

Analyses of the cost-effectiveness of pooled alendronate and risedronate, compared with strontium ranelate, raloxifene, etidronate and teriparatide. Updated following change in price of generic alendronate.

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#### Introduction

This report is an update of a similar report (compiled in July 2006) that takes the price change of generic alendronate into account. The previous report used a cost of £264 per annum for alendronate and risedronate treatment. This price used in the analyses has been reduced to £173, the price of generic alendronate. <sup>1</sup> Additionally for sensitivity analyses the price of proton pump inhibitors to counter side effects have been reduced to £6 per compliant patient. Excluding these change, the methodology and parameter values are identical in both reports.

#### The base-case scenario

The base-case is assumed to be that described in Table 1.

Table 1: The base-case scenario.

Parameter	Value	Source
Persistence at 5-years	50%	Estimated from the results of
		the accompanying literature
		review
The assumed relative risk	0.71 – 'hip'	Systematic Review and meta-
of bisphosphonates on	0.58 – 'spine'	analysis of alendronate and
osteoporotic fractures.	0.78 – 'prox hum'	risedronate data. See Appendix
	0.78 – 'wrist'	1.
Costs set to those used in	Age dependent, see	Updated costs used in previous
the initial report	previous report	NICE assessments of
		osteoporosis interventions.
Utility multiplier	Year 1 0.626	Kanis et al. Osteoporosis
associated with vertebral	Year 2+ 0.909	International 2004; 15 20-26.
fracture.		This source was used for all
		fracture types
Costs incurred over 5-	£4.50 per patient that	See earlier text
years via side effects	is compliant (costs	
associated with	for non-compliant	
bisphosphonate	patients are included	
	in our analyses)	
Utility multiplier	0.91	Groenveld et al <sup>2</sup>
associated with	(utility losses for	
bisphosphonate related GI	non-compliant	
symptoms	patients are included	
	in our analyses)	
Cost of bisphosphonate	£173 per annum	Price of alendronate.

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<sup>&</sup>lt;sup>1</sup> http://www.ppa.org.uk/edt/September 2006/mindex.htm. Accessed 01/09/06.

<sup>&</sup>lt;sup>2</sup> Groeneveld PW, Lieu TA, Fendrick M, Hurley LB, Ackerson LM, Levin TR and Allison JE. "Quality of life measurements clarifies the cost-effectiveness of Helicobacter Pylori eradication in peptic ulcer disease and uninvestigated dyspepsia" The American Journal of Gastroenterology. 2001 96 (2) 338 - 347

#### Summarised results for women identified through opportunistic assessment)

P	How scenario is different from the base-case.	Identification strategies potentially <sup>3</sup> costeffective from what age (years)?	Percentage of women age 50 or older that were opportunistically assessed that would be offered a BMD scan (%) <sup>V</sup>	Percentage of women age 50 or older that were opportunistically assessed that would be treated (%) <sup>∇Ψ</sup>
Base-case	- -	70	25.7	4.6
1	Persistence set to 25%	70	25.7	2.3
2	Persistence set to 75%	65	26.7	8.1
3	Efficacy assumed to be different in the osteoporotic, osteopenic and normal women, and equal to that from the FIT trial.	65	38.5	4.7
4	Efficacy of bisphosphonate set to 50% for clinical risk factor other than BMD and fracture status.	70	25.7	4.3
5	Efficacy of bisphosphonate set to 0% for clinical risk factor other than BMD and fracture status. Questions are not asked in this scenario.	75	22.2	3.2
6	The costs calculated by Stevenson et al <sup>4</sup> to be used instead of the older costs, including potential home help costs.	65	34.8	7.8
7	The costs calculated by Stevenson et al to be used instead of the older costs, excluded potential home help costs.	65	26.7	5.4
8	HRG costs to be used, including potential home help costs.	70	26.7	5.3
9	HRG costs to be used, excluding potential home help costs.	70	25.7	4.5
10	Vertebral disutility to be set to equal that associated with a hip fracture.	70	25.7	4.5
11	The costs and disutility from side effects to be double that estimated	70	25.7	4.5
12	The costs and disutility from side effects to be half that estimated	70	25.7	5.3
13	The costs and disutility from side effects to be set to zero	65	26.7	5.4
14	The disutility from side effects to be set to ten times that of the base-case	70	25.7	1.3
15	The costs from side effects increased were proton pump inhibitors are prescribed instead of H2 receptor antagonists.	70	25.7	4.6

<sup>&</sup>lt;sup>3</sup> Assuming a cost per QALY of £20,000 <sup>4</sup> Stevenson MD, Davis SE, Kanis JA. "The hospitalisation costs and out-patient costs of fragility fractures". Women's Health Medicine. In Press.

#### <u>Summarised results for self-identifying women</u>

	How scenario is different from the base-case.	Identification strategies potentially cost-effective from what age (years)?	Percentage of women age 50 or older that were opportunistically assessed that would be offered a BMD scan (%) $^{\nabla}$	Percentage of women age 50 or older that were opportunistically assessed that would be treated (%) $^{\nabla \psi}$
Base-case	-	50	64.4	19.3
1	Persistence set to 25%	55	55.5	9.1
2	Persistence set to 75%	50	64.4	30.1
3	Efficacy assumed to be different in the osteoporotic, osteopenic and normal women, and equal to that from the FIT trial.	50	100	11.6
4	Efficacy of bisphosphonate set to 50% for clinical risk factor other than BMD and fracture status.	60	60.1	18.3
5	Efficacy of bisphosphonate set to 0% for clinical risk factor other than BMD and fracture status.	65	59.3	15.8
6	The costs calculated by Stevenson et al <sup>6</sup> to be used instead of the older costs, including potential home help costs.	50	67.9	21.9
7	The costs calculated by Stevenson et al to be used instead of the older costs, excluded potential home help costs.	55	64.4	19.9
8	HRG costs to be used, including potential home help costs.	55	64.4	19.4
9	HRG costs to be used, excluding potential home help costs.	55	55.5	18.9
10	Vertebral disutility to be set to equal that associated with a hip fracture.	55	55.5	14.9
11	The costs and disutility from side effects to be double that estimated	55	55.5	18.3
12	The costs and disutility from side effects to be half that estimated	50	64.4	19.4
13	The costs and disutility from side effects to be set to zero	50	64.4	19.4
14	The disutility from side effects to be set to ten times that of the base-case	55	52.4	13.9
15	The costs from side effects increased were proton pump inhibitors are prescribed instead of H2 receptor antagonists.	50	55.9	18.9

<sup>&</sup>lt;sup>5</sup> Assuming a cost per QALY of £20,000 <sup>6</sup> Stevenson MD, Davis SE, Kanis JA. "The hospitalisation costs and out-patient costs of fragility fractures". Women's Health Medicine. In Press.

