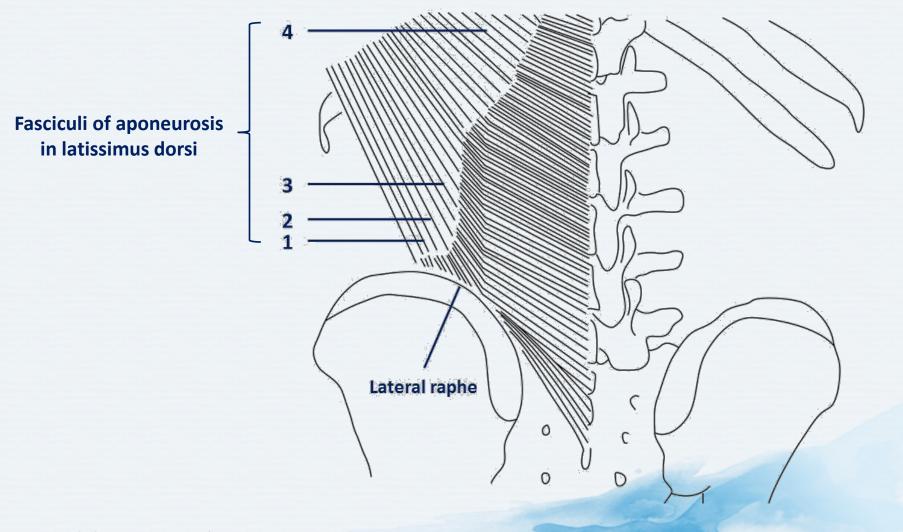


Anatomy of the Lumbar Muscles and their Fascias

В Α С D Ε F ٥ 0

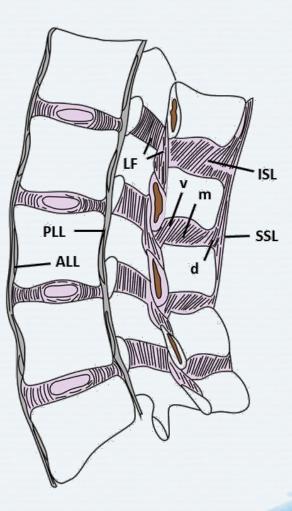
Bogduk N (ed). Clinical Anatomy of the Lumbar Spine and Sacrum. 4th ed. Elsevier; London, UK: 2005.

Superficial Thoracolumbar Fascia



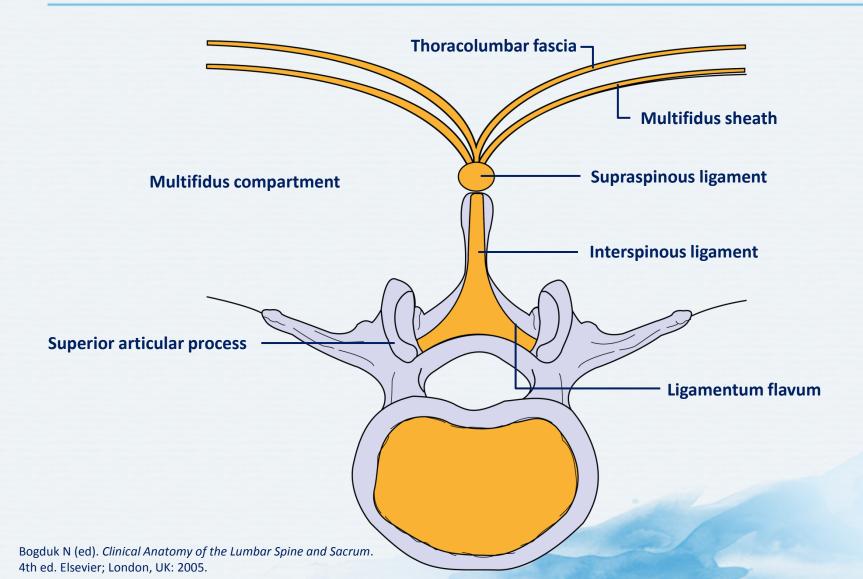
Bogduk N (ed). Clinical Anatomy of the Lumbar Spine and Sacrum. 4th ed. Elsevier; London, UK: 2005.

Sagittal View of Ligaments of the Lumbar Spine

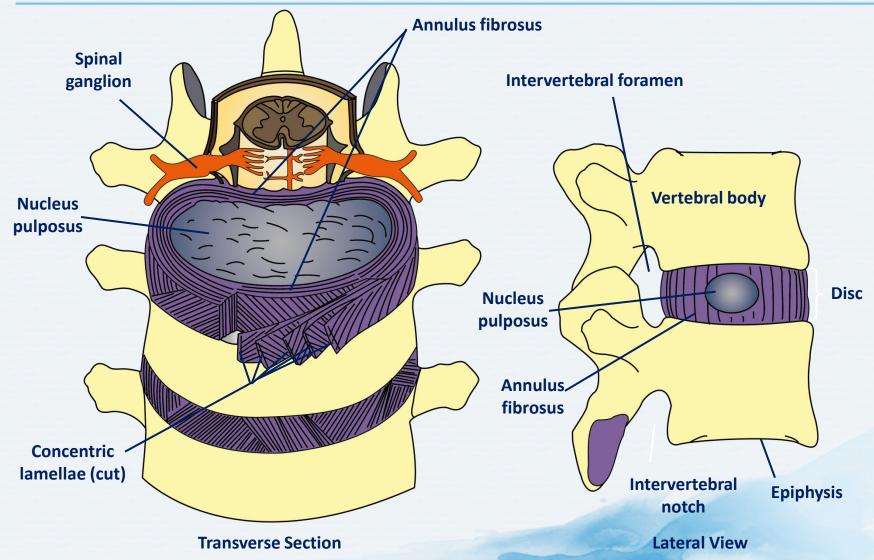


ALL = anterior longitudinal ligament; ISL = interspinous ligament; LF = ligamenta flava (yellow ligament); PLL = posterior longitudinal ligament; SSL = supraspinous ligament; v, m, and d are fasciculi of the ISL Bogduk N (ed). *Clinical Anatomy of the Lumbar Spine and Sacrum*. 4th ed. Elsevier; London, UK: 2005.

Complex Muscles, Ligaments, and Nerve Structures Associated with Low Back Pain

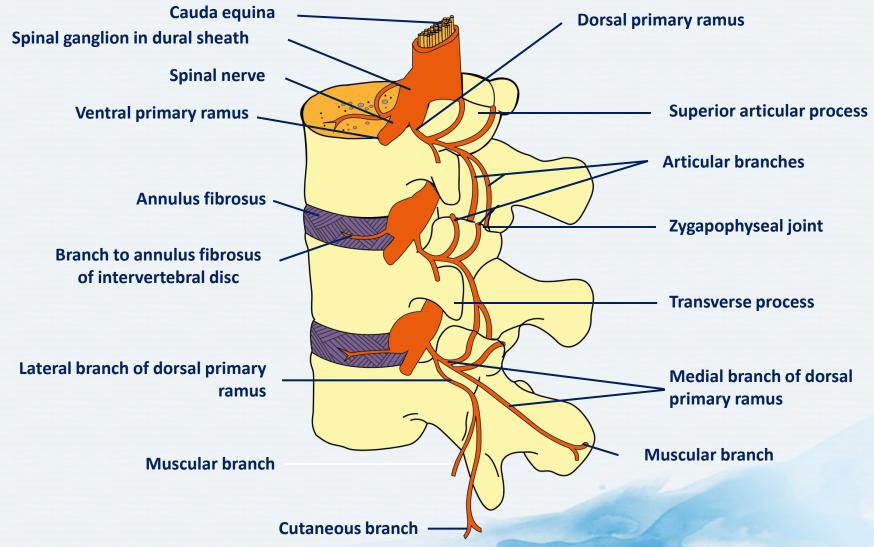


Anatomy of Intervertebral Disc and Ligaments: Transverse Section and Lateral View



