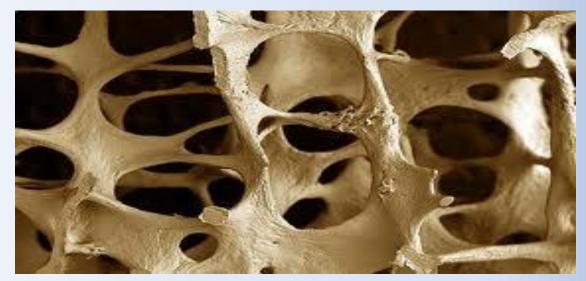


Osteoporosis Assessment & Management

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No disclosures to report





Epidemiology



- 2 million osteoporotic fractures in the United States each year
- about 50 million people in the U.S. are at risk for fracture
 - ► Diabetes: 34 million
 - Hyperlipidemia: 94 million
 - Hypertension: 75 million*



Epidemiology

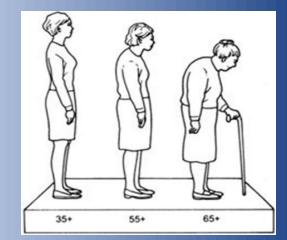


► Lifetime probability of a hip fracture in women is 10-15%

Lifetime probability of breast cancer is 12-13%



Morbidity & Mortality



- ► 15-20% will die within 1 year of a hip fracture
- ► 30% will have permanent disability
- ₩ 40% will be unable to walk independently
- ► 50% will no longer be able to live independently
- Mortality rates are higher for men than for women



Assessing Risk: Who to Screen?

- H DEXA scan
 - ► All women 65 and older
 - ► All men 70 and older*
 - ► Patients 50 and older with any nonphalangeal fracture
 - ► Patients at higher risk....



Patients at Increased Risk

- Hong-term glucocorticoid use
- H Smokers
- Heavy alcohol use
- H Inactivity
- Halabsorptive conditions
- Heumatologic conditions
- Hematologic conditions
- Heuromuscular diseases



Patients at Increased Risk

► Postmenopausal women <65

► FRAX score for major osteoporotic fracture >8.4% should get DEXA scan

ST (osteoporosis self-assessment tool)

Score < 2 should get DEXA scan

 \blacktriangleright OST = (weight in kg) – (age) 5

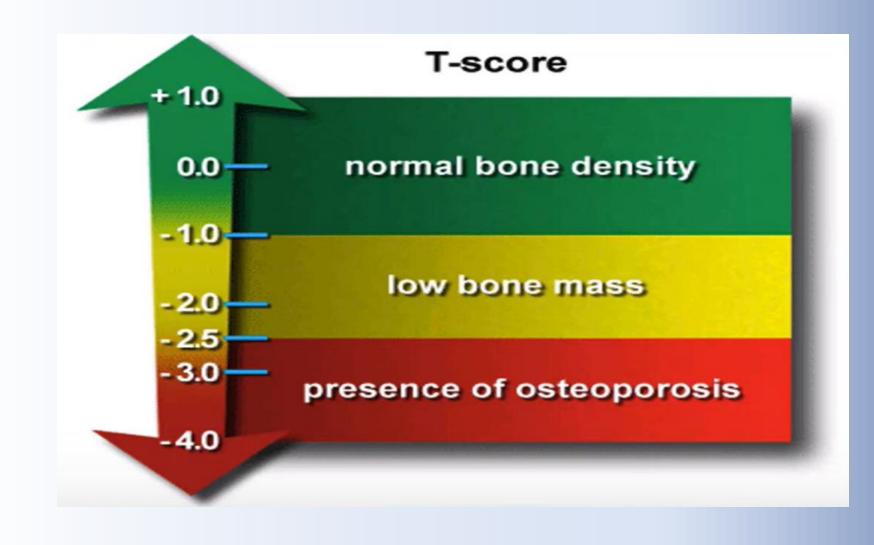


Patients at Increased Risk





DEXA Scan





DEXA Scan Rescreening

► For women 65 and older who are not taking prescription antifracture medication, suggested rescreening intervals are based on initial T-score:

Initial T-score	Suggested Minimum Interval
<u>≥</u> -1.4	10 years
-1.5 to -1.9	5 years
-2.0 to -2.4	2 years



Who to Treat?

