VERTEBRAL FRACTURE INITIATIVE Part I

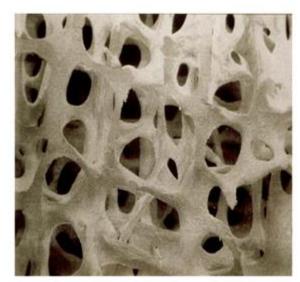
Overview of osteoporosis: Epidemiology and clinical management

By Pawel Szulc and Mary Bouxsein



Definition of osteoporosis

A disease characterized by low bone mass and micro-architectural deterioration of bone tissue leading to reduced bone strength and a consequent increase in fracture risk.



Normal bone



Osteoporosis



OSTEOPOROSIS: THE SIZE OF THE PROBLEM



Osteoporosis is a prevalent disease

- Estimated to affect 200 million women worldwide
 - 1/10 of women aged 60
 - 1/5 of women aged 70
 - 2/5 of women aged 80
 - 2/3 of women aged 90

From European, North American, Japanese and Australian cohorts¹

- Worldwide, an osteoporotic fracture occurs every 3 seconds, a vertebral fracture every 22 seconds²
- 20-25% Caucasian women and men over the age of 50 years have a prevalent vertebral fracture³, 12% in Latin American women and men⁴



Lifetime risk at the age of 50

	Women	Men
Osteoporotic fracture ^{1,2}	46-53%	21-22%
Hip fracture ^{2,3}	15-23%	5-11%
Radiographic vertebral fracture ⁴	27%	11%
Clinical vertebral fracture ²	15%	8%
Breast cancer	10-13%	
Prostate cancer	9-1	11%
	NB: variable between countries	