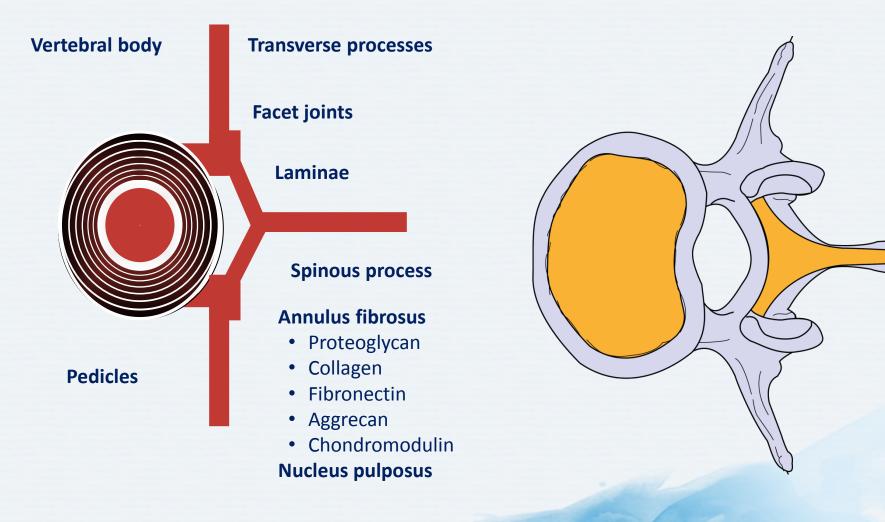
Bogduk N (ed). Clinical Anatomy of the Lumbar Spine and Sacrum. 4th ed. Elsevier; London, UK: 2005.

Anatomy of Intervertebral Disc and Ligaments: Posterolateral View



Bogduk N (ed). Clinical Anatomy of the Lumbar Spine and Sacrum. 4th ed. Elsevier; London, UK: 2005.

Simplified Axial Anatomy of the Spinal Column



Images courtesy of Dr. Ihab Gharzeddine

Bogduk N (ed). Clinical Anatomy of the Lumbar Spine and Sacrum. 4th ed. Elsevier; London, UK: 2005.

Intervertebral Disc Components: Structure and Location

	Annulus fibrosus	Nucleus pulposus	Vertebral end plates
Structure and location	 Fibers arranged in 10 to 12 layers (lamellae) forming concentric rings encircling nucleus pulposus Lamellae are thicker anteriorly and laterally and thinner posteriorly Fibers lie parallel within each lamella but are oriented at 65–70 degrees from vertical, successively in opposite directions between layers 	 Oval-shaped mass in central or posterior-central disc Enclosed by collagen fibers from inner layers of annulus fibrosus 	 Boundary between cancellous core of vertebral body and intervertebral disc Vertebral end plates extend centrally from an apophyseal ring and completely enclose nucleus pulposis from above and below

Chiarello CM. In: Cameron M (ed). *Physical Medicine Assessment and Intervention: An Evidence Based Approach.* Elsevier; Philadelphia, PA: 2007.

Intervertebral Disc Components: Composition and Properties

	Annulus fibrosus	Nucleus pulposus	Vertebral End plates
Composition	 60–70% water Dry components are 50–60% collagen (mostly type I) and 20% proteoglycans to bind water Chondrocytes near nucleus and fibroblasts near annular periphery synthesize collagen and proteoglycans 	 Hydrated, gelatinous, semifluid mass (70–90% water) Dry components are 65% proteoglycans and 15–20% collagen (mostly type II) Chondrocytes near vertebral end plates synthesize proteoglycans and collagen 	 Primarily hyaline cartilage in area closest to vertebral body and fibrocartilage near nucleus pulposis Also contain proteoglycans, collagen fibers and chondrocytes, with more water and proteoglycans and less collagen centrally
Properties	 Resists tensile loads Half the lamellae resist torsional loads in each direction 	 Deforms under pressure but cannot be compressed. 	 Strongest and stiffest posterolaterally and weakest centrally

Chiarello CM. In: Cameron M (ed). *Physical Medicine Assessment and Intervention: An Evidence Based Approach*. Elsevier; Philadelphia, PA: 2007.

Intervertebral Disc Components: Function

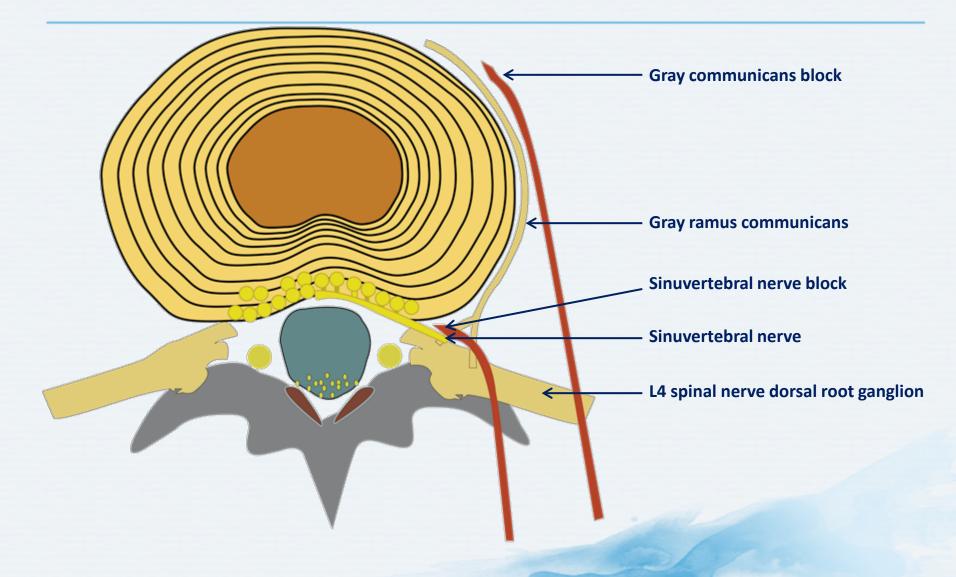
	Annulus fibrosus	Nucleus pulposus	Vertebral end plates
Function	 Principal load-bearing component of intervertebral disc 	 Reallocates applied loads in all directions to annulus fibrosus and vertebral end plates 	 Anchor intervertebral disc to vertebral bodies Prevent extrusion of nucleus pulposus into vertebral body Distribute and transfer load to vertebral body Site for diffusion of nutrients to intervertebral disc

Intervertebral Disc Components: Degenerative Changes

	Annulus fibrosus	Nucleus pulposus	Vertebral end plates
Degenerative changes	 Border between nucleus pulposus and annulus fibrosus becomes difficult to differentiate 3 types of tears occur: Peripheral: isolated to outer layers, parallel and adjacent to vertebral end plates Circumferential: rupture between lamellae Radial: continuous with clefts that radiate from nucleus pulposus 	 As water and proteoglycan levels decrease, nucleus pulposus becomes dryer, more fibrotic and less distinct from annulus fibrosus Disc becomes weaker as nucleus pulposus becomes less able to distribute loads Horizontal clefts develop between vertebral end plates and disc center 	 Thinning, fissures, horizontal cleft formation and fractures increase with age Ossification and local calcification reduce disc nutrition