- Start treatment in patients with hip or vertebral fractures**
- Start treatment in patients with a T-score <-2.5 SD at femoral neck, total hip, or spine on DEXA
- Treat postmenopausal women and men aged 50+ with osteopenia (T-score -1 to -2.4) with increased fracture risks
 - ► a 10-year hip fracture probability of 3% or more or
 - ► a 10-year major osteoporosis-related fracture probability of 20% or more
- Steoporosis based on clinical judgment

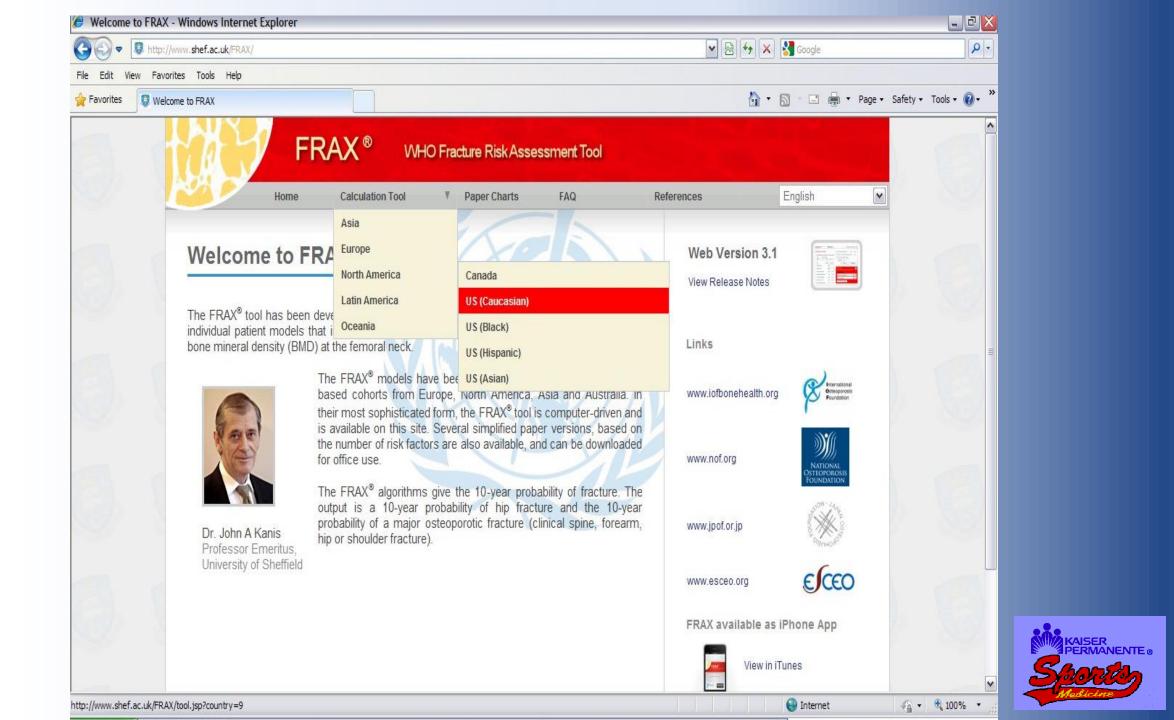


FRAX

► www.shef.ac.uk/FRAX

- Here the gold standard tool for assessing fracture risk
- In some cases, can be calculated without a DEXA score
- Can be quite useful for some of us if we aren't sure





Calculation Tool			200	
Please answer the questions below to calc	ulate the ten year probability of fracture with BMD.			
Country: US (Caucasian) Name/ID:	About the risk factors ()	1.07		
Questionnaire:	10. Secondary osteoporosis 💿 No 🔘 Yes		CDAH	
1. Age (between 40-90 years) or Date of birth	11. Alcohol 3 or more units per day 💿 No 🔘 Yes		Weight Conversion	
Age: Date of birth:	12. Femoral neck BMD (g/cm²)		Pounds 🔶 Kgs	
2. Sex 💿 Male 💿 Female	Clear Calculate			
3. Weight (kg)			Height Conversion	
4. Height (cm)			Inches 🔶 Cms	
5. Previous fracture 💿 No 🔵 Yes			Convert	
6. Parent fractured hip 💿 No 🕤 Yes				
7. Current smoking 💿 No 🕥 Yes				
8. Glucocorticoids 💿 No 🔵 Yes				

Done



Calculation Tool

Please answer the questions below to calculate the ten year probability of fracture with BMD.