#### <u>Detailed results for women found through opportunistic assessment.</u>

#### Scenario Base-case 1

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£11,259
		treat where T-	treat where T-	treat where T-	
		Score <-2.5	Score <-1.0	Score <-0.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£8,774
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-2.0	Score <-0.5	Score < 0.5SD	
	SD	SD	SD		

Sensitivity Analysis 1-1: Base-case, bar persistence set to 25%

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£12,053
		treat where T-	treat where T-	treat where T-	
		Score <-2.5	Score <-1.5	Score <-0.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£10,177
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-2.0	Score <-0.5	Score < 0.0	
	SD	SD	SD	SD	

#### Sensitivity Analysis 2-1: Base-case, bar persistence set to 75%

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
65-69 years	Do not BMD	Do not BMD	BMD and	BMD and	£18,829
			treat where T-	treat where T-	
			Score <-2.0	Score <-1.5	
			SD	SD	
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£11,482
		treat where T-	treat where T-	treat where T-	
		Score <-2.0	Score <-1.0	Score <-0.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£8,451
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-1.5	Score <-0.5	Score < 0.5	
	SD	SD	SD	SD	

# Sensitivity Analysis 3-1: Base-case, bar efficacy assumed to be different in the osteoporotic, osteopenic and normal women, and equal to that from the FIT trial.

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
65-69 years	Do not BMD	BMD and	BMD and	BMD and	£12,169
		treat where T-	treat where T-	treat where T-	
		Score <-2.5	Score <-2.5	Score <-2.5	
		SD	SD	SD	
70-74 years	BMD and	BMD and	BMD and	BMD and	£7,246
	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-2.5	Score <-2.5	Score <-2.5	
	SD	SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£969
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-2.5	Score <-2.5	Score <-2.5	
	SD	SD	SD	SD	

### Sensitivity Analysis 4-1: Base-case, bar efficacy of bisphosphonate set to 50% for clinical risk factor other than BMD and fracture status.

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£15,357
		treat where T-	treat where T-	treat where T-	
		Score <-2.5	Score <-2.0	Score <-1.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£11,344
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-2.0	Score <-1.5	Score <-0.5	
	SD	SD	SD	SD	

# Sensitivity Analysis 5-1: Base-case, bar efficacy of bisphosphonate set to 0% for clinical risk factor other than BMD and fracture status. Questions on clinical risk factors are assumed not to be asked.

Age (years)	0 Clinical Risk Factors	1 Clinical Risk Factor	2 Clinical Risk Factors	3 Clinical Risk Factors	Cost Per QALY of strategy		
75 years and over	BMD and treat	BMD and treat where T-Score <-2.5 SD					

## Sensitivity Analysis 6-1: Base-case, bar using the costs calculated by Stevenson et al to be used instead of the older costs, including potential home help costs

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
65-69 years	Do not BMD	Do not BMD	BMD and	BMD and	£16,732
			treat where T-	treat where T-	
			Score <-2.0	Score <-1.5	
			SD	SD	
70-74 years	BMD and	BMD and	BMD and	BMD and	£13,340
	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-2.0	Score <-1.0	Score <-0.5	
	SD	SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£7,848
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.0	Score <-1.5	Score < 0.0	Score < 0.5	
	SD	SD	SD	SD	

## Sensitivity Analysis 7-1: Base-case, bar using the costs calculated by Stevenson et al to be used instead of the older costs, excluding potential home help costs

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
65-69 years	Do not BMD	Do not BMD	BMD and	BMD and	£17,956
			treat where T-	treat where T-	
			Score <-2.5	Score <-1.5	
			SD	SD	
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£10,001
		treat where T-	treat where T-	treat where T-	
		Score <-2.0	Score <-1.0	Score <-0.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£7,097
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-1.5	Score <-0.5	Score < 0.5	
	SD	SD	SD	SD	

## Sensitivity Analysis 8-1: Base-case, bar using HRG costs including potential home help costs

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£11,218
		treat where T-	treat where T-	treat where T-	
		Score <-2.0	Score <-1.0	Score <-0.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£8,823
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-1.5	Score <-0.5	Score < 0.5	
	SD	SD	SD	SD	

## Sensitivity Analysis 9-1: Base-case, bar using HRG costs excluding potential home help costs

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£11,430
		treat where T-	treat where T-	treat where T-	
		Score <-2.5	Score <-1.5	Score <-0.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£10,171
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-2.0	Score <-0.5	Score < 0.5	
	SD	SD	SD	SD	

## Sensitivity Analysis 10-1: Base-case, bar vertebral fracture disutility reduced to that of hip fracture

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£12,434
		treat where T-	treat where T-	treat where T-	
		Score <-2.5	Score <-1.5	Score <-1.0	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£10,178
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-2.0	Score <-1.0	Score < 0.0	
	SD	SD	SD	SD	

## Sensitivity Analysis 11-1: Base-case, bar disutility and costs associated with side effects are doubled.

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£11,281
		treat where T-	treat where T-	treat where T-	
		Score <-2.5	Score <-1.5	Score <-0.5	
		SD	SD	SD	
75 years	Do not BMD	BMD and	BMD and	BMD and	£9,207
and over		treat where T-	treat where T-	treat where T-	
		Score <-2.5	Score <-0.5	Score < 0.0	
		SD	SD	SD	