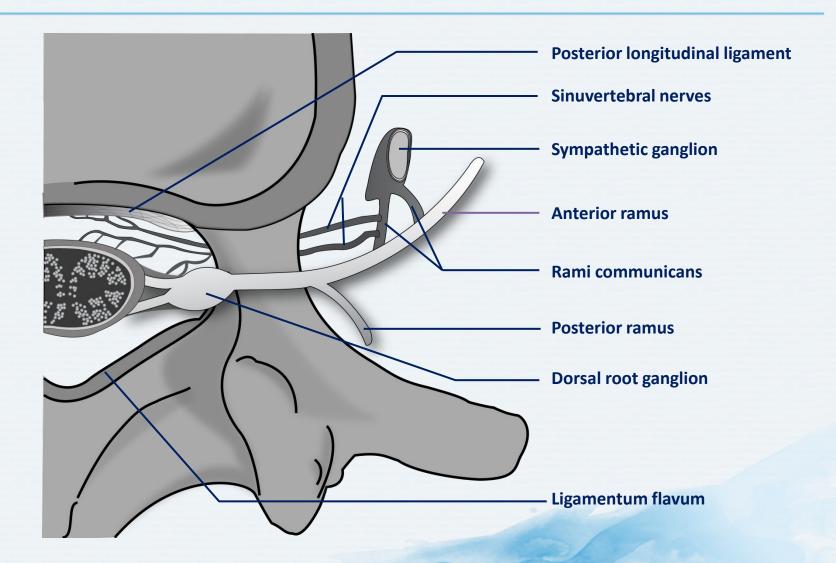
Chiarello CM. In: Cameron M (ed). *Physical Medicine Assessment and Intervention: An Evidence Based Approach.* Elsevier; Philadelphia, PA: 2007.

Intervertebral Disc Innervation

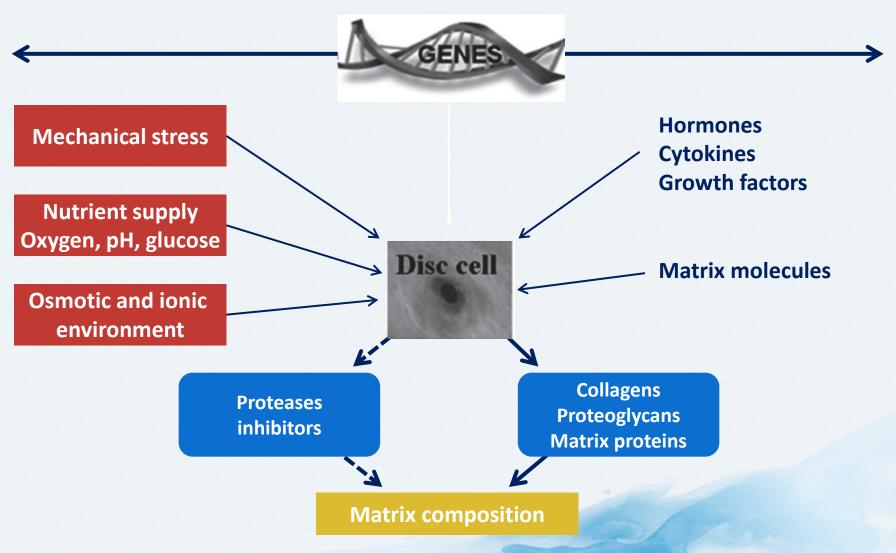


Vivian D. In: Lennard TA et al (eds). Pain Procedures in Clinical Practice. 3rd ed. Elsevier; Philadelphia, PA: 2011.

Sinuvertebral Nerves

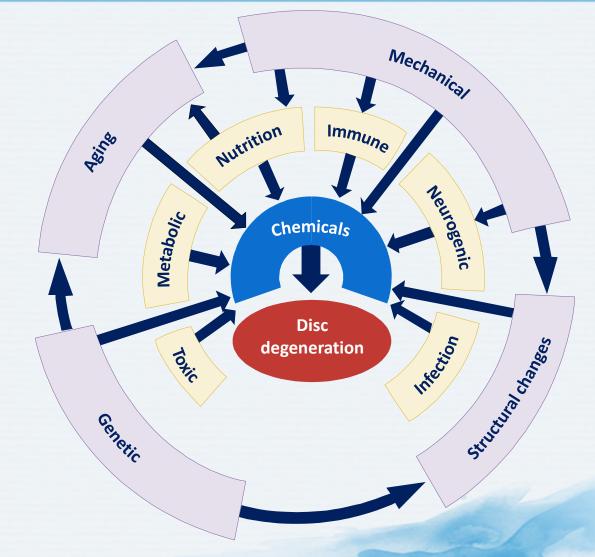


Cellular Activity of the Disc Influence of Environmental Factors as Potent Modifiers



Hughes SP et al. J Bone Joint Surg Br 2012; 94(10):1298-304.

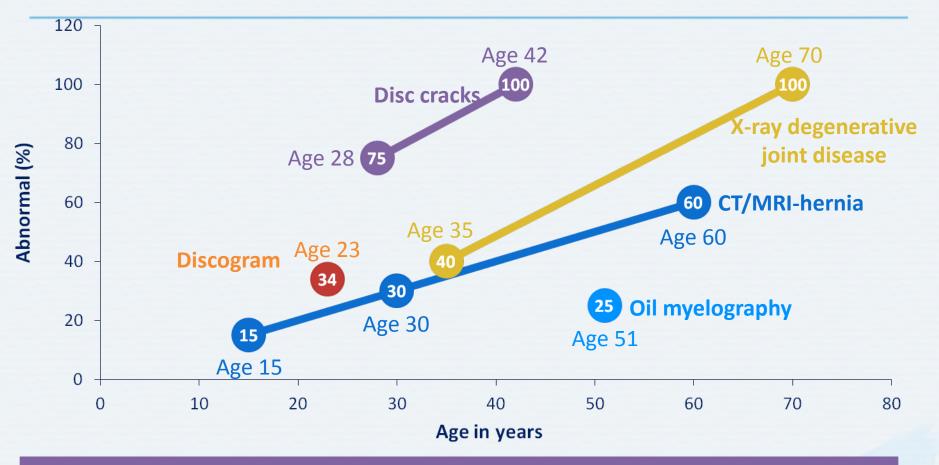
Multifactorial Pathophysiology of Disc Degeneration Model: Initiators, Promoters or Both



Hadjipavlou AG et al. J Bone Joint Surg Br 2008; 90(10):1261-70.

NORMAL IMAGING OF THE LUMBAR SPINE

Disc Findings in Normal Subjects



Find the patient's *age in years line* to determine the chance of a finding being present before symptoms begin.

CT = computed tomography; MRI = magnetic resonance imaging Bigos SJ, Davis GE. *J Orthop Sports Phys Ther* 1996; 24(4):192-207.