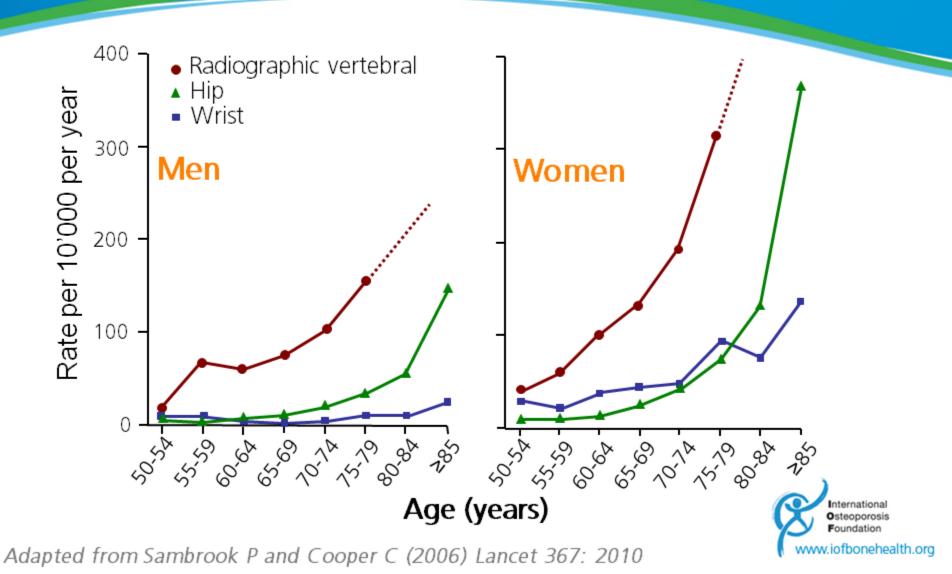
¹Van Staa TP et al (2001) Bone 29: 517

²Kanis JA et al (2000) Osteoporos Int 11: 669

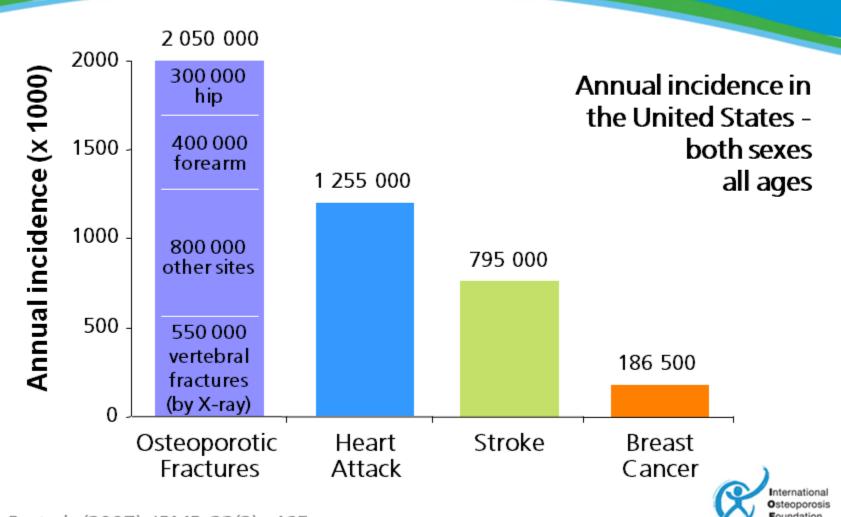
³Samelson EL et al (2007) J Bone Miner Res 22: 1449 ⁴Samelson EL et al (2006) J Bone Miner Res 21: 1207



Age-specific and sex-specific incidence of osteoporotic fractures



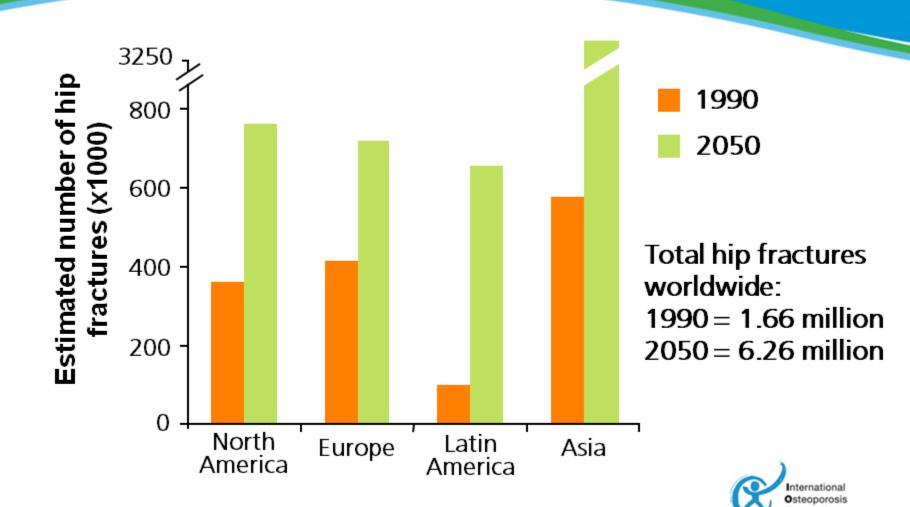
Comparison of osteoporotic fractures with other common diseases



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Burge R et al. (2007) JBMR 22(3): 465 United States Cancer Statistics 2005, American Heart Association 2006

Increased burden of osteoporotic fractures worldwide

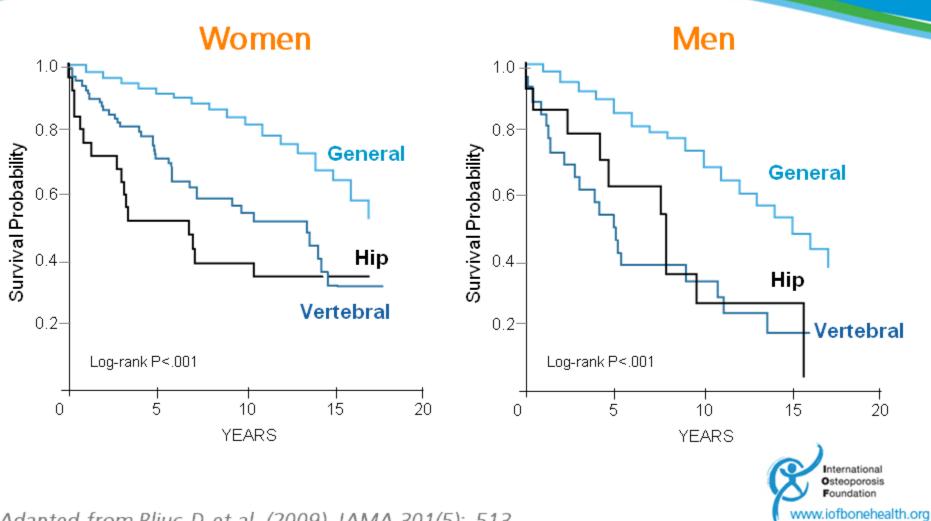


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OSTEOPOROSIS: MORTALITY & MORBIDITY