## Sensitivity Analysis 12-1: Base-case, bar disutility and costs associated with side effects are halved.

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£11,311
		treat where T-	treat where T-	treat where T-	
		Score <-2.0	Score <-1.0	Score <-0.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£11,860
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-1.5	Score <-0.5	Score < 0.5	
	SD	SD	SD	SD	

## Sensitivity Analysis 13-1: Base-case, bar disutility and costs associated with side effects are set to zero.

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
65-69 years	Do not BMD	Do not BMD	BMD and	BMD and	£19,977
			treat where T-	treat where T-	
			Score <-2.5	Score <-2.0	
			SD	SD	
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£11,417
		treat where T-	treat where T-	treat where T-	
		Score <-2.0	Score <-1.0	Score <-0.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£8,550
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-1.5	Score < 0.0	Score < 0.5	
	SD	SD	SD	SD	

## Sensitivity Analysis 14-1: Base-case, bar disutility from bisphosphonate side effects set to 10 times that of the base-case

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£11,449
		treat where T-	treat where T-	treat where T-	
		Score <-3.5	Score <-2.5	Score <-1.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£7,886
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-3.5	Score <-3.0	Score <-2.0	Score <-1.0	
	SD	SD	SD	SD	

# Sensitivity Analysis 15-1: Base-case, bar costs associated with side effects increased to £6 per patient due to the assumption that proton pump inhibitors are prescribed instead of H2 receptor agonists

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£11,293
		treat where T-	treat where T-	treat where T-	
		Score <-2.5	Score <-1.0	Score <-0.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£8,854
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-2.0	Score <-0.5	Score < 0.5	
	SD	SD	SD	SD	

#### Detailed results for self-identifying women

#### Scenario Base-case 2

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
50-54 years	Do not BMD	Do not BMD	BMD and treat	£19,818
			where T-Score	
			<-2.0 SD	
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£14,414
			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£15,321
		where T-Score	where T-Score	
		<-2.5 SD	<-2.0 SD	
65 - 69 years	BMD and treat	BMD and treat	BMD and treat	£15,087
	where T-Score	where T-Score	where T-Score	
	<-3.0 SD	<-2.5 SD	<-1.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£10,028
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.0 SD	<0.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£6,373
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-0.5 SD	<0.5 SD	

#### Sensitivity Analysis 1-2: Base-case, bar persistence set to 25%

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£15,043
			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£15,830
		where T-Score	where T-Score	
		<-2.5 SD	<-2.0 SD	
65 - 69 years	Do not BMD	BMD and treat	BMD and treat	£13,881
		where T-Score	where T-Score	
		<-2.5 SD	<-2.0 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£11,171
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.5 SD	<-0.5 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£7,628
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-1.0 SD	<0.5 SD	

#### Sensitivity Analysis 2-2: Base-case, bar persistence set to 75%

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
50 - 54 years	Do not BMD	Do not BMD	BMD and treat	£19,229
			where T-Score	
			<-2.0 SD	
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£14,674
			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£14,804
		where T-Score	where T-Score	
		<-2.5 SD	<-1.5 SD	
65 - 69 years	BMD and treat	BMD and treat	BMD and treat	£15,170
	where T-Score	where T-Score	where T-Score	
	<-2.5 SD	<-2.0 SD	<-1.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£9,533
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.0 SD	<0.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£5,970
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-0.5 SD	<0.5 SD	

# Sensitivity Analysis 3-2: Base-case, bar efficacy assumed to be different in the osteoporotic, osteopenic and normal women, and equal to that from the FIT trial.

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
50-54 years	BMD and treat	BMD and treat	BMD and treat	£13,314
	where T-Score	where T-Score	where T-Score	
	<-2.5 SD	<-2.5 SD	<-2.5 SD	
55 - 59 years	BMD and treat	BMD and treat	BMD and treat	£7,589
	where T-Score	where T-Score	where T-Score	
	<-2.5 SD	<-2.5 SD	<-2.5 SD	
60-64 years	BMD and treat	BMD and treat	BMD and treat	£6,084
	where T-Score	where T-Score	where T-Score	
	<-2.5 SD	<-2.5 SD	<-2.5 SD	
65 - 69 years	BMD and treat	BMD and treat	BMD and treat	£4,320
	where T-Score	where T-Score	where T-Score	
	<-2.5 SD	<-2.5 SD	<-2.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£202
	where T-Score	where T-Score	where T-Score	
	<-2.5 SD	<-2.5 SD	<-2.5 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	Dominating
over	where T-Score	where T-Score	where T-Score	
	<-2.5 SD	<-2.5 SD	<-2.5 SD	

## Sensitivity Analysis 4-2: Base-case, bar efficacy of bisphosphonate set to 50% for clinical risk factor other than BMD and fracture status.

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
60 - 64 years	Do not BMD	Do not BMD	BMD and treat	£15,667
			where T-Score	
			<-2.5 SD	
65 - 69 years	BMD and treat	BMD and treat	BMD and treat	£17,440
	where T-Score	where T-Score	where T-Score	
	<-3.0 SD	<-2.5 SD	<-2.0 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£11,320
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.5 SD	<-1.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£8,007
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-1.0 SD	<0.0 SD	

# Sensitivity Analysis 5-2: Base-case, bar efficacy of bisphosphonate set to 0% for clinical risk factor other than BMD and fracture status. Questions on clinical risk factors are assumed not to be asked.

