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Models of care for osteoporosis: A systematic scoping review of efficacy and implementation characteristics

Alicia R Jones^{a,b}, Madhuni Herath^{b,c}, Peter R Ebeling^{b,d}, Helena Teede^{a,b}, Amanda J Vincent^{a,b,*}

^a Monash Centre for Health Research and Implementation, Monash University, Locked Bag 29, Clayton, Vic 3168, Australia

^b Department of Endocrinology, Monash Health, Melbourne, Australia

^c Hudson Institute of Medical Research, Melbourne, Australia

^d Department of Medicine, School of Clinical Sciences, Monash University, Melbourne, Australia

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ABSTRACT

Background: Osteoporosis affects over half of adults over 50 years worldwide. With an ageing population, osteoporosis, fractures and their associated costs are increasing. Unfortunately, despite effective therapies, many with osteoporosis remain undiagnosed and untreated. Models of care (MoC) to improve outcomes include fracture liaison services, screening, education, and exercise programs, however efficacy for these is mixed. The aim of this study is to summarise MoC in osteoporosis and describe implementation characteristics and evidence for improving outcomes.

Methods: This systematic scoping review identified articles via Ovid Medline and Embase, published in English between 01/01/2009 and 15/06/2021, describing MoC for adults aged \geq 18 years with, or at risk of, osteoporosis and / or health professionals caring for this group. All included at least one of clinical, consumer or clinician outcomes, with fractures and bone mineral density (BMD) change the primary clinical outcomes. Exclusion criteria were studies assessing pharmaceuticals or procedures without other interventions, or insufficient operational details. All study designs were included, with no comparator necessary. Title and abstract were reviewed by two reviewers. Full text review and data extraction was performed by these reviewers for 20% of article and, thereafter by a single author. As the review was predominantly descriptive, no comparator statistics were used.

Findings: 314 articles were identified describing 289 MoC with fracture liaison services (n=89) and education programs (n=86) predominating. The population had prior fragility fracture in 77 studies, the median (IQR) patient number was 210 (87, 667) and the median (IQR) follow-up duration for outcome assessment was 12 (6, 12.5) months. Fracture reduction was reported by 65 studies, with 16 (37%) graded as high quality, and 19 / 47 studies with a comparator group found a reduction in fractures. BMD change was reported by 73 studies, with 41 finding improved BMD. Implementation characteristics including reach, fidelity and loss to follow-up were under-reported, and consumer and clinician perspectives rare.

Interpretation: This comprehensive review of MoC for osteoporosis demonstrated inconsistent evidence for improving outcomes despite similar types of models. Future studies should include implementation outcomes, consumer and clinician perspectives, and fracture or BMD outcomes with sufficient duration of follow-up. Authors should consider pragmatic trial designs and co-design with clinicians and consumers.

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1. Introduction

Osteoporosis and low bone mass (osteopenia) is estimated to effect more than 50% of adults aged over 50 years [1,2]. Osteoporosis causes minimal symptoms prior to a fracture, and, in older adults, most fractures are the result of osteoporosis [3,4]. In 2000, 9 million

* Corresponding author at: Monash Centre for Health Research and Implementation, Monash University, Locked Bag 29, Clayton, Vic 3168, Australia. *E-mail address:* Amanda.vincent@monash.edu (A.J. Vincent). osteoporotic fractures occurred worldwide; the lifetime risk of hip fracture for adults aged 50 years is equivalent to the risk of stroke, and the risk of any major osteoporotic fracture is similar to the risk of cardiovascular disease [5,6]. Morbidity and mortality following fracture is substantial, and recent evidence suggests the burden from osteoporotic fractures is greater than many other non-communicable diseases, including chronic obstructive pulmonary disease and stroke [5]. The cost to the healthcare system for fractures is large; among six European countries, expenditure on osteoporotic fractures was \in 37.5 billion in 2017, or up to 6.4% of healthcare expenditure [5].

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Research in context

Evidence before this study

Models of care for improving outcomes for people with, or at risk of, osteoporosis include fracture liaison, screening, education and exercise programs. However, the evidence for improving clinical outcomes is mixed, and there is a paucity of data on the most critical outcome of fracture reduction. We performed a systematic scoping review of models of care for adults with or at risk of osteoporosis, using Ovid Medline and Ovid Embase, of articles published between 01/01/2009-15/06/2021.

Added value of this study

To our knowledge, this is the largest review of models of care in osteoporosis. We have provided a comprehensive summary of published evidence and have used a validated system for classifying models of care, which can be replicated in other studies.

Implications of all the available evidence

We suggest future reports on models of care for osteoporosis consider the study design, and inclusion of an appropriate comparison group, provide longitudinal follow-up to allow assessment of fracture reduction, or consider using bone mineral density changes as a surrogate marker for this, and include details of delivery and implementation characteristics, which may assist in scaling models to other settings. Lastly, we suggest the inclusion of consumer or clinician perspectives, as key to the success of complex interventions.

