What Every Specialist Should Know About Osteoporosis Today



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High-Risk Populations Require Special Attention

Rheumatoid Arthritis

22-50% prevalence due to chronic inflammation and glucocorticoid therapy

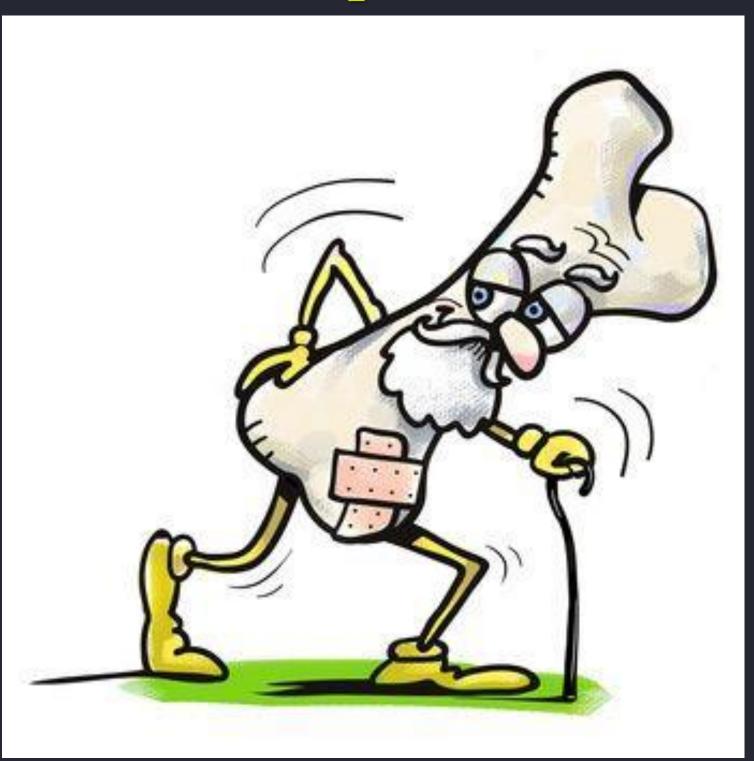
Chronic Kidney Disease

33-81% develop osteopenia in stages3-5D from metabolic disruptions

Post-Transplant Patients

Elevated risk from immunosuppressive medications and metabolic complications

Osteoporosis



Osteoporosis Overview



Bone Deterioration



Population Impact

Low bone mass and microarchitectural breakdown characterize disease

1 in 3 women, 1 in 5 men affected after ag e 50



Economic Burden

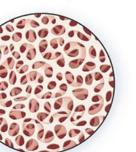
Annual fracture costs exceed \$20 billion in US alone



Osteoporosis

High bone density

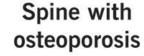


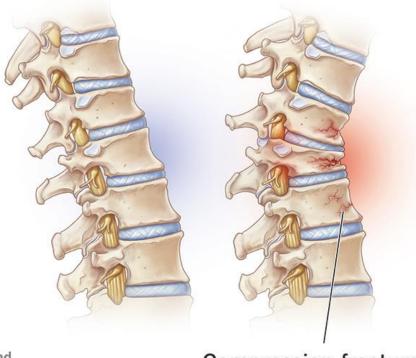


Low bone density Osteoporosis



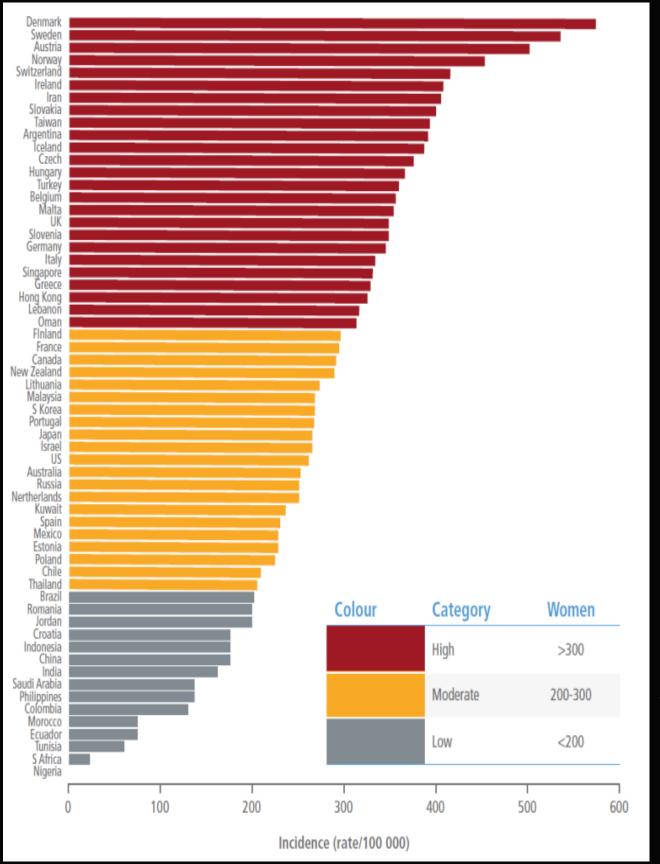
Healthy spine







Compression fracture



The Global Burden By the Numbers

200M

30%

\$20B

Global Prevalence

Postmenopausal Women

Annual U.S. Costs

People worldwide affected by osteoporosis

At risk for osteoporotic fractures

Healthcare expenditure for osteoporotic fractures

Fracture-related mortality reaches 20-30% in the first year following hip fracture, making osteoporosis a critical public health priority across all specialties.