Age (years)		CPQ of
		strategy
65 - 69 years	BMD and treat where T-Score <-3.0 SD	£19,992
70-74 years	BMD and treat where T-Score <-2.0 SD	£12,234
75 years and	BMD and treat where T-Score <-1.5 SD	£9,507
over		

## Sensitivity Analysis 6-2: Base-case, bar using the costs calculated by Stevenson et al to be used instead of the older costs, including potential home help costs

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
50-54 years	Do not BMD	Do not BMD	BMD and treat	£16,954
			where T-Score	
			<-2.0 SD	
55 - 59 years	Do not BMD	BMD and treat	BMD and treat	£16,462
		where T-Score	where T-Score	
		<-2.5 SD	<-1.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£13,225
		where T-Score	where T-Score	
		<-2.5 SD	<-1.5 SD	
65 - 69 years	BMD and treat	BMD and treat	BMD and treat	£7,414
	where T-Score	where T-Score	where T-Score	
	<-2.5 SD	<-2.0 SD	<-1.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£7,414
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.0 SD	<0.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£4,571
over	where T-Score	where T-Score	where T-Score	
	<-1.0 SD	<-0.5 SD	<1.0 SD	

## Sensitivity Analysis 7-2: Base-case, bar using the costs calculated by Stevenson et al to be used instead of the older costs, excluding potential home help costs

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£17,888
			where T-Score	
			<-2.0 SD	
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£13,594
			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£13,349
		where T-Score	where T-Score	
		<-2.5 SD	<-1.5 SD	
65 - 69 years	BMD and treat	BMD and treat	BMD and treat	£14,161
	where T-Score	where T-Score	where T-Score	
	<-2.5 SD	<-2.5 SD	<-1.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£8,424
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.0 SD	<0.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£4,755
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-0.5 SD	<0.5 SD	

## Sensitivity Analysis 8-2: Base-case, bar using HRG costs including potential home help costs

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£19,430
			where T-Score	
			<-2.0 SD	
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£14,497
			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£14,652
		where T-Score	where T-Score	
		<-2.5 SD	<-1.5 SD	
65 - 69 years	BMD and treat	BMD and treat	BMD and treat	£14,121
	where T-Score	where T-Score	where T-Score	
	<-3.0 SD	<-2.5 SD	<-1.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£9,211
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.0 SD	<0.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£6,103
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-0.5 SD	<0.5 SD	

## Sensitivity Analysis 9-2: Base-case, bar using HRG costs excluding potential home help costs

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£14,663
			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£14,991
		where T-Score	where T-Score	
		<-2.5 SD	<-2.0 SD	
65 - 69 years	Do not BMD	BMD and treat	BMD and treat	£12,188
		where T-Score	where T-Score	
		<-2.5 SD	<-1.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£10,455
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.0 SD	<0.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£7,437
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-0.5 SD	<0.5 SD	

## Sensitivity Analysis 10-2: Base-case, bar vertebral fracture disutility reduced to that of hip fracture

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
55-59 years	Do not BMD	Do not BMD	BMD and treat	£15,817
			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£16,143
		where T-Score	where T-Score	
		<-2.5 SD	<-2.0 SD	
65 - 69 years	Do not BMD	BMD and treat	BMD and treat	£13,038
		where T-Score	where T-Score	
		<-2.5 SD	<-2.0 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£9,546
	where T-Score	where T-Score	where T-Score	
	<-2.5 SD	<-2.0 SD	<-0.5 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£8,896
over	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.0 SD	<0.0 SD	

## Sensitivity Analysis 11-2: Base-case, bar disutility and costs associated with side effects are doubled.

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£14,683
			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£15,632
		where T-Score	where T-Score	
		<-2.5 SD	<-2.0 SD	
65 - 69 years	Do not BMD	BMD and treat	BMD and treat	£12,147
		where T-Score	where T-Score	
		<-2.5 SD	<-1.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£10,286
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.5 SD	<-0.5 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£6,776
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-1.0 SD	<0.5 SD	

## Sensitivity Analysis 12-2: Base-case, bar disutility and costs associated with side effects are halved.

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
50-54 years	Do not BMD	Do not BMD	BMD and treat	£19,319
			where T-Score	
			<-2.0 SD	
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£14,850
-			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£14,968
-		where T-Score	where T-Score	
		<-2.5 SD	<-1.5 SD	
65 - 69 years	BMD and treat	BMD and treat	BMD and treat	£14,715
	where T-Score	where T-Score	where T-Score	
	<-3.0 SD	<-2.5 SD	<-1.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£9,752
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.0 SD	<0.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£6,210
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-0.5 SD	<0.5 SD	

## Sensitivity Analysis 13-2: Base-case, bar disutility and costs associated with side effects are set to zero.

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
rige (years)	Factor	Factors	Factors	strategy
50-54 years	Do not BMD	Do not BMD	BMD and treat	£18,825
-			where T-Score	
			<-2.0 SD	
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£14,667
			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£14,609
		where T-Score	where T-Score	
		<-2.5 SD	<-1.5 SD	
65 - 69 years	BMD and treat	BMD and treat	BMD and treat	£14,372
	where T-Score	where T-Score	where T-Score	
	<-3.0 SD	<-2.5 SD	<-1.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£9,631
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.0 SD	<0.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£5,995
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-0.5 SD	<0.5 SD	

## Sensitivity Analysis 14-2: Base-case, bar disutility from bisphosphonate side effects set to 10 times that of the base-case

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
55-59 years	Do not BMD	Do not BMD	BMD and treat	£16,856
			where T-Score	
			<-2.5 SD	
60-64 years	Do not BMD	Do not BMD	BMD and treat	£12,607
-			where T-Score	
			<-2.0 SD	
65 - 69 years	Do not BMD	BMD and treat	BMD and treat	£11,339
		where T-Score	where T-Score	
		<-3.0 SD	<-2.0 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£9,319
-	where T-Score	where T-Score	where T-Score	
	<-2.5 SD	<-2.0 SD	<-1.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£5,683
over	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.5 SD	<-0.0 SD	

# Sensitivity Analysis 15-2: Base-case, bar costs associated with side effects increased to £6 per patient due to the assumption that proton pump inhibitors are prescribed instead of H2 receptor agonists

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
50-54 years	Do not BMD	Do not BMD	BMD and treat	£19,908
			where T-Score	
			<-2.0 SD	
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£14,500
			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£15,412
		where T-Score	where T-Score	
		<-2.5 SD	<-2.0 SD	
65 - 69 years	Do not BMD	BMD and treat	BMD and treat	£11,881
		where T-Score	where T-Score	
		<-2.5 SD	<-1.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£10,105
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.0 SD	<0.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£6,443
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-0.5 SD	<0.5 SD	

### <u>4.1 Detailed analysis for each intervention following opportunistic assessment of clinical risk factors.</u>

Summarised results are given in Table 4 followed by the individual results for each intervention. (Tables 5 to 9)

Table 4. Summarised strategies for each intervention for women <u>identified through</u> opportunistic assessment (base-case scenario).