Questionnaire:		10. Secondary osteoporosis 💿 No 🔘 Yi
1. Age (between 40-90 ye	ars) or Date of birth	11. Alcohol 3 or more units per day 🛛 💿 No 🔘 Yi
Age: Date of birt	h:	12. Femoral neck BMD (g/cm²)
65 Y:	M: D:	T-Score -2.1
2. Sex 🔘	Male 💿 Female	Clear Calculate
3. Weight (kg)	68.04	
4. Height (cm)	167.64	BMI 24.2
5. Previous fracture	💿 No 🕜 Yes	The ten year probability of fracture (%)
6. Parent fractured hip	💿 No 🕤 Yes	Major osteoporotic 11
7. Current smoking	💿 No 🕥 Yes	
	🖲 No 🛛 Yes	Hip fracture 1.8







Calculation Tool

Home

Please answer the questions below to calculate the ten year probability of fracture with BMD.

Questionnaire:			🖲 No 🔘 Ye 🖲 No 🔘 Ye
1. Age (between 40-90 ye: Age: Date of birt		12. Femoral neck BMD (g/cm²)	
65 Y:	M: D: D: Male 👵 Female	T-Score -2.1 Clear Calc	ulate
3. Weight (kg)	68.04	()()
4. Height (cm)	167.64	BMI 24.2	
5. Previous fracture	⊙ No () Yes	The ten year probability of fracture (%)	$\tilde{}$
6. Parent fractured hip	💿 No 🔵 Yes	Major osteoporotic	12
7. Current smoking	🔘 No 🛛 💿 Yes	Hip fracture	3.1
8. Glucocorticoids	💿 No 🕥 Yes		0.1
9. Rheumatoid arthritis	🖲 No 🕥 Yes		

ferences	English	•
1	2	
46		
X	Weight Conversion	
	Pounds 🔶 kg	
	150 Convert	
	Height Conversion	
	66 Convert	
6	01141754	
	Individuals with fracture risk assessed since 1st June 2011	



Treatment – Primary Prevention

- According to the Bone Health and Osteoporosis Foundation (BHOF), all people aged 50 and above should: ► be encouraged to engage in regular weightbearing and muscle strengthening exercises to reduce the risk of falls
 - safety-proof their home

- avoid smoking and excessive alcohol
- take 1000-1200mg of calcium daily*
- 🛏 take 800-1000 IU of vitamin D



Treatment – Primary Prevention

- According to the United States Preventive
 - Services Task Force (USPSTF):
 - Hexercise to prevent falls
 - insufficient evidence to assess balance risks/benefits for vitamin D and calcium for primary prevention



recommends against < 400 IU/day of vitamin D or < 1000 mg/day of calcium in postmenopausal women



Calcium

- Dietary supplementation is considered safest
 No more than 2000mg/day of supplements
- H Calcium carbonate
 - 🛏 cheapest
 - max absorption with 500mg doses
 - Hetter absorption after meals
 - Helps with heartburn
- H Calcium citrate
 - Hest absorption
 - preferred if on acid-blocking medication
 - Herefore preferred if history of renal stones



Treatment

- First-line treatment of osteoporosis is with **bisphosphonates**
 - 🛏 alendronate (Fosamax)
 - ₩70mg, once weekly
 - 🛏 ibandronate (Boniva)
 - ► 150mg, once monthly
 - Hack of evidence for non-vertebral fx risk
 - **zolendronic** acid (Reclast)
 - HIV, once yearly
 - Fisedronate (Actonel)
 - Hest tolerated
 - 🛏 most expensive





Bisphosphonates

- Bisphosphonates reduce the incidence of vertebral fractures by almost 50% over 3 years
- Alendronate has a little better data than the other bisphosphonates
- The number needed to treat to prevent one hip fracture per year is about 100



Estrogen receptor modulator Raloxifene (Evista)

May reduce vertebral fractures by 30-55%
 Not clinically proven to reduce hip fractures
 Incidentally lowers breast cancer risk
 Increased risk of hot flashes, leg cramps, and blood clots/DVTs



Calcitonin

🛏 Intranasal

- Can help the pain associated with vertebral fractures
- Questionably decreases risk for new vertebral fractures in established osteoporosis, but no evidence of significant effect on hip fractures
- Mot frequently used for fracture prevention



Parathyroid hormone analogs

- Heriparatide (Forteo)
- Habaloparatide (Tymlos)
- Hoaily subcutaneous injections
- ► May reduce hip fractures by 65%, and other nonvertebral fractures by 53%
- May increase risk of osteosarcoma
- HUsed for a maximum of two years





Other Medications Receptor activator of nuclear factor kappa-B (RANK) ligand inhibitor