Key Performance Metrics

30%

Fracture Reduction

Fall prevention strategies effectiveness rate

\$19B

Annual Costs

Healthcare expenses from fractures

10M

Americans Affected

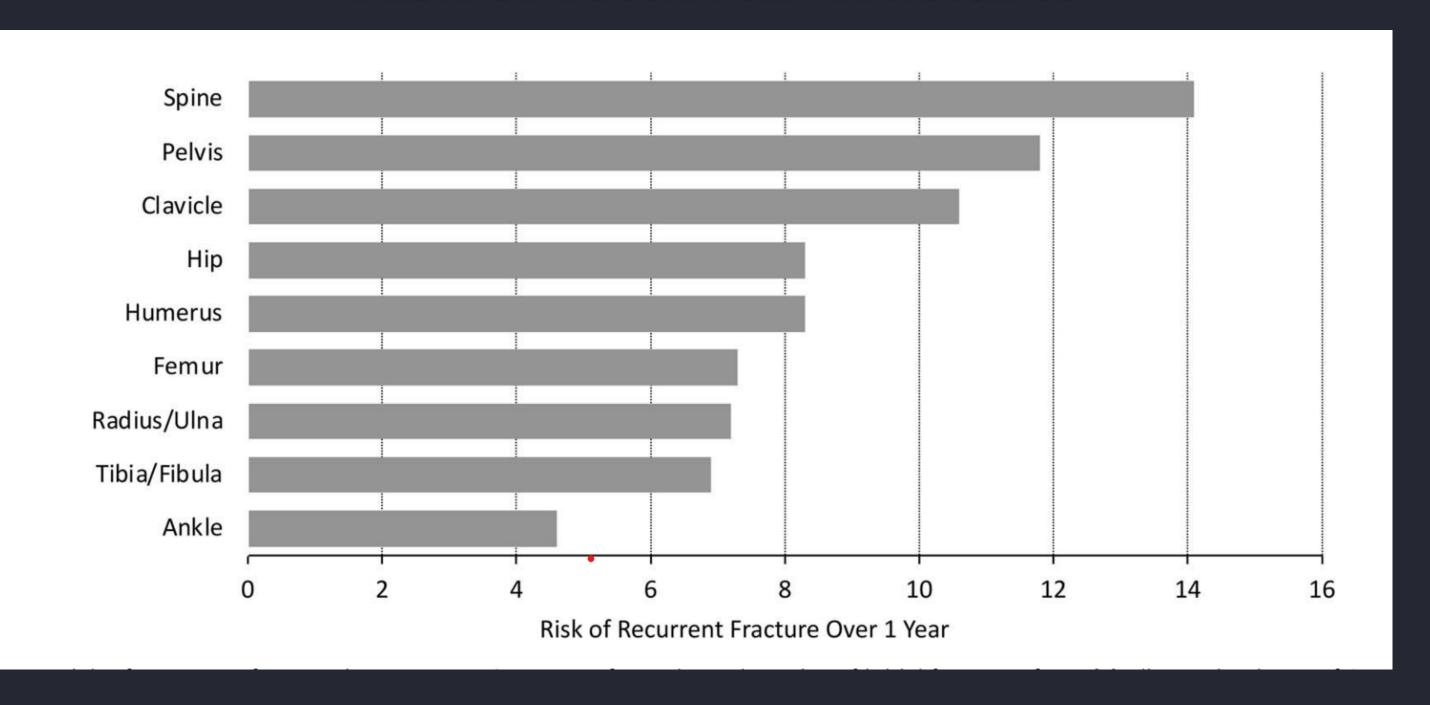
Current osteoporosis patient population

44M

At-Risk Population

Americans with low bone mass

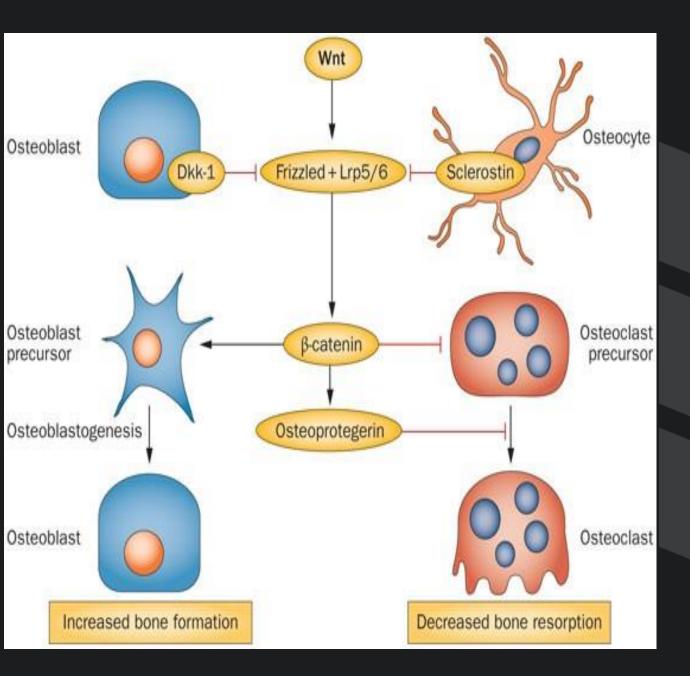
Risk of recurrent fractures



MENOPAUSE (SANDROPAUSE)