Intervention analysed	Identification	Percentage of	Percentage of
	strategies cost-	women age 50 or	women age 50 or
	effective from what	older that would	older that would
	age (years)?	be offered a	be treated (%) $\nabla \psi$
		BMD scan (%) $^{\nabla}$	
Pooled alendronate and risedronate	70	25.7	4.6
Strontium ranelate	75	5.7	0.2
Raloxifene	None	0.0	0.0
Etidronate	70	25.7	9.1
Teriparatide	75	1.2	0.0

<sup>&</sup>lt;sup>▽</sup> These are the BMD Scans and people treated assuming that all women with a prior fracture were opportunistically screened immediately. Once this had been achieved, the numbers will be significantly reduced, assuming that opportunistic assessment regarding clinical risk factors would be undertaken once every 5 years, and that people on treatment would not be re-assessed.

Table 5. The base-case results for pooled alendronate and risedronate in women identified by opportunistic assessment. (same as matrix 'base-case 1')

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£11,259
		treat where T-	treat where T-	treat where T-	
		Score <-2.5	Score <-1.0	Score <-0.5	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£8,774
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-2.5	Score <-2.0	Score <-0.5	Score < 0.5SD	
	SD	SD	SD		

<sup>&</sup>lt;sup>Ψ</sup> These numbers have taken persistence into account. Thus, where persistence is 50%, double this number would be offered treatment.

Table 6. The base-case results for strontium ranelate in women identified by opportunistic assessment.

Age (years)	0 Clinical Risk	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Factor	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
75 years	Do not BMD	BMD and	BMD and	BMD and	£15,848
and over		treat where T-	treat where T-	treat where T-	
		Score <-3.5	Score <-2.5	Score <-2.0	
		SD	SD	SD	

Table 7. The base-case results for raloxifene in women identified by opportunistic assessment.

Opportunistic assessment strategies have cost per QALYs of >£20,000 at all ages.

Table 8. The base-case results for etidronate in women identified by opportunistic assessment.

Age (years)	0 Clinical	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Risk Factors	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
70-74 years	Do not BMD	BMD and	BMD and	BMD and	£17,064
		treat where T-	treat where T-	treat where T-	
		Score <-1.5	Score <-0.5	Score <1.0	
		SD	SD	SD	
75 years	BMD and	BMD and	BMD and	BMD and	£16,490
and over	treat where T-	treat where T-	treat where T-	treat where T-	
	Score <-1.5	Score <-1.0	Score < 0.0	Score <1.0	
	SD	SD	SD	SD	

Table 9. The base-case results for teriparatide in women identified by opportunistic assessment.

Age (years)	0 Clinical Risk	1 Clinical	2 Clinical	3 Clinical	Cost Per
	Factor	Risk Factor	Risk Factors	Risk Factors	QALY of
					strategy
75 years	Do not BMD	Do not BMD	BMD and	BMD and	£16,830
and over			treat where T-	treat where T-	
			Score <-4.0	Score <-3.0	
			SD	SD	

Illustrative cost per QALY values for each intervention compared with no treatment are given in Table 10 for women of different ages, at a T-Score in the range –2.5 to – 3.0 SD, assuming that they have no clinical risk factors. These values include neither the costs of assessment nor the costs of BMD scanning.

Table 10. Cost per QALY values for each intervention compared with no treatment. For women with a T-Score in the range -2.5 to -3.0 SD and no clinical risk factors.

	50	60	70	75
Pooled alendronate	£71,344	£48,900	£21,213	£14,893
and risedronate				
Strontium ranelate	£218,040	£140,125	£57,939	£47,144
Raloxifene	£788,772	£312,558	£83,367	£70,977
Etidronate	£124,373	£59,866	£17,243	£13,744
Teriparatide	£522,441	£369,429	£179,154	£148,713

Table 11 gives the incremental cost-effectiveness of moving from each intervention to pooled alendronate and risedronate. Pooled alendronate and risedronate is considered more cost-effective than both strontium ranelate and raloxifene. However because of the lower price of etidronate a move from etidronate to pooled alendronate and risedronate would not be considered cost-effective as the cost per QALY ratio is greater than £60,000 in the examples provided.

Table 11. Cost per QALY values for pooled alendronate and risedronate compared with each intervention. For women with a T-Score in the range –2.5 to –3.0 SD and no clinical risk factors.

	50	60	70	75
Strontium	Pooled	Pooled	Pooled	Pooled
ranelate	alendronate and	alendronate and	alendronate and	alendronate and
	risedronate	risedronate	risedronate	risedronate
	dominates	dominates	dominates	dominates
	strontium	strontium ranelate	strontium	strontium
	ranelate		ranelate	ranelate
Raloxifene	Pooled	Pooled	Pooled	Pooled
	alendronate and	alendronate and	alendronate and	alendronate and
	risedronate	risedronate	risedronate	risedronate
	dominates	dominates	dominates	dominates
	raloxifene	raloxifene	raloxifene	raloxifene
Etidronate	£42,686	£37,708	£37,659	£18,689
Teriparatide *	£5.1 m	£4.2 m	£2.7 m	£2.2 m

<sup>\*</sup> Teriparatide provides more QALYs than pooled alendronate and risedronate, but costs more. In this circumstance cost per QALY ratios greater than £20,000 are desirable.

Thus, given our current efficacy and pricing assumptions that etidronate could be considered the most cost-effective treatment for younger women with a BMD of -2.5 to -3.0 SD.