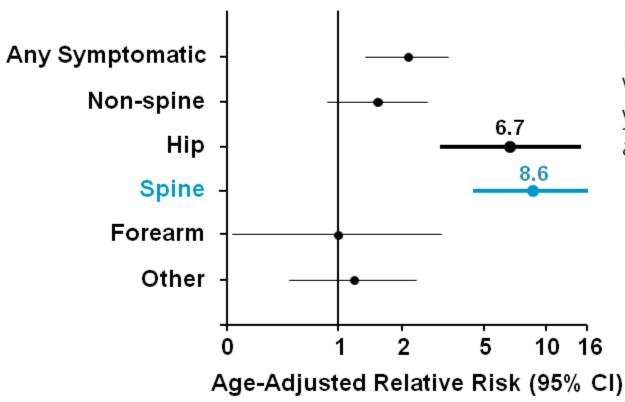


Similar mortality in patients with vertebral fractures and in those with hip fractures



Relative risk of death following clinical osteoporotic fractures

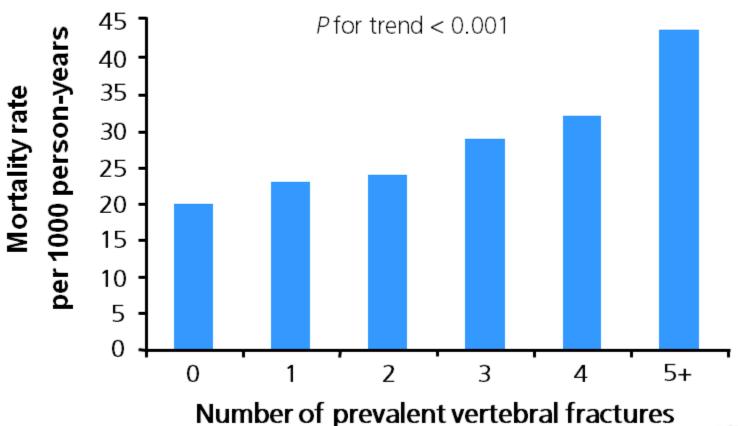
Fracture Intervention Trial (FIT)*



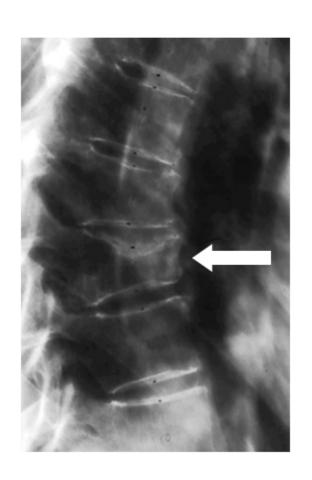
*6459 postmenopausal women ages 55-81 years followed for an average of 3.8 years



Mortality rate increases with the number of prevalent vertebral fractures



Vertebral fractures increase the risk of subsequent fragility fracture



 Women with vertebral fractures have a 5-fold increased risk of a new vertebral fracture and a 2-fold increased risk of hip fracture

Black DM et al. (1999) J Bone Miner Res 14: 821 Melton LJ 3rd et al. (1999) Osteoporos Int 10: 214

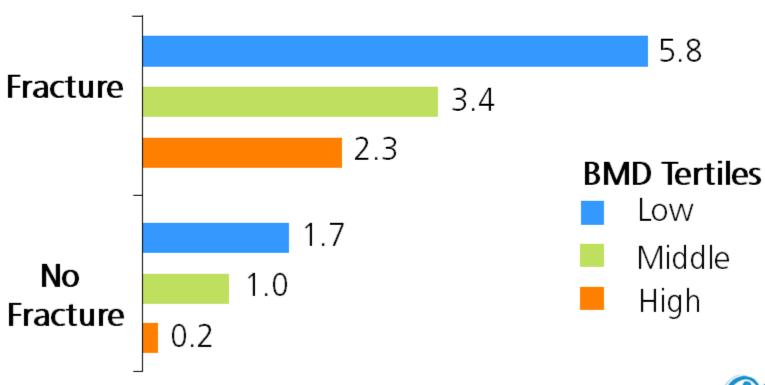
 Of women who suffer a vertebral fracture, 1 in 5 will suffer another vertebral fracture within a year