Pathogenesis Cellular and Molecular Insights





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Disrupted osteoblast formation and osteoclast resorption

Pathway Disruption

Abnormal metabolic and molecular signaling identified

RANK Signaling

RANK/RANKL/OPG pathway controls bone remodeling

Pathophysiology

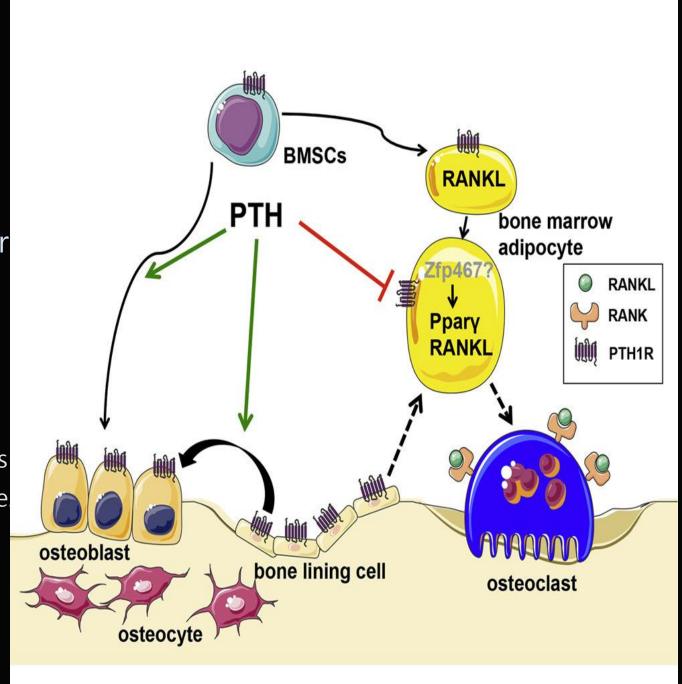
Bone Remodeling Imbalance

Osteoporosis develops when osteoclast activity exceeds osteoblast function. Key drivers include:

- Aging-related hormone decline
- Inflammatory cytokines (TNFα, IL-6)
- Glucocorticoid-induced suppression

CKD-Mineral Bone Disorder

Secondary hyperparathyroidism, reduced calcitriol activation, and uremic toxins create complex high- or low-turnover bone diseas patterns in chronic kidney disease.



Understanding Bone Remodeling



Wnt Signaling Pathway

Dysfunction in this critical disrupts bone formation and resorption balance



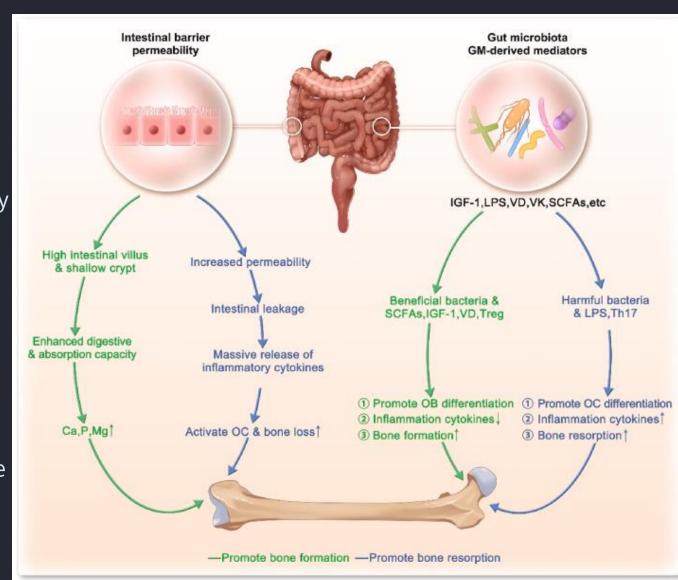
Genetic Factors

Genome-wide studies & hundreds of susceptibility loci.