The strategy for opportunistically assessing women and subsequently providing BMD scans for women with a self-identifying fracture is however unaffected by the choice of bisphosphonate, i.e. it is identical for the pooled alendronate and risedronate and for etidronate and is thus unaffected by whichever intervention was chosen as first line treatment. From our data, etidronate could be cost-effectively prescribed to women at less severe T-Score thresholds than pooled alendronate and risedronate. However for more severe patients (particular at risk of hip fracture) the pooled alendronate and risedronate treatment option becomes more cost-effective.

## 4.2 Detailed analysis for each intervention for women presenting with a self-identifying risk factor.

Summarised results are given in Table 12 followed by the individual results for each intervention. (Tables 13 to 17)

Table 12. Summarised strategies for each intervention for women presenting with a self- identifying risk factor (base-case scenario)

Intervention analysed	BMD scanning	Percentage of	Percentage of
	strategies cost-	women age 50 or	women age 50 or
	effective from what	older that would	older that would
	age (years)?	be offered a	be successfully
		BMD scan (%) $^{\nabla}$	treated (%) $\nabla_{\Psi}$
Pooled alendronate and risedronate	50	64.4	19.3
Strontium ranelate	65	44.1	3.5
Raloxifene	None	0.0	0.0
Etidronate	55	58.2	27.5
Teriparatide	70	2.5	0.1

<sup>&</sup>lt;sup>V</sup> These are the BMD Scans and people treated assuming that all women with a prior fracture were opportunistically screened immediately. Once this had been achieved, the numbers will be significantly reduced, assuming that opportunistic assessment regarding clinical risk factors would be undertaken once every 5 years, and that people on treatment would not be re-assessed.

 $<sup>^{\</sup>Psi}$  These numbers have taken persistence into account. Thus, where persistence is 50%, double this number would be offered treatment.

Table 13. The base-case results for pooled alendronate and risedronate in women with a one self-identifying risk factor (same as matrix 'base-case 2')

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
50-54 years	Do not BMD	Do not BMD	BMD and treat	£19,818
			where T-Score	
			<-2.0 SD	
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£14,414
			where T-Score	
			<-2.0 SD	
60-64 years	Do not BMD	BMD and treat	BMD and treat	£15,321
		where T-Score	where T-Score	
		<-2.5 SD	<-2.0 SD	
65 - 69 years	BMD and treat	BMD and treat	BMD and treat	£15,087
	where T-Score	where T-Score	where T-Score	
	<-3.0 SD	<-2.5 SD	<-1.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£10,028
	where T-Score	where T-Score	where T-Score	
	<-2.0 SD	<-1.0 SD	<0.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£6,373
over	where T-Score	where T-Score	where T-Score	
	<-1.5 SD	<-0.5 SD	<0.5 SD	

Table 14. The base-case results for strontium ranelate in women with one self-identifying risk factor.

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
65 - 69 years	Do not BMD	Do not BMD	BMD and treat	£19,623
			where T-Score	
			<-3.0 SD	
70-74 years	Do not BMD	BMD and treat	BMD and treat	£14,816
-		where T-Score	where T-Score	
		<-3.0 SD	<-2.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£12,660
over	where T-Score	where T-Score	where T-Score	
	<-3.5 SD	<-3.0 SD	<-1.5 SD	

Table 15. The base-case results for raloxifene in women with one self- identifying risk factor.

Opportunistic assessment strategies have cost per QALYs of >£20,000 at all ages.

Table 16. The base-case results for etidronate in women with one self-identifying risk factor.

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
55 - 59 years	Do not BMD	Do not BMD	BMD and treat	£16,858
			where T-Score	
			<-1.5 SD	
60-64 years	Do not BMD	Do not BMD	BMD and treat	£16,960
			where T-Score	
			<-1.5 SD	
65 - 69 years	Do not BMD	BMD and treat	BMD and treat	£16,281
		where T-Score	where T-Score	
		<-2.0 SD	<-0.5 SD	
70-74 years	BMD and treat	BMD and treat	BMD and treat	£12,629
	where T-Score	where T-Score	where T-Score	
	<-0.0 SD	<1.0 SD	<1.0 SD	
75 years and	BMD and treat	BMD and treat	BMD and treat	£9,869
over	where T-Score	where T-Score	where T-Score	
	<0.5 SD	<1.0 SD	<1.0 SD	

Table 17. The base-case results for teriparatide in women with one self-identifying risk factor.

Age (years)	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk	CPQ of
	Factor	Factors	Factors	strategy
70-74 years	Do not BMD	Do not BMD	BMD and treat	£14,051
			where T-Score	
			<-3.5 SD	
75 years and	Do not BMD	Do not BMD	BMD and treat	£11,280
over			where T-Score	
			<-3.5 SD	

Illustrative cost per QALY values for each intervention compared with no treatment are given in Table 18 for women of different ages, at a T-Score in the range –2.5 to – 3.0 SD, assuming that they have one self-identifying risk factor. These values include neither the costs of assessment nor the costs of BMD scanning.

Table 18. Cost per QALY values for each intervention compared with no treatment. For women with a T-Score in the range −2. 5 to −3.0 SD and one self-identifying risk factor.

	50	60	70	75
Pooled alendronate	£30,710	£24,465	£11,421	£7,998
and risedronate				
Strontium ranelate	£99,056	£73,776	£33,971	£28,791
Raloxifene	£289,465	£153,113	£48,583	£42,883
Etidronate	£55,787	£31,173	£9,834	£8,039
Teriparatide	£246,896	£201,238	£109,130	£95,397

Table 19 gives the incremental cost-effectiveness of moving from each intervention to pooled alendronate and risedronate. Pooled alendronate and risedronate is considered more cost-effective than both strontium ranelate and raloxifene. However because of the lower price of etidronate a move from etidronate to pooled alendronate and risedronate would not be considered cost-effective as the cost per QALY ratio is greater than £30,000 in the examples provided.

Table 19. Cost per QALY values for pooled alendronate and risedronate compared with each intervention. For women with a T-Score in the range –2.5 to –3.0 SD and one self-identifying risk factor.