Lindsay et al. (2001) JAMA 285(3): 320

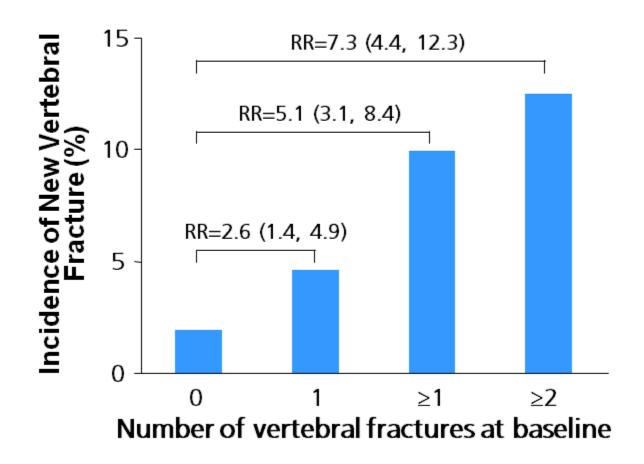


Prior fracture increases the risk for future vertebral fracture, independent of BMD

Risk of vertebral fractures (% per year)



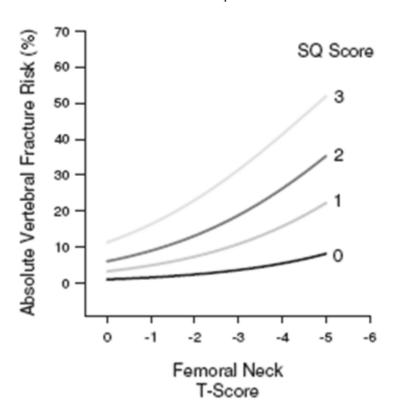
Prior vertebral fracture increases the risk of subsequent vertebral fracture

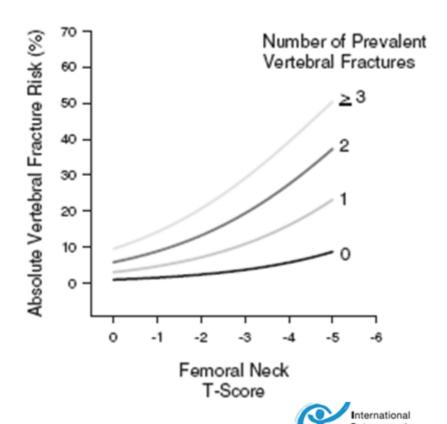




Incident vertebral fracture risk: effect of BMD and prevalent vertebral fractures

SQ score: worst grade of any fractured vertebra according to Genant semi-quantitative scale



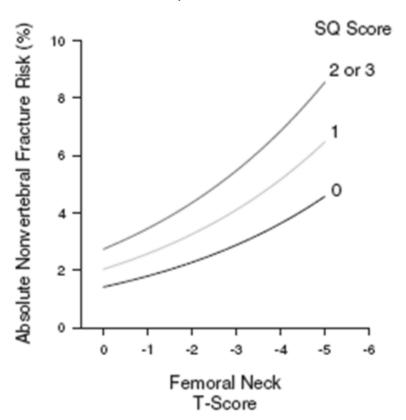


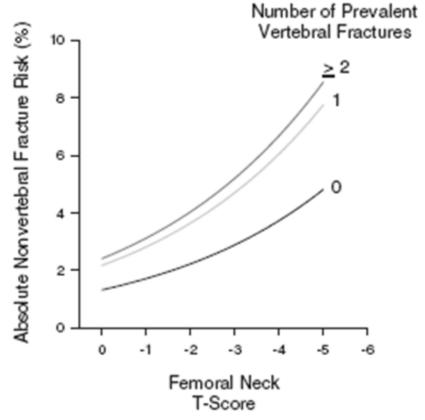
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Siris ES et al. (2007) Osteoporosis Int 18: 761 With kind permission from Springer Science+Business Media

Incident non-vertebral fracture risk: effect of BMD and prevalent vertebral fractures