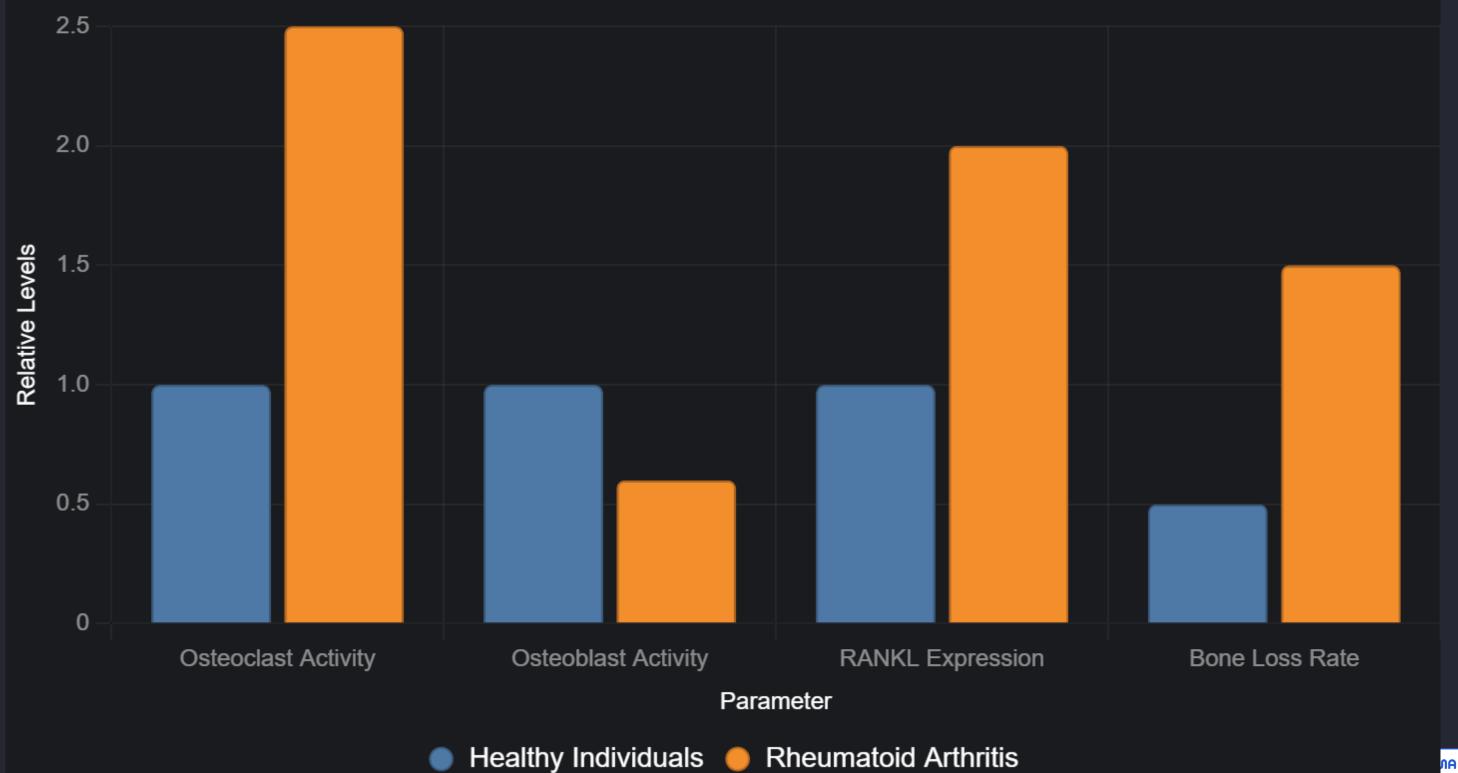


Emerging Mechanisms

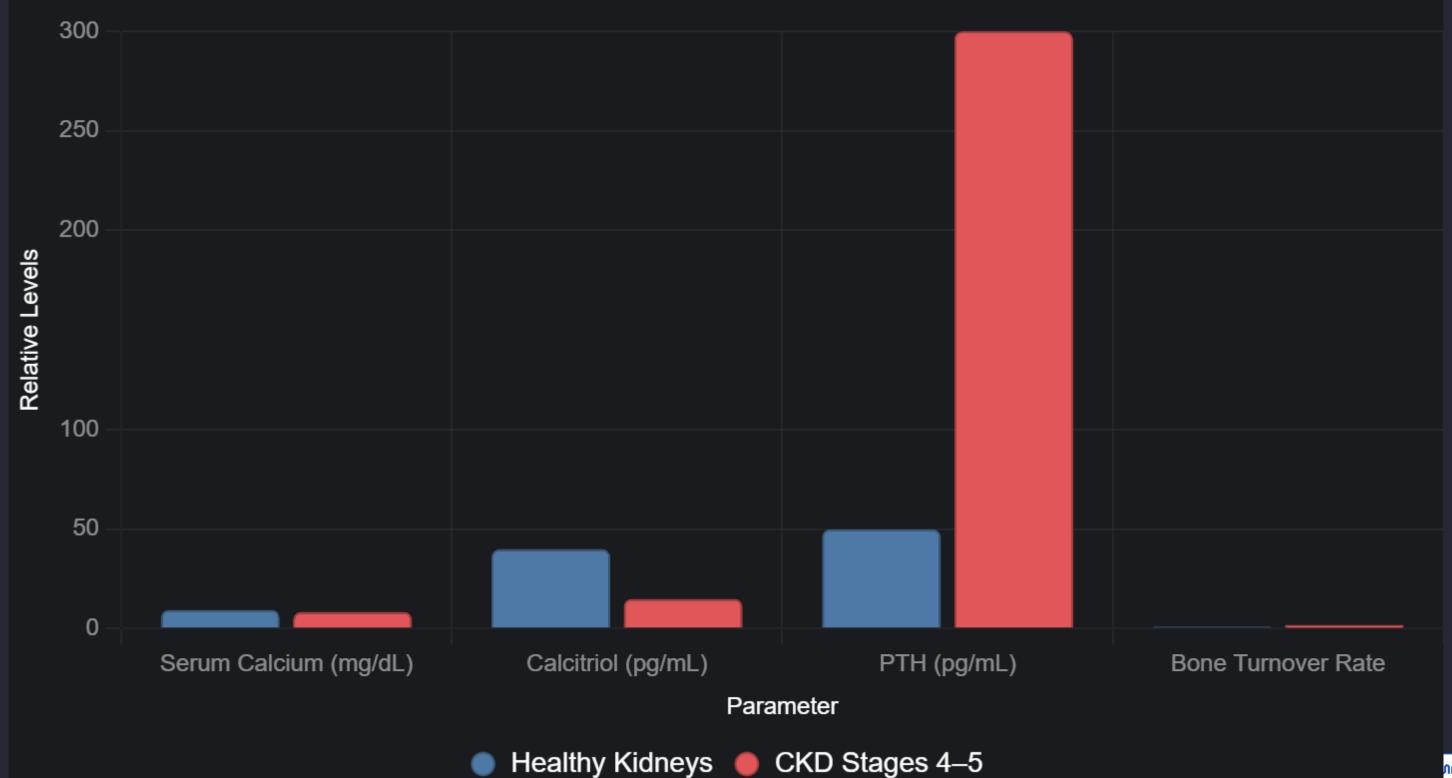
Gut microbiome, autophagy, iron balance, and cellular senescence