Plain-Film Radiology of a Normal Spinal Column with Physiological Lordosis

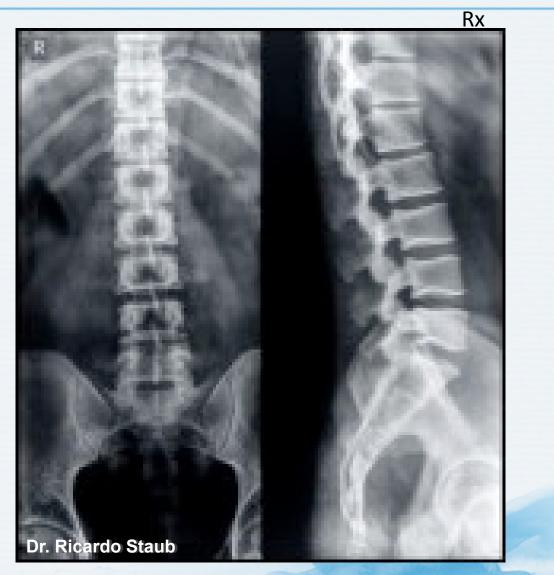


Image courtesy of Dr. Ricardo Staub

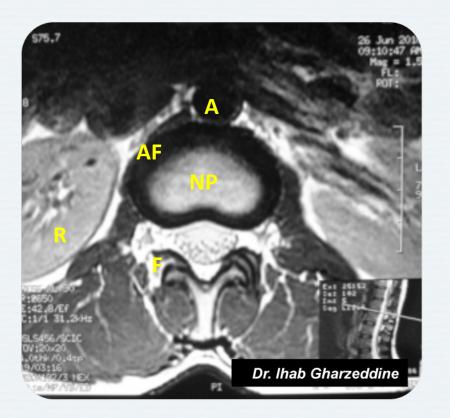
Magnetic Resonance Imaging of Sagittal Slices in T1 and T2

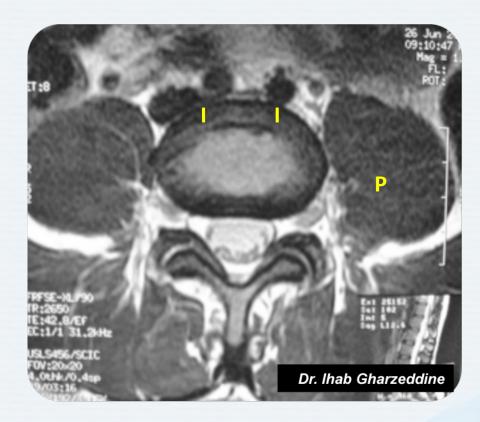
Dr. Ihab Gharzeddine Dr. Ihab Gharzeddine

T1 and T2 refer to repetition period during imaging.

Images courtesy of Dr. Ihab Gharzeddine FONAR. *MRI Glossary*. Available at: http://fonar.com/glossary.htm. Accessed: October 15, 2013.

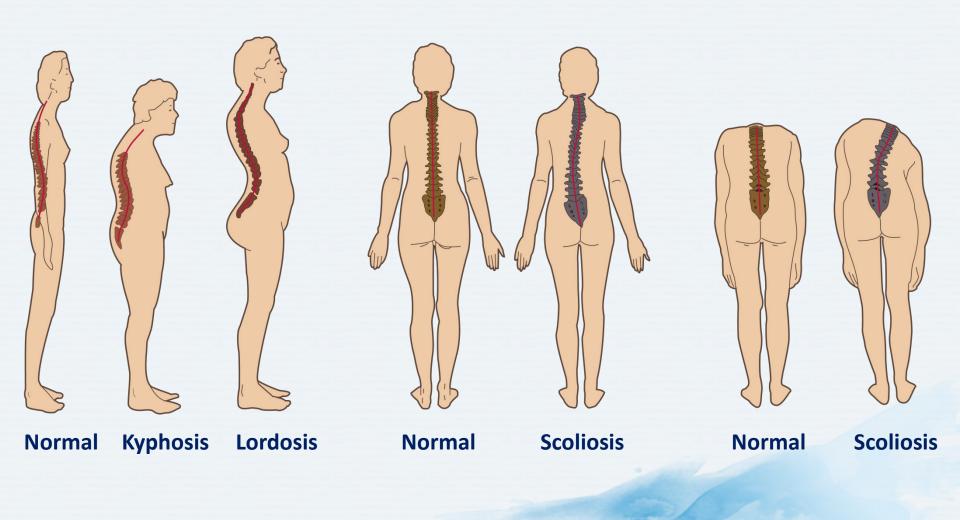
Axial Slices



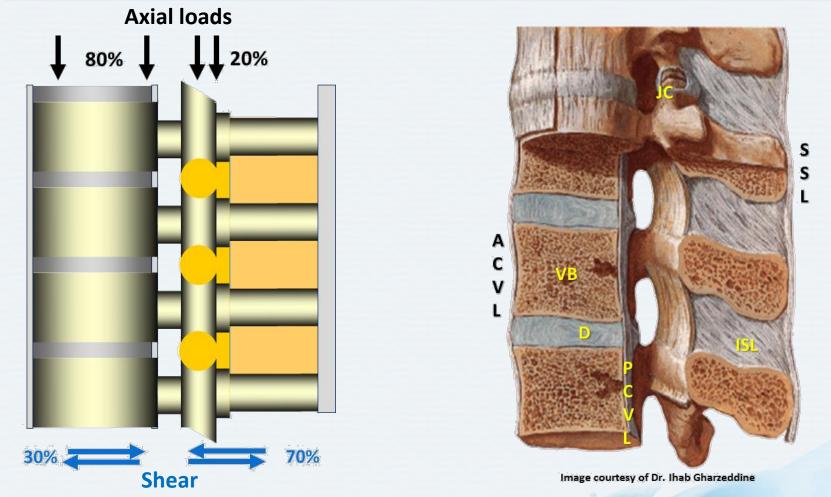


A = aorta; AF = annulus fibrosus; F = joint facet; I = bifurcation of the iliac artery; NP = nucleus pulposus; R = right kidney Images courtesy of Dr. Ihab Gharzeddine

Normal and Pathological Variants of the Vertebral Axis



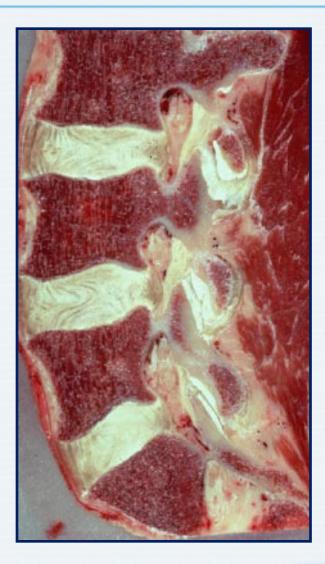
Simplified Spinal Anatomy in Sagittal View

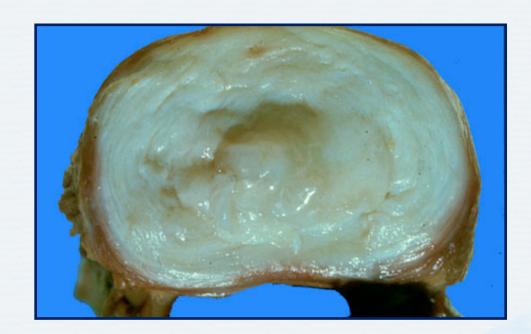


ACVL = anterior common vertebral ligament; D = intervertebral discs; ISL = interosseous sacroiliac ligament; JC = joint capsule; LF = yellow ligament; LIE = interspinous ligaments; LCVP = posterior common vertebral ligament; SSL = supraspinous ligaments; VB = vertebral bodies