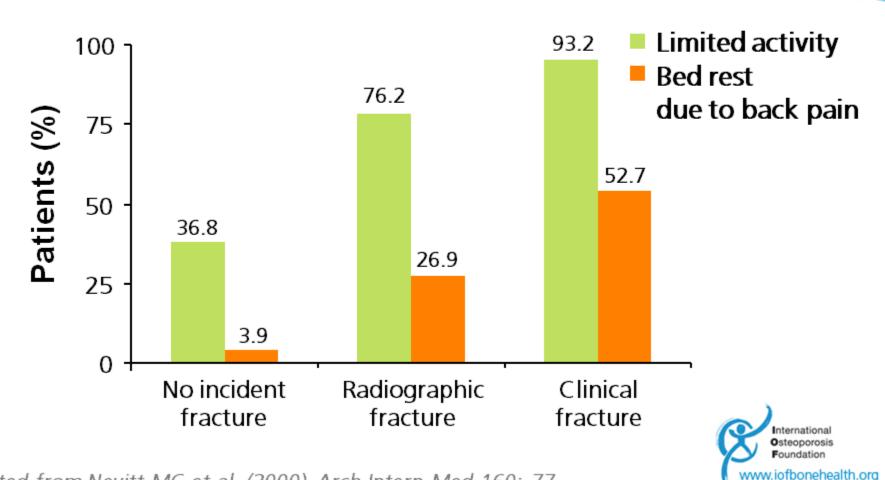
SQ score: worst grade of any fractured vertebra according to Genant semi-quantitative scale





Siris ES et al. (2007) Osteoporosis Int 18: 761 With kind permission from Springer Science+Business Media

All types of vertebral fractures are associated with morbidity



Consequences of vertebral fractures



- Kyphosis
- Loss of height
- Bulging abdomen
- Acute and chronic back pain
- Breathing difficulties
- Depression
- Reflux and other GI symptoms
- Difficulty with activity of daily living (bending, rising, dressing, climbing stairs)
- Need to use a walking aid



REDUCED INDEPENDENCE
AND QUALITY OF LIFE

Treatment of symptomatic vertebral fractures

- Bed rest (in the case of severe pain)
- Pharmacological treatment
 - analgesics
 - opioids
- Physical therapy
- Bracing
- Local steroid injections
- Vertebral augmentation
 - Vertebroplasty
 - Kyphoplasty



Summary

Vertebral Fractures

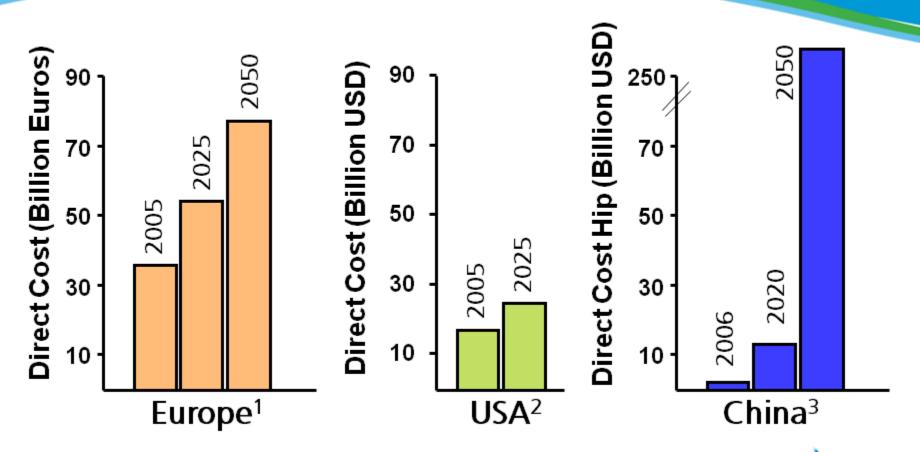
- Are the most common osteoporotic fracture
- Are associated with excess mortality
- Are associated with significant morbidity, even if they do not come to clinical attention
- Increase the risk of subsequent vertebral fractures by 5-fold and the risk of other fragility fractures (including hip) by 2 to 4-fold



COST OF OSTEOPOROSIS



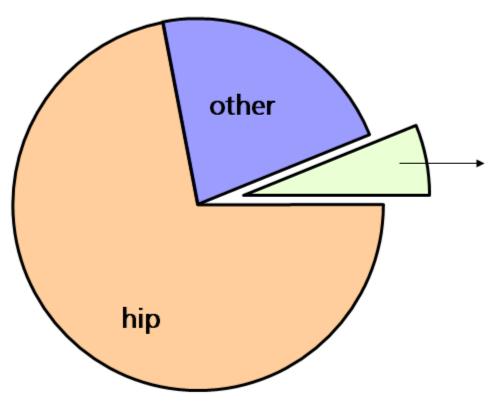
Projected costs of osteoporotic fractures