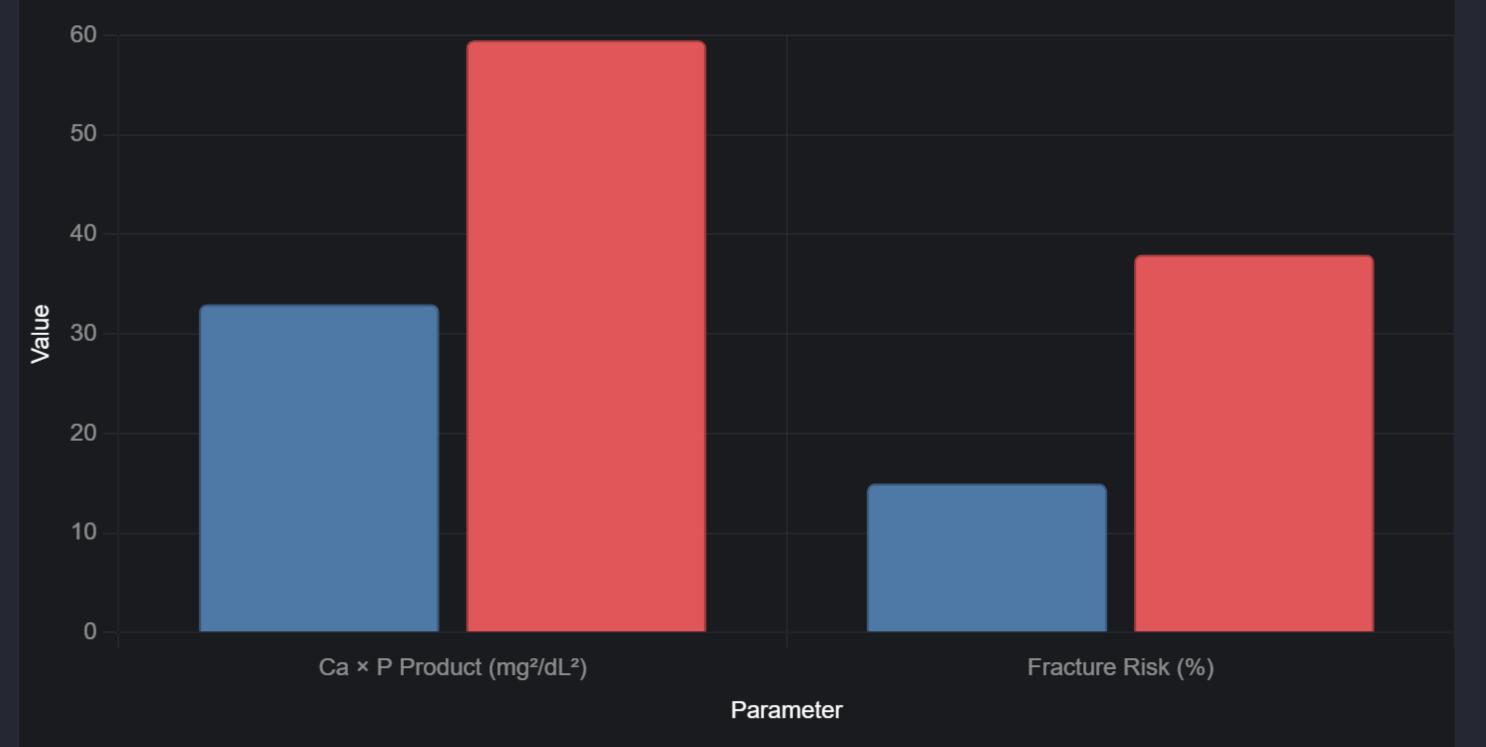
Bone Remodeling in Autoimmune Disease (RA)