Images courtesy of Dr. Ihab Gharzeddine

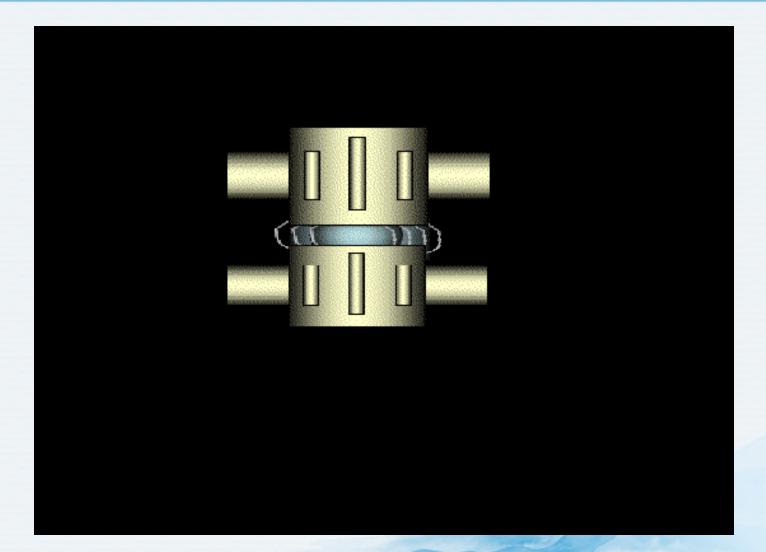
Pathological Anatomy of the Spine



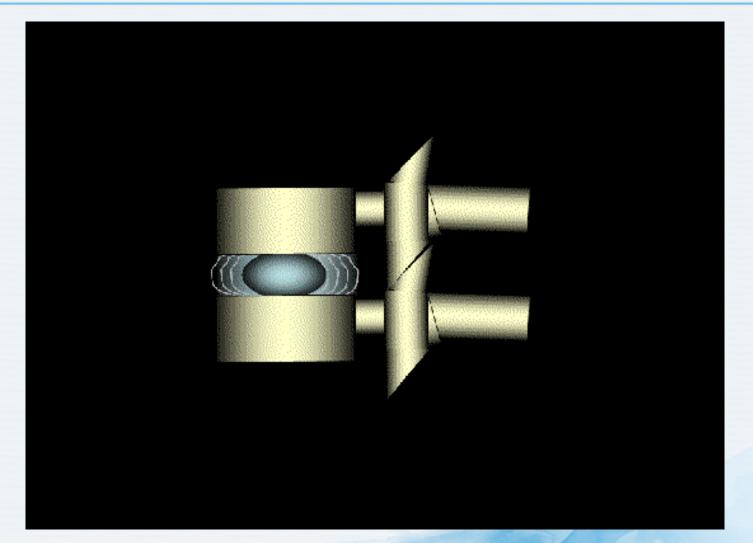


Images courtesy of Dr. Ihab Gharzeddine

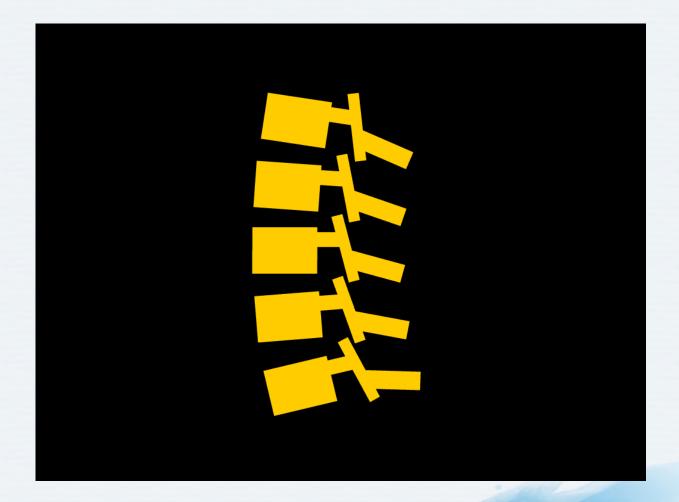
Vertebral Unit under Pressure or Compression



Vertebral Unit during Flexion and Extension

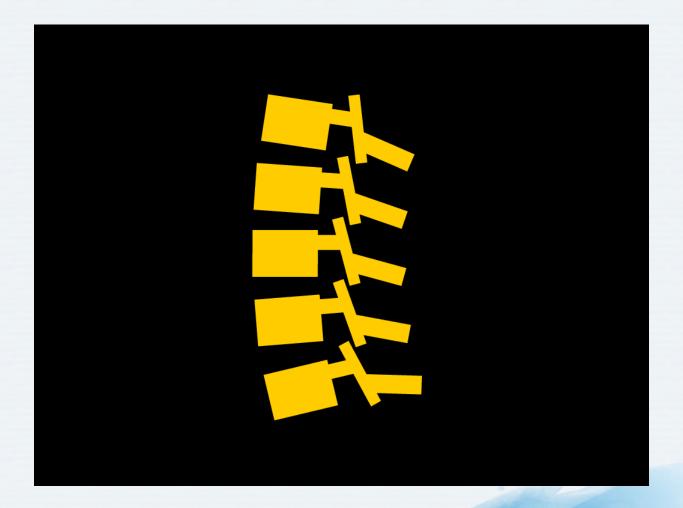


Combined Rotation and Translation in Normal Flexion-Extension in the Lumbar Spine

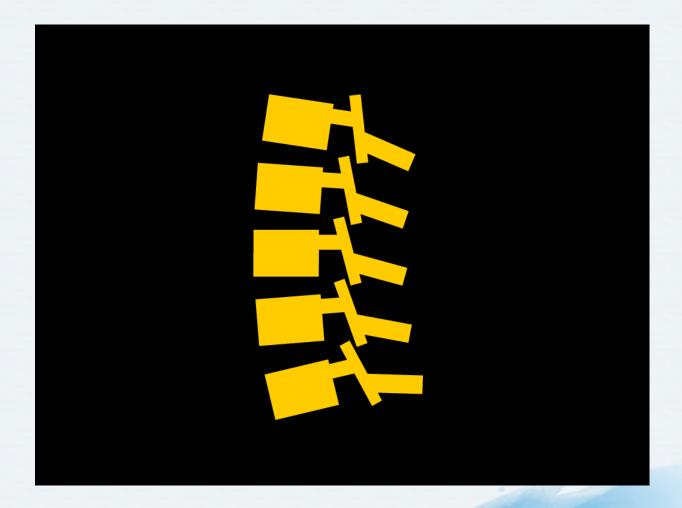


Animation courtesy of Dr. Ihab Gharzeddine

Theoretical Isolation of Rotational Movement during Flexion-Extension in the Lumbar Spine