With an ageing population worldwide, the prevalence of osteoporosis, low bone mass, and osteoporotic fractures is predicted to increase, and by 2040 it is expected that over 300 million people will be at high risk for osteoporotic fracture [7]. Therefore, it is critical that measures are taken to prevent fractures, and ensure that people who suffer a fracture receive appropriate care to prevent recurrent fractures. Unfortunately, a treatment gap exists in osteoporosis, with low screening and treatment rates, and poor adherence to treatment [5,8–10]. Models of care (MoC) can be defined as operationalising how specific care should be delivered to a group of people at a disease, service or systems level [11]. MoC for primary fracture prevention include screening, education initiatives for clinicians and / or consumers, and exercise programs [12-14]. The efficacy of these initiatives is unclear, and may be related to differences in program characteristics, the population studied, and control group used [13,15,16]. The gold standard MoC for secondary prevention of osteoporotic fractures is a fracture liaison service (FLS). An FLS employs a dedicated coordinator to identify, inform and assess all patients with an osteoporotic fracture within a health system. Different FLS have been classified as Type A (identify patients, investigate for secondary causes of osteoporosis and initiate appropriate treatment), Type B (identify and investigate, but refer to primary care physician for treatment), Type C (identify and inform patient and their primary care physician) and Type D (identify and inform the patient only) [17]. Reviews of FLS have shown an improvement in dual energy X-ray absorptiometry (DXA) screening and treatment rates, which vary by the type of FLS model, being highest for the Type A FLS model [16–19]. Whilst increased treatment may be presumed to lead to a reduction in refractures due to the known benefits of antiresorptive therapy, adherence to treatment started in an FLS is variable, ranging between 34 and 95% [17]. Indeed, evidence for fracture reduction using an FLS is unclear, limited by study size, an appropriate control group and duration of follow-up [17]. Recently, changes in bone mineral density

(BMD) has been proposed as a surrogate marker for fractures for therapeutic trials in osteoporosis, and this may also prove useful for more complex interventions such as FLS [20].

A limitation of published research on osteoporosis MoC is failure to include delivery and implementation characteristics. Operational characteristics for delivery include the frequency, duration and method of contact, the setting, and whether participants are seen individually or in a group. Implementation characteristics include factors such as acceptability, uptake, fidelity, cost and sustainability [21]. Studies of osteoporosis MoC can be viewed as hybrid effectiveness-implementation trials, as they use a targeted implementation strategy (such as education or coordination of care) to try to change behaviour (such as medication initiation, DXA screening) and ultimately improve bone health (reduce fractures or increase BMD). Guidelines exist on designing and reporting on implementation trials, and frameworks such as RE-AIM (Reach, Effectiveness, Adoption, Implementation, Maintenance) can be used to assess and compare implementation characteristics in real-world interventions [22–24]. Differences in implementation characteristics may contribute to variable outcomes between similar MoC, and impact the ability to scale up MoC to other settings.

Despite advances in screening and treatment for osteoporosis, a global increase in fractures in the coming years due to populations ageing is predicted, and so implementing effective models of care is essential [5,25,26]. The aims of this review are to: (i) summarise MoC for people with or at risk of osteoporosis; (ii) outline and compare the implementation characteristics of different MoC; and (iii) compare whether different MoC improve a variety of outcomes including reductions in fractures and increases in BMD. We hope this will assist those people planning, implementing and reporting on osteoporosis interventions in the future.

2. Methods

2.1. Search strategy and selection criteria

A scoping review methodology was chosen to enable a broad overview of MoC that have been trialled in osteoporosis, and to describe the evidence for each of these. The scoping review protocol adhered to the Joanna Briggs Institute guidelines for scoping reviews [27]. Inclusion criteria were English language publications, published between 01/01/2009 and 15/06/2021. This date range was chosen to include the most contemporary MoC using currently available technology and therapeutics. All study designs were included. The population was defined as either (i) adults aged \geq 18 years with, or at risk of, low bone mineral density with or without fracture; and / or (ii) any health professional, including allied health. The intervention comprised any MoC for osteoporosis. No comparator was necessary for inclusion. Outcomes needed to include at least one of clinical, consumer or clinician outcomes. The primary clinical outcome was fractures; the secondary clinical outcome was increase in BMD. Other outcomes included consumer (medication use and adherence, calcium supplement use / calcium intake, vitamin D supplement use, DXA rates, osteoporosis knowledge, osteoporosis self-efficacy, osteoporosis health beliefs), clinician (prescribing rates for medications and vitamin D, screening rates for DXA, osteoporosis knowledge), health service satisfaction, implementation characteristics and cost. Implementation characteristics were broadly based on the RE-AIM framework [24]-Reach (the proportion of people who participated in the MoC, of those eligible), Effectiveness (outcomes as mentioned), Adoption (where applicable, the proportion of settings / institutions who participated in the MoC, of those invited), Implementation (fidelity to the intervention) and Maintenance (the longest time point reported was included in results). Exclusion criteria were studies assessing individual or combination pharmaceuticals or procedures without other interventions, or insufficient detail provided to specify operational characteristics of the MoC.