- Henosumab (Prolia, Xgeva)
- Monoclonal antibody bone-modifying agent, used for bony metastases
- Subcutaneous injection every 6 months
- Cardiovascular, neurologic, and gastrointestinal side effects are not uncommon
- Severe hypocalcemia if advanced CKD
- ► Possibly worse bone density and increased fracture risk after discontinuation
- Possible alternative for women at high risk for fracture who cannot take bisphosphonates



Sclerostin Inhibitors

- Romosoxumab (Evenity)
- Monoclonal antibody bone-modifying agent
- Subcutaneous injection monthly for 12 months
- H Increased risk of MI, stroke, and cardiovascular death
- Possible alternative for women at high risk for fracture who cannot take bisphosphonates
- HUNIKE the PTH analogs, can be used again after stopping



Treatment

- First check: creatinine, calcium, albumin, and vitamin D level
 - GFR should be <a>>30
 - Hypocalcemia can be worsened by bisphosphonates
 - Hold Albumin to get the corrected calcium
 - Vitamin D level should be above 20 ng/ml before initiating treatment with bisphosphonates
 - ► If vitamin D level is below 20 ng/ml, treat with vitamin D2 --50,000 units once weekly for 6-12 weeks
 - Re-check a level before starting bisphosphonates



Contraindications to Alendronate

- HTrue Allergy
- 🛏 Renal
 - ₩ GFR <30
- Gastrointestinal esophageal stricture
 - 🛏 achalasia



- inability to remain upright for 30 minutes
- History of bariatric surgery
- HIV bisphosphonate may be a good choice
- Hendocrine Endocrine
 - 🛏 hypocalcemia



Bisphosphonates – Adverse Effects

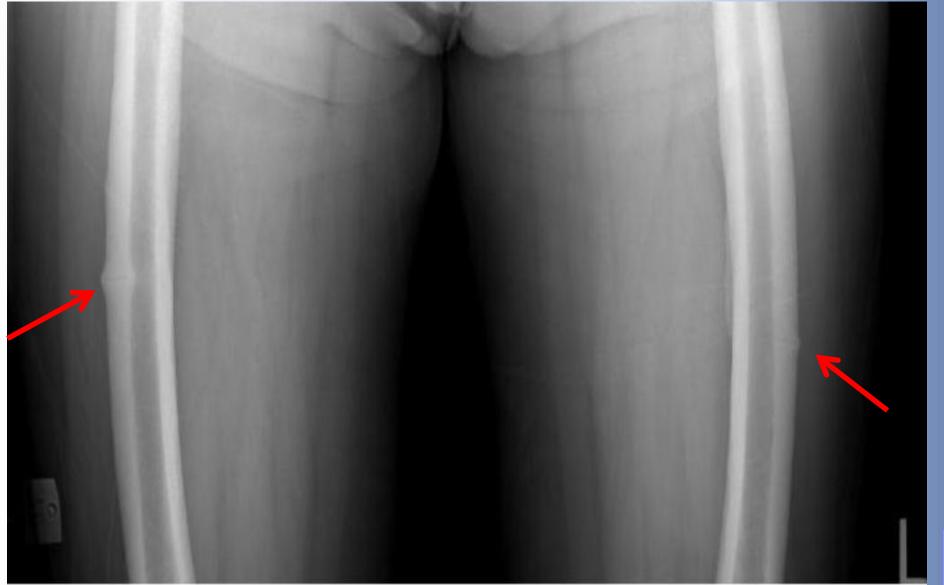
Gastrointestinal issues
 Difficulty swallowing
 Gastric ulcer



- **Esophageal inflammation**
- ► Atrial fibrillation?
- Steonecrosis of the jaw
- ► Atypical femur fractures*