	50	60	70	75
Strontium	Pooled	Pooled	Pooled	Pooled
ranelate	alendronate and	alendronate and	alendronate and	alendronate and
	risedronate	risedronate	risedronate	risedronate
	dominates	dominates	dominates	dominates
	strontium	strontium	strontium ranelate	strontium ranelate
	ranelate	ranelate		
Raloxifene	Pooled	Pooled	Pooled	Pooled
	alendronate and	alendronate and	alendronate and	alendronate and
	risedronate	risedronate	risedronate	risedronate
	dominates	dominates	dominates	dominates
	raloxifene	raloxifene	raloxifene	raloxifene
Etidronate	£16,030	£17,084	£19,216	£7,827
Teriparatide *	£2.5 m	£2.4 m	£1.8 m	£1.5 m

<sup>\*</sup> Teriparatide provides more QALYs than pooled alendronate and risedronate, but costs more. In this circumstance cost per QALY ratios greater than £20,000 are desirable.

Although pooled alendronate and risedronate are more cost-effective at a T-Score of – 2.5SD to –3.0 SD, etidronate could be cost-effectively prescribed to women at less

severe T-Score thresholds than pooled alendronate and risedronate. This arises as these women have a relatively higher risk of vertebral fracture (where we have assumed that etidronate has the greater effect) and because etidronate is cheaper than alendronate.

#### 5. Estimating the cost-effectiveness of potential second line interventions

Based on the current guidance for the secondary prevention of osteroporotic fracture<sup>7</sup> and previous appraisal consultation documents<sup>8,9</sup>, it is conceivable that an analysis for second-line interventions is required. For women who have been identified, the number of risk factors summated, a BMD scan performed and begun treatment on alendronate or risedronate but cannot tolerate this intervention, the T-Score threshold at which other interventions become cost-effective has therefore been calculated. In this circumstance no additional assessment or BMD scanning costs are incurred as these costs have already been accounted for, i.e. that the risk factors and BMD of women considered for pooled alendronate and risedronate treatment are already known. In this instance only the cost-effectiveness of treatment itself is relevant.

As an example, in isolation strontium ranelate is cost effective for women aged 70 years with 3 clinical risk factors (none of which were self-identifying). However if strontium was considered as the first line therapy these patients would not be treated as the costs of opportunistically assessing and then providing BMD scans to women with 3 clinical risk factors were prohibitive. Where pooled alendronate and risedronate were assumed first line therapy, women can be cost-effectively assessed and those with 3 clinical risk factors provided with BMD scans. If women with T-scores <-2.5 SD could not tolerate bisphosphonates then strontium ranelate could be cost-effectively initiated.

The T-Score thresholds may differ between women previous identified by opportunistic screening and those presenting with a self-identifying risk factor as the coefficient of increased risk for future fractures is different between clinical risk factors. As such the T-Score thresholds are presented separately for women who were identified by opportunistic assessment and for those with self-identifying risk factors.

For comparative purposes the T-Score threshold at which pooled alendronate and risedronate is considered a cost-effective treatment is provided in italics in the tables.

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<sup>&</sup>lt;sup>7</sup> http://www.nice.org.uk/page.aspx?o=TA087

<sup>8</sup> http://www.nice.org.uk/page.aspx?o=273457

http://www.nice.org.uk/page.aspx?o=273846

## 5.1 T-Score threshold analysis for each intervention for women who had previously been identified by opportunistic assessment.

Using pooled alendronate and risedronate as a first-line treatment, at a cost per QALY of £20,000 it was considered cost-effective to opportunistically assess all women aged 70 years and over and to BMD scan all these women bar those aged 70-74 years and without a clinical risk factor. Using this strategy as a base-case the T-Score thresholds at which women that had previously been identified by opportunistic assessment could be cost effectively treated with an alternative intervention is given in Tables 20 and 21.

The most negative T-Score that was analysed was women in the group −5.0 to −5.5SD since very few women have T-Scores more severe than this. Where the cost per QALY of an intervention was greater than £20,000 for women with T-Scores of -4.75 to -5.25SD, the phrase "Cost per QALY >£20,000 for all T-Scores" has been used in the tables.

Table 20. Treatment thresholds for which each intervention can be considered cost-effective in women aged 70-74 years of age who had previously been identified by opportunistic assessment.

	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk
	Factor	Factors	Factors
Pooled	Treat where T-	Treat where T-	Treat where T-
alendronate	<i>Score</i> <-2.5	Score <1.0 SD	<i>Score</i> <-0.5
and	SD		SD
risedronate			
Strontium	Treat where T-	Treat where T-	Treat where T-
ranelate	Score <-4.0	Score <-3.0 SD	Score <-2.5 SD
	SD		
Raloxifene	Cost per	Treat where T-	Treat where T-
	QALY	Score <-5.0 SD	Score <-4.5 SD
	>£20,000 for		
	all T-Scores		
Etidronate	Treat where T-	Treat where T-	Treat where T-
	Score <-1.5	Score <-0.5 SD	Score < 1.0 SD
	SD		
Teriparatide	Treat where T-	Treat where T-	Treat where T-
	Score <-5.0	Score <-4.5 SD	Score <-3.5 SD
	SD		

Table 21. Treatment thresholds for which each intervention can be considered cost-effective in women aged 75-79 years of age who had previously been identified by opportunistic assessment.

	0 Clinical Risk	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk
	Factors	Factor	Factors	Factors
Pooled	Treat where T-	Treat where T-	Treat where T-	Treat where T-
alendronate	<i>Score</i> <-2.5	<i>Score</i> <-2.0	<i>Score</i> <-0.5	Score < 0.5 SD
and	SD	SD	SD	
risedronate				
Strontium	Treat where T-	Treat where T-	Treat where T-	Treat where T-
ranelate	Score <-4.0	Score <-3.5	Score <-2.5 SD	Score <-2.0 SD
	SD	SD		
Raloxifene	Cost per	Cost per	Treat where T-	Treat where T-
	QALY	QALY	Score <-5.0 SD	Score <-4.0 SD
	>£20,000 for	>£20,000 for		
	all T-Scores	all T-Scores		
Etidronate	Treat where T-	Treat where T-	Treat where T-	Treat where T-
	Score <-1.5	Score <-1.0	Score < 0.0 SD	Score <1.0 SD
	SD	SD		
Teriparatide	Cost per	Treat where T-	Treat where T-	Treat where T-
	QALY	Score <-5.0	Score <-4.0 SD	Score <-3.0 SD
	>£20,000 for	SD		
	all T-Scores			

## <u>5.2 T-Score threshold analysis for each intervention for women who had</u> previously presented with a self-identifying risk factor.