Calcium Metabolism: Healthy Kidneys vs. CKD

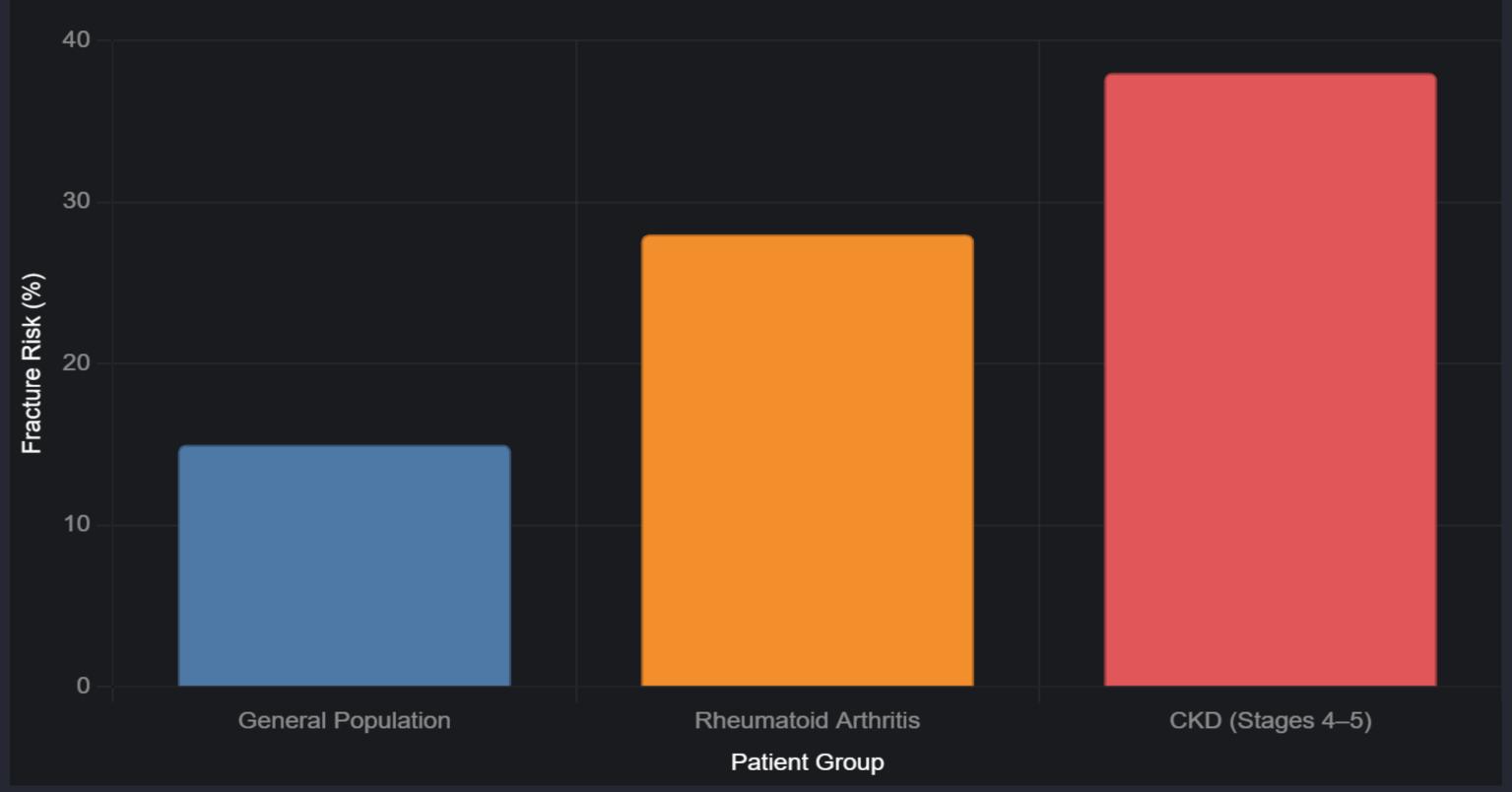


Ca × P Product and Fracture Risk: Healthy vs. CKD



Healthy Kidneys CKD Stages 4–5

Osteoporotic Fracture Risk in Rheumatology and Nephrology



Clinical Implementation Challenges

30%

50%+

Fracture Reduction

Underdiagnosis

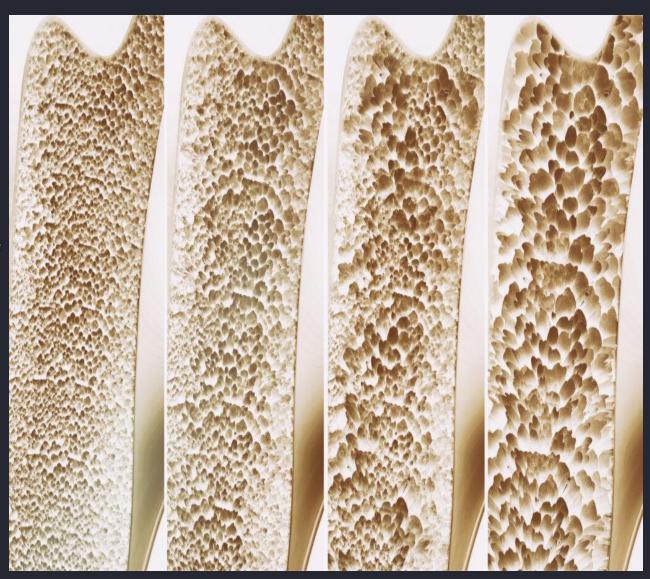
Raloxifene decreases vertebral fracture risk

Many patients only identified after fracture

40%

Non-adherence

Limits effectiveness of available treatments



Updated Screening Guidelines (2025)

Expanded USPSTF Recommendations

All women aged 65 and older require screening.