- 1. Kanis JA, Johnell O (2005) Osteoporos Int 16: 229
- 2. Burge R et al. (2007) J Bone Miner Res 22: 465
- 3. Mithal A et al. (2009) The Asian Audit, IOF: 15



Direct costs of vertebral fractures



Vertebral fractures:

6% of the total direct cost of osteoporotic fractures, costing

1 billion USD in the US in 2005

In Europe, the cost was 719 million EUR in 2005



Vertebral fractures induce back pain and disability

Risk associated with incident vertebral fracture (Odd's Ratio) 7223 women >65 yrs, 3.7 yrs follow-up

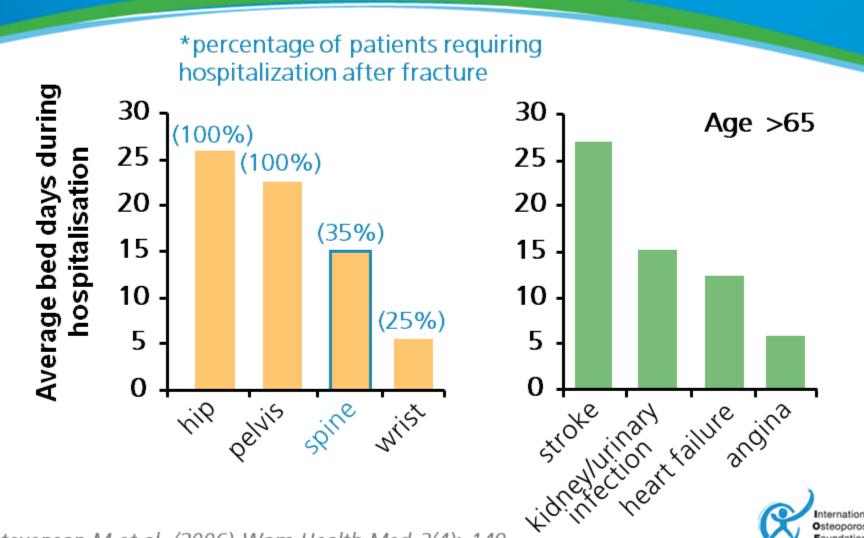
Vertebral fracture at baseline	NO	YES
Back pain	2.0	2.7
Back disability	1.8	4.1
≥ 1 day bed rest	4.9	7.5
≥ 7 days limited activity	2.7	7.4

Incident ______
vertebral fracture

+ 1 to 2 days per year of bed rest

+ 10 days per year of limited activity

Average hospital utilisation for osteoporotic fractures in the UK



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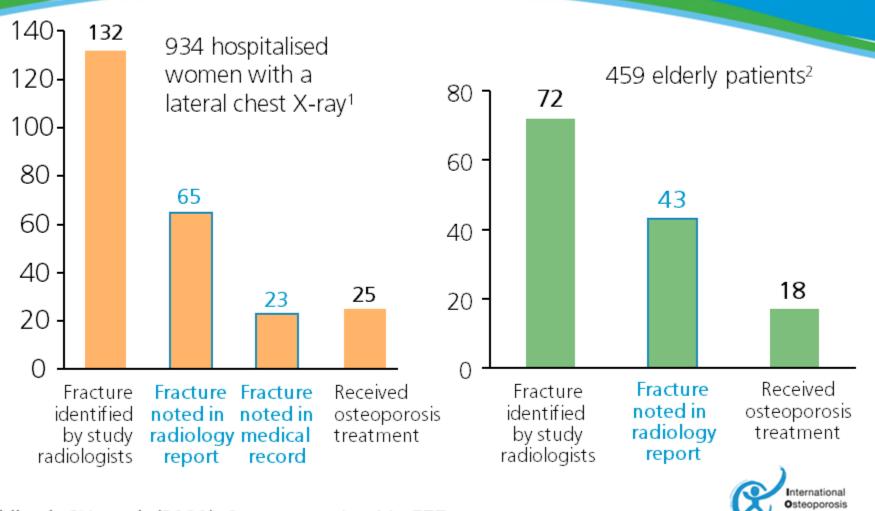
Stevenson M et al. (2006) Wom Health Med 3(4): 149 Ham C et al. (2003) Brit Med J 327(7426): 1257