Atypical Femur Fractures



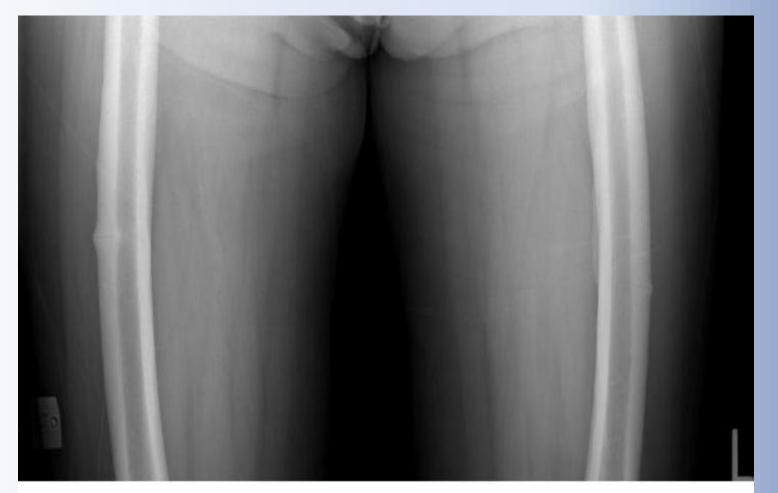


Atypical Femur Fractures





Overall Benefit of Bisphosphonates NNT for alendronate is about 100 patients NNH for alendronate is about 5000 patients*



Years of Rx with a oral Bisphosphonate	Risk/Year per 100,000 for an Atypical Femur Fracture
<1 year	2 in 100,000
1 - 1.9 years	2 in 100,000
2 - 2.9 years	3 in 100,000
3 - 3.9 years	12 in 100,000
4 - 4.9 years	16 in 100,000
5 - 5.9 years	24 in 100,000
6 - 6.9 years	43 in 100,000
7 - 7.9 years	78 in 100,000

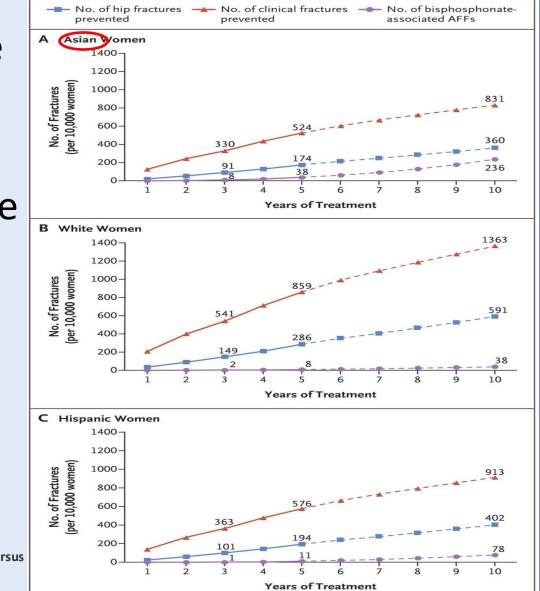
Dell R, Greene D, Ott S, et al. A retrospective analysis of all atypical femur fractures seen in a large California HMO from the years 2007 to 2009. ASBMR 2010 Annual Meeting, Toronto, Canada. 2010

Atypical Femur Fractures

- Highest incidence in Asian women
- Glucocorticoid use > 1 year also a risk factor

Benefit still outweighs risk

D.M. Black, E.J. Geiger, R. Eastell, *et al*. Atypical femur fracture risk versus fragility fracture prevention with bisphosphonates N. Engl. J. Med., 383 (8) (2020), pp. 743-753





Drug Holiday



After 5 years of oral bisphosphonate therapy (or 3 years of IV bisphosphonate), there should be a drug holiday if:

► DEXA T-score is better than -2.5, and

- Mo history of fragility fracture, and
- ► Not on bone-losing medication
- Reassess with DEXA scan every 2 years
 - Resume treatment for any of the above changing or for bone loss >5% between tests



Osteoporotic hip fractures affect as many as 1 in 8 women, resulting in 10-20% excess mortality

Screening with DEXA is recommended in:

- Hall women 65 and older
- ► all men 70 and older*
- ► fracture patients 50 and older
- Higher risk patients (50 and older)



- ► Primary prevention in people 50+ (BHOF):
 - ➡ 1000-1200mg of calcium daily
 - 🛏 800-1000 IU of vitamin D
 - regular weightbearing and muscle strengthening exercises to reduce the risk of falls
 - safety-proofing the home
 - avoid smoking and excessive alcohol



Start treatment in patients with: ► a hip or a vertebral fracture ► a Dexa T-score of -2.5 or lower ► a FRAX 10-year probability of: ► 3% or more for hip fracture or ► 20% or more for major osteoporotic fracture



- Bisphosphonates are the medication of choice to treat osteoporosis
 - Most side effects are gastrointestinal in nature
 - Serious side effects are rare but notable
 - Benefits outweigh serious risks by 50 to 1
 - A 5-year timespan of treatment is currently recommended, with reassessment and a possible 2-year drug holiday
 - A repeat DXA scan can be checked at 2 years



Questions/Comments