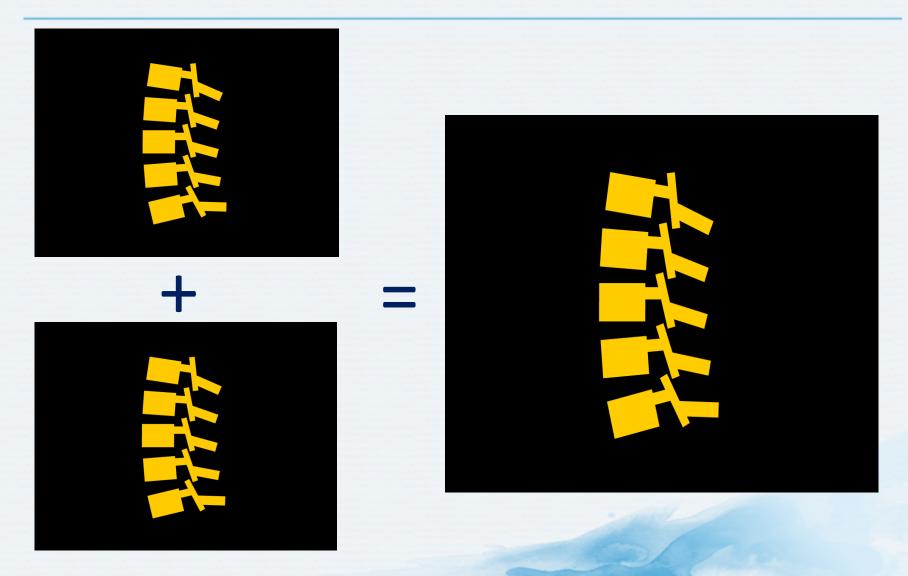


Theoretical Isolation of Translational Movement during Flexion-Extension in the Lumbar Spine



Animation courtesy of Dr. Ihab Gharzeddine

Flexion-Extension of the Spinal Column



Images and animation courtesy of Dr. Ihab Gharzeddine

PROTRUSION, EXTRUSION AND SEQUESTRATION

Protrusion of the Spinal Column

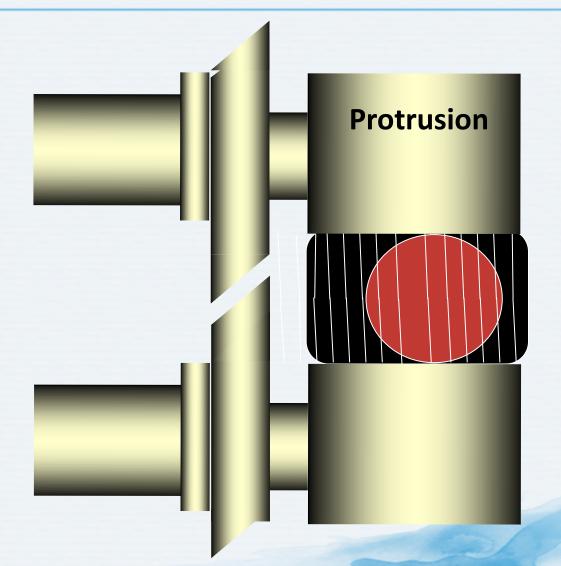


Image courtesy of Dr. Ihab Gharzeddine

Sagittal and Axial Views of Central Protrusion





Extrusion of the Spinal Column

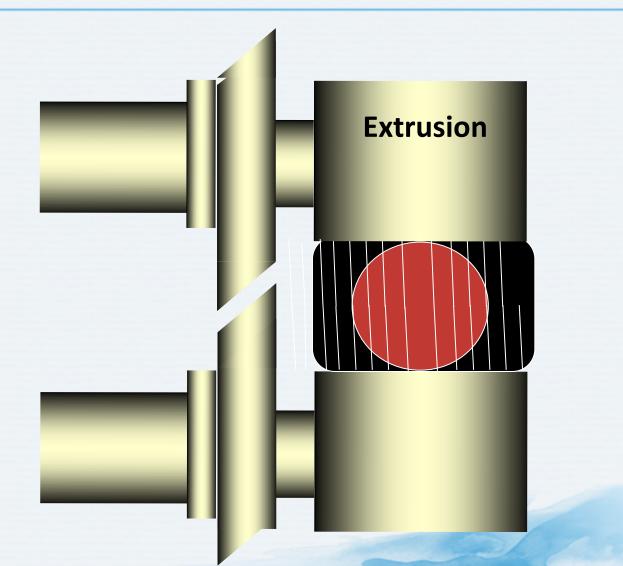


Image courtesy of Dr. Ihab Gharzeddine

Disc Extrusion





Disc Sequestration

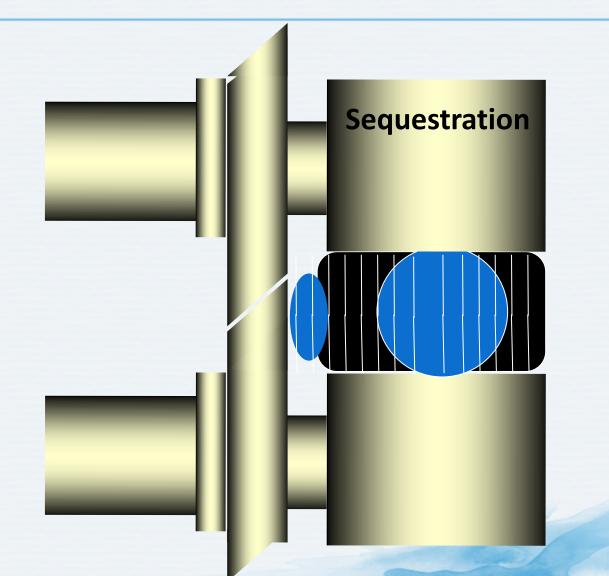
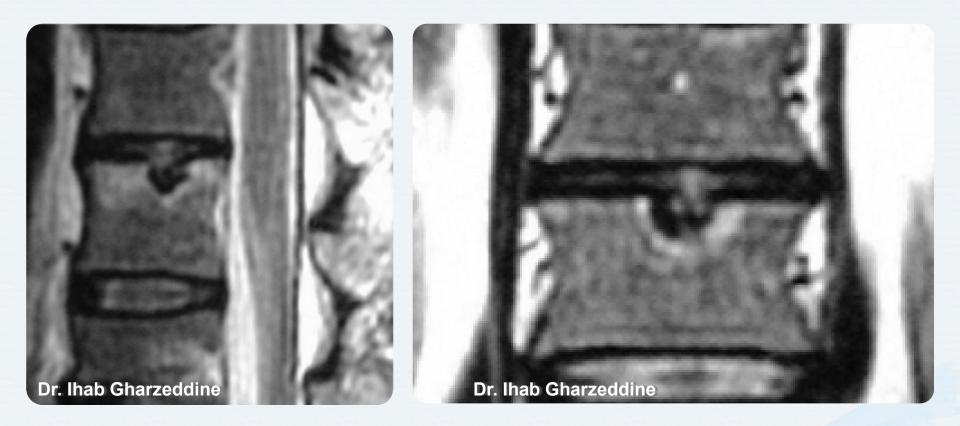


Image courtesy of Dr. Ihab Gharzeddine

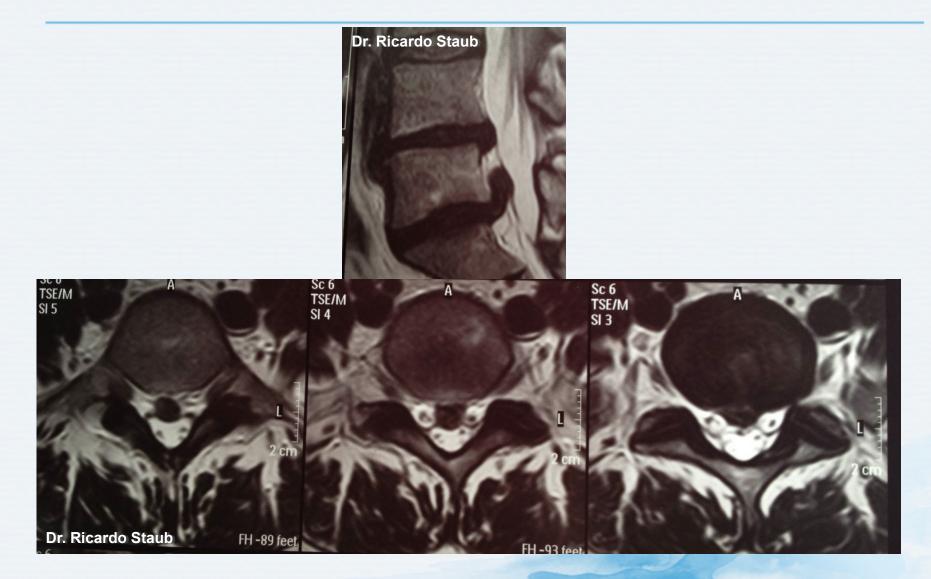
Disc Sequestration



Schmorl Nodules



What is this?