Assuming pooled alendronate and risedronate as a first-line treatment, at a cost per QALY threshold of £20,000 it was considered cost-effective to selectively BMD scan all women aged 55 years and over. As age increased the number of clinical risk factors required to receive a BMD scan decreased. Using this strategy as a base-case the T-Score thresholds at which women that had previously been identified by opportunistic assessment could be cost effectively treated with an alternative intervention is given in Tables 22 and 26.

Table 22. Treatment thresholds for which each intervention can be considered cost-effective in women aged 50-54 years of age who had previously presented with a self-identifying risk factor.

	3 Clinical Risk
	Factors
Pooled alendronate and	Treat where T-Score
risedronate	<-2.0 SD
Strontium ranelate	Treat where T-Score
	<-3.5 SD
Raloxifene	Cost per QALY
	>£20,000 for all T-
	Scores
Etidronate	Treat where T-Score
	<-3.5 SD
Teriparatide	Treat where T-Score
	<-4.0 SD

Table 23. Treatment thresholds for which each intervention can be considered cost-effective in women aged 55-59 years of age who had previously presented with a self-identifying risk factor.

	3 Clinical Risk
	Factors
Pooled alendronate and	Treat where T-Score
risedronate	<-2.0 SD
Strontium ranelate	Treat where T-Score
	<-3.5 SD
Raloxifene	Cost per QALY
	>£20,000 for all T-
	Scores
Etidronate	Treat where T-Score
	<-1.5 SD
Teriparatide	Treat where T-Score
_	<-4.0 SD

Table 24. Treatment thresholds for which each intervention can be considered cost-effective in women aged 60-64 years of age who had previously presented with a self-identifying risk factor.

	2 Clinical Risk	3 Clinical Risk
	Factors	Factors
Pooled alendronate and	Treat where T-Score	Treat where T-Score
risedronate	<-2.5 SD	<-2.0 SD
Strontium ranelate	Treat where T-Score	Treat where T-Score
	<-4.0 SD	<-3.5 SD
Raloxifene	Cost per QALY	Cost per QALY
	>£20,000 for all T-	>£20,000 for all T-
	Scores	Scores
Etidronate	Treat where T-Score	Treat where T-Score
	<-3.0 SD	<-1.5 SD
Teriparatide	Treat where T-Score	Treat where T-Score
	<-4.5 SD	<-4.0 SD

Table 25. Treatment thresholds for which each intervention can be considered cost-effective in women aged 65-69 years of age who had previously presented with a self-identifying risk factor.

	1 Clinical Risk	2 Clinical Risk	3 Clinical Risk
	Factors	Factors	Factors
Pooled	Treat where T-Score	Treat where T-Score	Treat where T-Score
alendronate and	<-3.0 SD	<-2.5 SD	<-1.5 SD
risedronate			
Strontium	Treat where T-Score	Treat where T-Score	Treat where T-Score
ranelate	<-4.5 SD	<-4.0 SD	<-3.0 SD
Raloxifene	Cost per QALY	Cost per QALY	Cost per QALY
	>£20,000 for all T-	>£20,000 for all T-	>£20,000 for all T-
	Scores	Scores	Scores
Etidronate	Treat where T-Score	Treat where T-Score	Treat where T-Score
	<-2.0 SD	<-2.0 SD	<-0.5 SD
Teriparatide	Treat where T-Score	Treat where T-Score	Treat where T-Score
	<-5.0 SD	<-5.0 SD	<-4.5 SD

Table 26. Treatment thresholds for which each intervention can be considered cost-effective in women aged 70-74 years of age who had previously presented with a self-identifying risk factor.

	1 Clinical Risk Factor	2 Clinical Risk	3 Clinical Risk
		Factors	Factors
Pooled	Treat where T-Score	Treat where T-Score	Treat where T-Score
alendronate and	<-2.0 SD	<-1.0 SD	<-0.0 SD
risedronate			
Strontium	Treat where T-Score	Treat where T-Score	Treat where T-Score
ranelate	<-3.5 SD	<-3.0 SD	<-2.0 SD
Raloxifene	Treat where T-Score	Treat where T-Score	Treat where T-Score
	<-5.0 SD	<-4.5 SD	<-4.0 SD
Etidronate	Treat where T-Score	Treat where T-Score	Treat where T-Score
	<0.0 SD	<1.0 SD	<1.0 SD
Teriparatide	Treat where T-Score	Treat where T-Score	Treat where T-Score
	<-5.0 SD	<-4.5 SD	<-3.5 SD

Table 27. Treatment thresholds for which each intervention can be considered cost-effective in women aged 75 years of age and older who had previously presented with a self-identifying risk factor.

	1 Clinical Risk Factor	2 Clinical Risk	3 Clinical Risk
		Factors	Factors
Pooled	Treat where T-Score	Treat where T-Score	Treat where T-Score
alendronate and	<-1.5 SD	<-0.5 SD	<-0.5 SD
risedronate			
Strontium	Treat where T-Score	Treat where T-Score	Treat where T-Score
ranelate	<-3.5 SD	<-3.0 SD	<-1.5 SD
Raloxifene	Treat where T-Score	Treat where T-Score	Treat where T-Score
	<-4.5 SD	<-4.5 SD	<-3.5 SD
Etidronate	Treat where T-Score	Treat where T-Score	Treat where T-Score
	<0.5 SD	<1.0 SD	<1.0 SD
Teriparatide	Treat where T-Score	Treat where T-Score	Treat where T-Score
	<-5.0 SD	<-4.5 SD	<-3.5 SD