Post-menopausal women now qualify more frequently.

Risk Assessment Tools

FRAX calculator provides
standardized risk
evaluation. Earlier
screening recommended
for high-risk patients.

Enhanced Coverage

Medicare improvements support broader screening access. More patients qualify for preventive care.

Annals of Internal Medicine



VITAMIN D AND CALCIUM SUPPLEMENTATION TO PREVENT FRACTURES IN ADULTS CLINICAL SUMMARY OF U.S. PREVENTIVE SERVICES TASK FORCE RECOMMENDATION

Population	Men or premenopausal women	Community-dwelling postmenopausal women at doses of >400 IU of vitamin D ₃ and >1000 mg of calcium	Community-dwelling postmenopausal women at doses of ≤400 IU of vitamin D ₃ and ≤1000 mg of calcium	
Recommendation	No recommendation. Grade: I statement	No recommendation. Grade: I statement	Do not supplement. Grade: D	
Preventive Medications	determine the effect	f vitamin D and calcium are essential to overall hea tot of combined vitamin D and calcium supplementa premenopausal women. nce that daily supplementation with 400 IU of vitat the incidence of fractures in postmenop	ition on the incidence of fractures in men or D_3 and 1000 mg of calcium has no effect on	
	There is inadequate evidence regarding the effect of higher doses of combined vitamin D and calcium supplementation on fracture incidence in community-dwelling postmenopausal women.			
Balance of Benefits and Harms	Evidence is lacking regarding the benefit of daily vitamin D and calcium supplementation for the primary prevention of fractures, and the balance of benefits and harms cannot be determined.	Evidence is lacking regarding the benefit of daily supplementation with >400 IU of vitamin D ₃ and >1000 mg of calcium for the primary prevention of fractures in postmenopausal women, and the balance of benefits and harms cannot be determined.	Daily supplementation with ≤400 IU of vitamin D ₃ and ≤1000 mg of calcium has no net benefit for the primary prevention of fractures.	
Other Relevant USPSTF Recommendations	The USPSTF has made recommendations on screening for osteoporosis and vitamin D supplementation to prevent falls in community-dwelling older adults. These recommendations are available at www.uspreventiveservicestaskforce.org.			

or a summary of the evidence systematically reviewed in making this recommendation, the full recommendation statement, and supporting documents, pleas o to www.uspreventiveservicestaskforce.org

Country : UK	Name / ID :	About the risk factors (i)
Questionnaire) :	10. Secondary osteoporosis No Yes
1. Age (between 40-90	years) or Date of birth	11. Alcohol 3 more units per day No Yes
Age: Date of birth: Y: M: D:		12. Femoral neck BMD Select
2. Sex 3. Weight (kg)	Male Female	Clear Calculate
4. Height (cm)		BMI The ten year probability of fracture (%)
5. Previous fracture	ONo OYes	without BMD
6. Parent fractured hip	○No ○Yes	■ Major osteoporotic
7. Current smoking	ONo OYes	■ Hip fracture
8. Glucocorticoids	ONo OYes	View NOGG Guidance
9. Rheumatoid arthritis	○No ○Yes	

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Diagnostic Approach

DXA Scanning Standards

Dual-energy X-ray absorptiometry remains the gold standard. Provides precise bone mineral density measurements.

T-Score Interpretation

- Normal: >-1.0
- Low bone mass: -1.0 to -2.5
- Osteoporosis: ≤-2.5

Monitoring Schedule

Test every 1-2 years after treatment initiation. Longer intervals for low-risk patients with favorable scores.



Diagnostic Approach and Screening

DEXA Screening Protocol

Gold standard BMD measurement.
Screen women ≥65, men ≥70, or
earlier with risk factors

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FRAX Risk Assessment

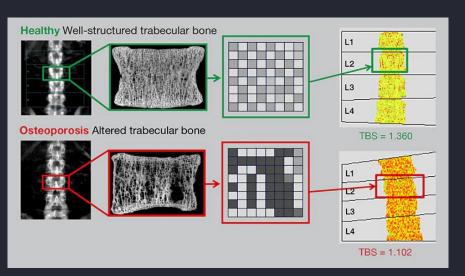
Calculate 10-year fracture probability, though may underestimate risk in CKD and autoimmune diseases

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CKD-Specific Considerations

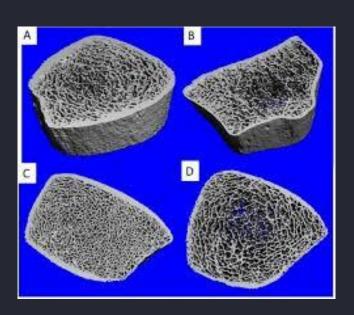
DEXA less reliable in stages 4-5D; consider quantitative CT or trabecular bone score

Diagnostic Innovations



Early Detection

TBS and HR-pQCT identify changes before DXA miRNA biomarkers enable earlier diagnosis



Cone-Beam CT

New imaging quantifies mandibular cortical measurements to identify low bone density.