DEGENERATIVE DISC DISEASE ON MULTIPLE LEVELS

Vertebra-Disc-Vertebra Functional Unit

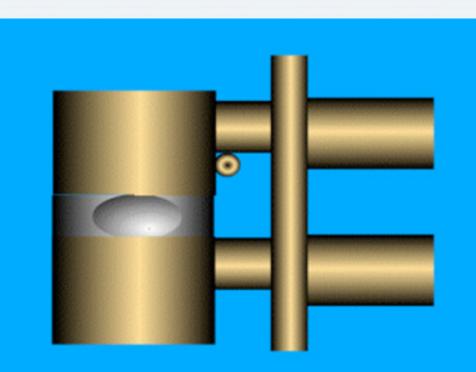
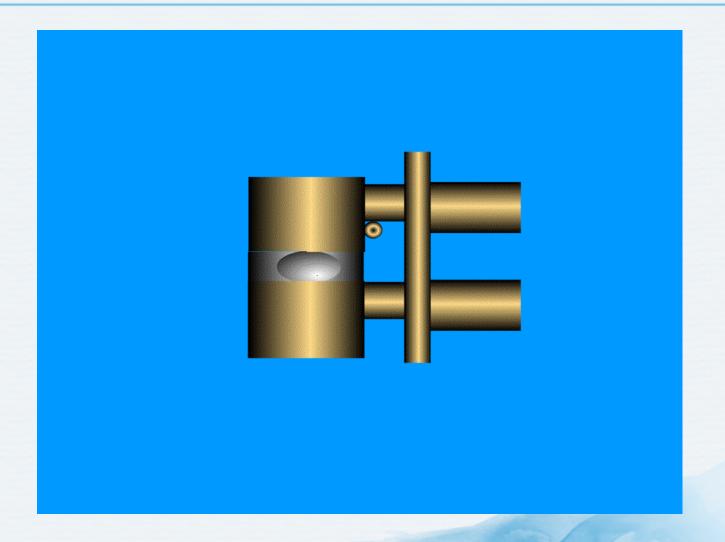


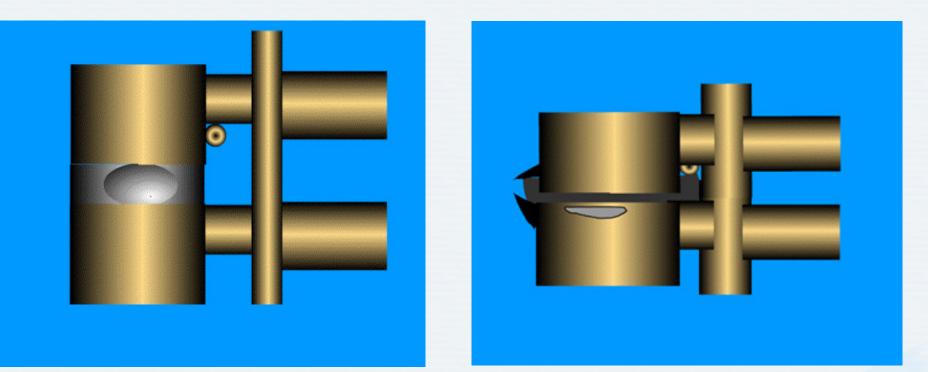
Image courtesy of Dr. Ihab Gharzeddine

History of the Vertebral Column



Animation courtesy of Dr. Ihab Gharzeddine

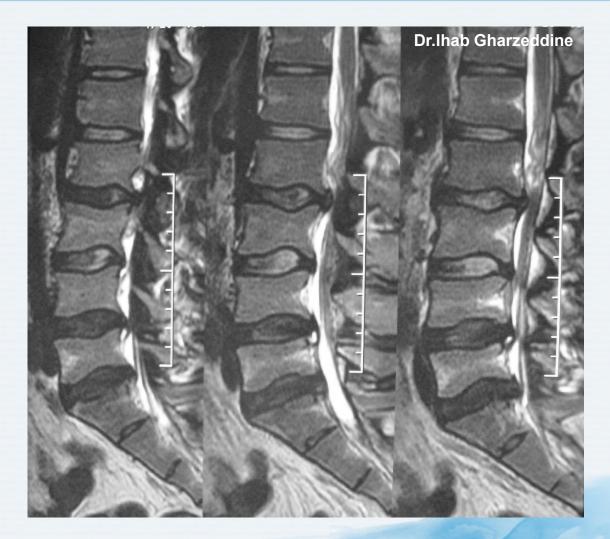
Normal vs. Degenerated Functional Unit of the Vertebral Column