UNDER-DIAGNOSIS OF VERTEBRAL FRACTURES:

A MISSED OPPORTUNITY



Under-diagnosed vertebral fractures



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¹Gehlbach SH et al. (2000) Osteoporos Int 11: 577 ²Majumdar SR et al. (2005) Arch Intern Med 165: 905

Under-diagnosis of vertebral fractures is a worldwide problem

Vertebral fractures are under-diagnosed in all geographic regions

|--|

North America	45.2%
Latin America	46.5%
Europe/South Africa/Australia	29.5%
Global rate	34.0%

 Attributable both to poor identification of vertebral fractures and, when a fracture is identified, use of ambiguous terminology in the patient's medical record that does not clearly state vertebral fracture.



Awareness and treatment of vertebral fractures is low despite...

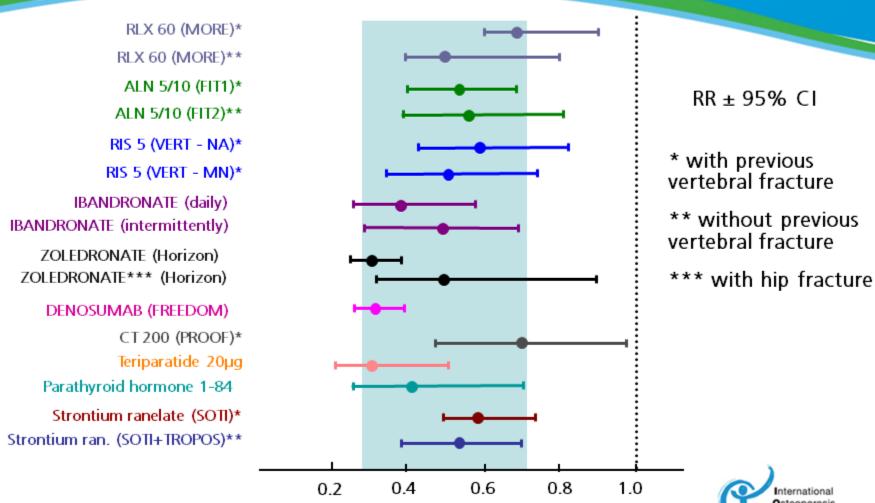
- Known disability associated with vertebral fractures
- Excess mortality associated with vertebral fractures
- Every year, over 1billion USD spent in the US and over 750 million EUR in Europe for treating vertebral fractures alone (direct costs)
- Validated radiologic techniques for diagnosis
- Effective and safe treatments
- Evidence based guidelines for diagnosis and management of osteoporosis, including vertebral fractures



Effective therapies are widely available and can reduce vertebral fractures by 30% to 70%.



Effect of osteoporosis drugs on new vertebral fractures



Conclusions

- Vertebral fractures are a common complication of osteoporosis and markedly increase the likelihood of subsequent fractures
- Mild and moderate vertebral fractures are often not being recognised and reported, leading to under-diagnosis and under-treatment
- Lateral spine radiographs or by lateral DXA imaging are the best ways to confirm the presence of vertebral fractures in clinical practice
- Early radiographic diagnosis followed by appropriate therapy will help prevent subsequent fractures
- Effective therapies are widely available and can reduce vertebral, hip and other fragility fractures by 30% to 70%.

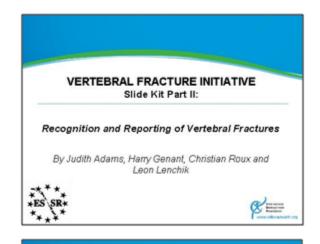


Action is needed by radiologists and other clinicians to ensure:

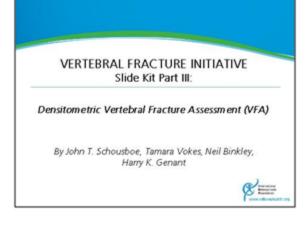
- Recognition of vertebral fractures using radiography,
 DXA-VFA or other imaging techniques
- Reporting as FRACTURED to avoid ambiguity
- These actions will help patients receive effective treatment and prevent subsequent fractures



RECOGNITION & REPORTING OF VERTEBRAL FRACTURES



Part II
Recognition and Reporting of
Vertebral Fractures



Part III

DXA-based Vertebral Fracture

Assessment