Dual-Energy X-ray

Remains the standard for bone mineral density assessment.

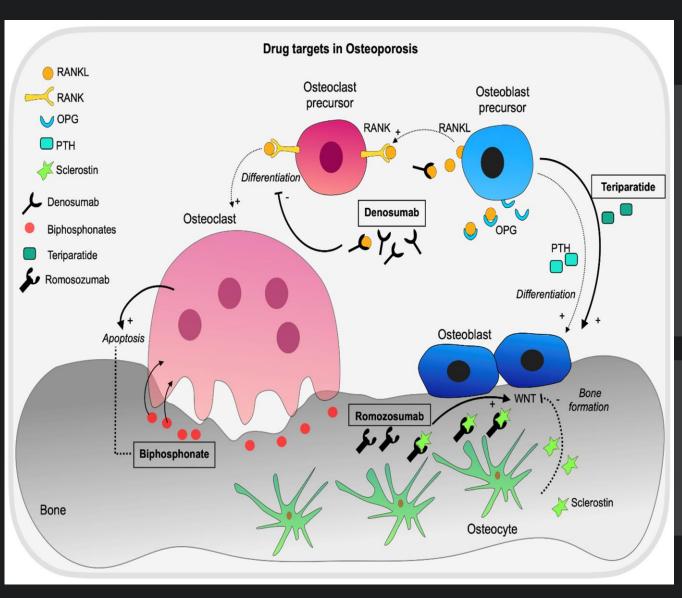
Fracture Risk Tool

Integrates clinical risk factors for better decision-making.

Advanced Imaging

Trabecular Bone Score and HR-pQCT reveal microstructure

Emerging Therapeutic Targets



Signaling Pathways

Wnt/β-catenin and RANKL inhibition show promise. Targeted pathway modulation improves outcomes.

Novel Agents

Sclerostin inhibitors and cathepsin K blockers. Phase II trials demonstrate efficacy.

RNA Therapies

Experimental RNA-based treatments under investigation. Promising preclinical results emerging.

Current Treatment Options



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Alendronate, risedronate, ibandronate, zoledronic acid



RANK Ligand Inhibitor

Denosumab for patients intolerant to bisphosphonates



Anabolic Agents

Teriparatide for severe osteoporosis cases



Alternative Therapies

Raloxifene, hormone therapy, calcitonin options

Safety Considerations



Non-Pharmacologic Management





Essential Nutrients

Calcium 1000-1200mg daily.
Vitamin D 800-1000 IU daily



Exercise Program

Weight-bearing and resistance exercises 3-5 times weekly. Progressive training



Fall Prevention

Home safety modifications and balance training.
Remove hazards and improve lighting.



Lifestyle Changes

Smoking cessation and alcohol limitation. Regular monitoring

Pharmacologic Treatment Indications

Fracture History

Any history of hip or vertebral fractures, whether clinical or asymptomatic, indicates immediate treatment need.

T-Score Criteria

Scores ≤-2.5 at femoral neck, total hip, or lumbar spine require pharmacologic intervention.

High-Risk Osteopenia

- 10-year hip fracture probability ≥3%
- Major osteoporosis fracture probability ≥20%
- Enhanced Medicare coverage supports treatment



Advanced CKD Management Challenges

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Drug Clearance Issues

Bisphosphonates contraindicated if eGFR <35 mL/min/1.73 m² due to accumulation risk

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Safer Alternative: Denosumab

Not renally cleared, but requires careful monitoring for hypocalcemia with vitamin D/calcium supplementation

CKD-MBD Targeted Therapy

Phosphate binders, calcimimetics (cinacalcet), and calcitriol analogs for secondary hyperparathyroidism

LOW risk

Normal BMD, no history of fracture, and low fracture probability

No pharmacologic treatment

MODERATE risk

Osteopenic range BMD, no history of fracture, and low fracture probability

No pharmacologic treatment; consider bisphosphonate

HIGH risk

Osteoporosis range BMD, osteopenic range BMD + spine or hip fracture, high fracture probability

> Alendronate Denosumab Risedronate Zoledronic acid

VERY HIGH risk

Endocrine Society:

Osteoporosis range BMD with spine or hip fracture, multiple spine fractures

> Abaloparatide Romosozumab Teriparatide

AACE:

Fracture within prior 12 months, fracture on osteoporosis medication, multiple fractures, fractures on bone-toxic medications, T-score < -3.0, high risk for falls, very high risk of fracture probability by FRAX^b

Abaloparatide
Denosumab
Romosozumab
Teriparatide
Zoledronic acid



Key Takeaways and Next Steps



Implement New Guidelines

Apply 2025 USPSTF screening recommendations. Expand screening to newly eligible populations.



Utilize Risk Tools

Incorporate FRAX calculator into routine assessment. Quantify individual fracture probability accurately.



Coordinate Care

Establish Fracture Liaison Services. Improve post-fracture care coordination and follow-up.



Personalize Treatment

Tailor therapy duration and selection. Monitor effectiveness with regular reassessment protocols.