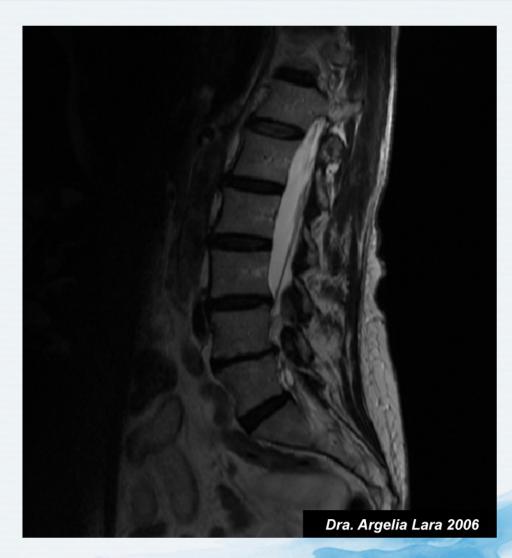
Images courtesy of Dr. Ihab Gharzeddine

PATHOLOGICAL DISC DEGENERATION

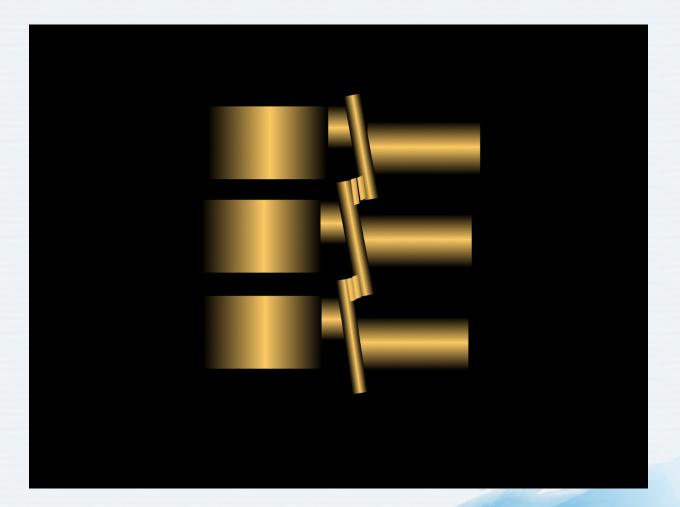
Disc Degeneration on Multiple Levels



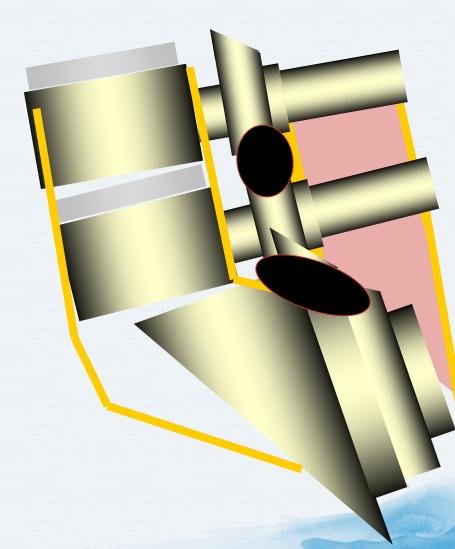
Narrow Lumbar Canal (Lumbosacral Stenosis)



Spondylolysis and Spondylolisthesis



Applied Anatomy and Biomechanics: Situation of Anatomical Structures in Listhesis



Degenerative Disc Disease with Spondylolisthesis





Disc Herniation, Narrow Canal and Spondylolisthesis



Image courtesy of Dr. Argelia Lara

Degenerative Scoliosis



De Novo Scoliosis



Scoliosis and Scoliotic Posture

Idiopathic Scoliosis



Scoliotic Posture



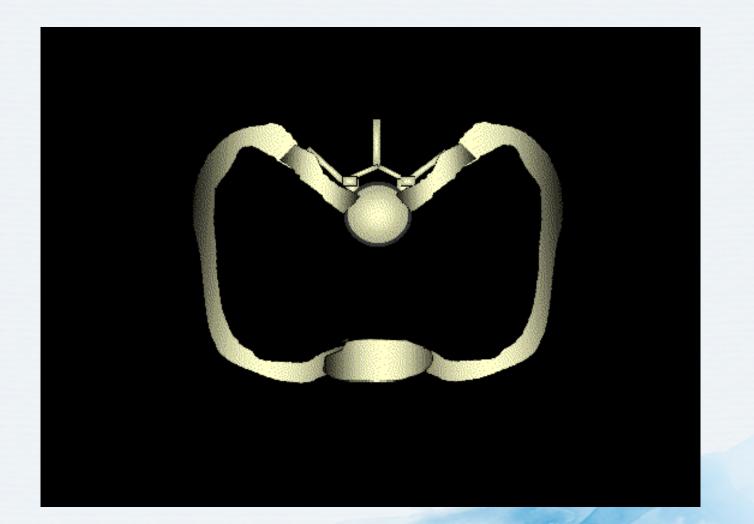
Images courtesy of Dr. Ihab Gharzeddine

Scoliosis



Image courtesy of Dr. Ihab Gharzeddine

Rotation of Thoracic Vertebrae



Asymmetrical Rib Cage in Scoliosis

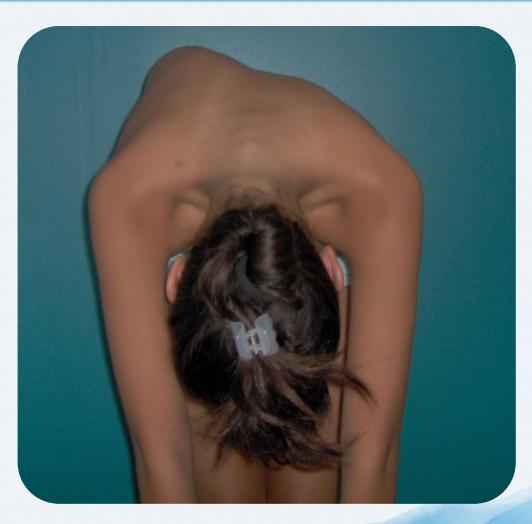
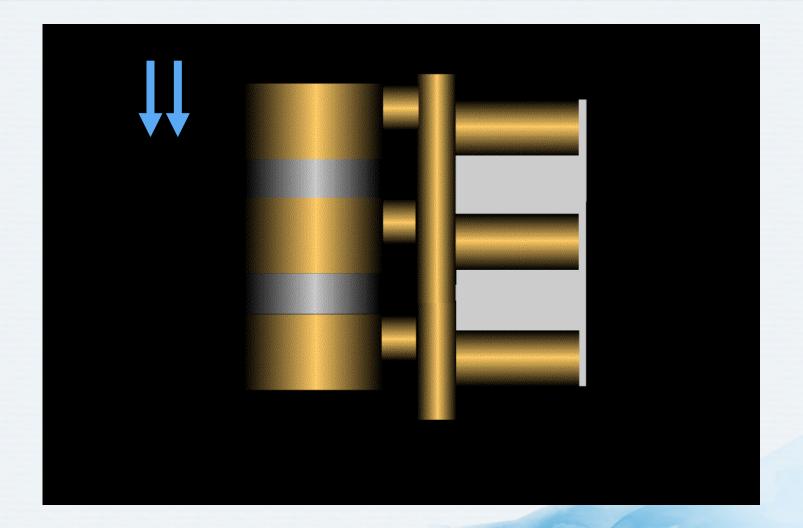


Image courtesy of Dr. Ihab Gharzeddine

FRACTURES

Compression Fractures



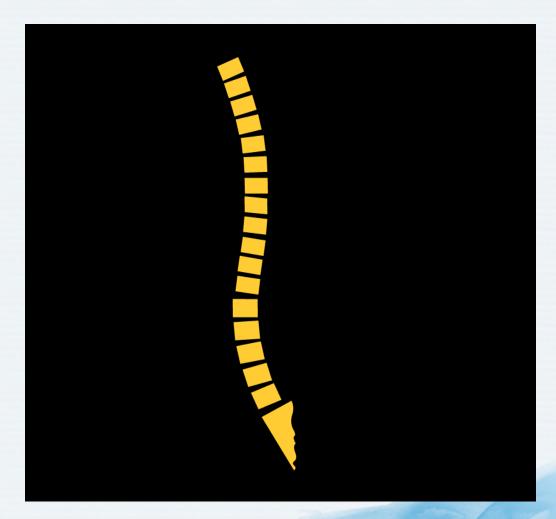
Animation courtesy of Dr. Ihab Gharzeddine

Vertebral Fracture



Image courtesy of Dr. Ihab Gharzeddine

Domino Effect in Vertebral Fractures



Animation courtesy of Dr. Ihab Gharzeddine

Summary

Pathophysiology of Low Back Pain: Summary

- Low back pain is most frequently attributable to a mechanical cause brought on by overuse or repetitive trauma
 - Other potential causes include neurogenic causes, referred visceral pain or non-mechanical spinal conditions
- Low back pain may be classified according to duration as acute, subacute or chronic
- Low back pain may also be classified according to pathophysiology
- Multiple mechanisms may be involved in low back pain in what is referred to as a "mixed pain" state