A systematic search, based on the selection criteria and combining MeSH terms and text words, was developed for Ovid Medline and translated to Embase (Supplement 1). Hand searching of included articles' reference lists was also performed. Authors were contacted directly where full-text article could not be retrieved, or to clarify study details. Covidence (www.covidence.org) was used to manage search results, and for abstract and full text review. Two reviewers (AJ, MH) independently reviewed the titles, abstracts and keywords of every article retrieved by the search strategy according to the selection criteria. Full text of the articles were retrieved for further assessment if the information given suggests that the study meets the selection criteria or if there is any doubt regarding eligibility of the article based on the information given in the title and abstract. Full text review and data extraction was performed independently by two reviewers for 20% of articles, to achieve 100% agreement, thereafter performed by a single author (AJ). The study protocol was registered with Joanna Briggs Scoping reviews on 13/11/2019 (Supplement 2), and reporting adhered to the PRISMA-scoping review extension checklist.

2.2. Data analysis

Data extraction was performed in Microsoft Excel 2016. We adapted our data extraction table from the Cochrane Effective Practice and Organisation of Care (EPOC) framework for describing interventions, and a previously published scoping review on low-cost MoC [28,29]. Information collected included general details (title, authors, country, year of publication), participants and number, the MoC implemented, delivery characteristics [28] (contact method, frequency, setting, individual vs group care) and clinical outcomes as mentioned. MoC were categorized according to the Cochrane EPOC taxonomy of delivery arrangements and implementation strategies for health system interventions [30]. We also classified MoC by the primary type of activity, such as fracture liaison services (further classified into Types A to D as per Ganda [17], education, exercise, screening, orthogeriatric services (OGS), or specialist review. Where models

were multi-component, the primary activity was listed, followed by the other types. Where a single model, with the same participants, was described by different papers (Eg. different time points or outcomes), results were summarised together. The longest follow up time point reported was included in result tables. Where studies included a comparison group, p values for between groups, was included in results tables. Due to the number of studies included in our review, risk of bias assessment using the SIGN proforma [31], was performed for papers reporting fractures, our primary clinical outcome, only.

Given the primary aim of this review was descriptive, no comparative statistics were used. Categorical data are described as number (percentage, %). Continuous data are described as mean (standard deviation) where normally distributed, and median (interquartile range, IQR) when non-parametric. Studies were summarised (i) overall, and then by outcomes of (ii) fractures and (iii) BMD change.

2.3. Role of the funding source

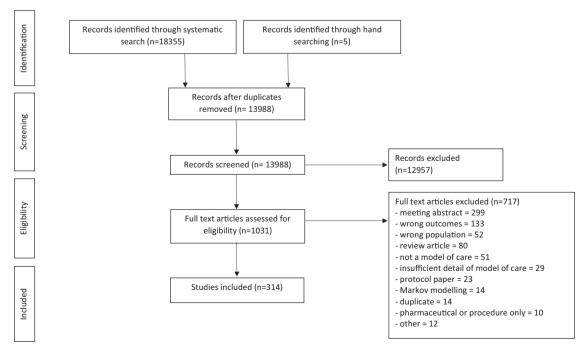
This study received no direct funding. All authors had full access to the data in the study and accept responsibility to submit for publication.

3. Results

3.1. Overall

Fig. 1 and Supplement 3 summarises our search strategy, which resulted in 314 articles included which reported on 289 models of care (25 articles were additional follow-up of the same model and participants). The majority of excluded studies at the title and abstract stage reported only on pharmaceuticals / surgical procedures, and at the full text review stage because they were an abstract only or reported the wrong outcomes.

Summary data for included studies are shown in Table 1, with complete study details shown in Table S1 and implementation characteristics are in Table 2.



Study design n(%)		Randomised trial	117 (40.5
		Non-randomised trial	16 (5.5)
		Cohort study	80 (27.7)
		Case study / series	38 (13.1)
		Pre-test post-test	23 (8.0)
		Other	15 (5.2)
Type of model of care n(%)		Education	86 (29.8)
		Fracture liaison service	89 (30.8)
		Туре А	54 (18.7)
		Туре В	13 (4.5)
		Type C	15 (5.2)
		Type D	4(1.4)
		Combination	3 (1.0)
		Exercise	68 (23·5)
		Screening	18 (6.2)
		Orthogeriatric service	11 (3.8)
		Other	17 (5.9)
Target Population n(%)	Patient (<i>n</i> =290)	Prior fragility fracture (any)	77 (26.6)
		Post-menopausal women	44 (15-2)
		Prior hip fracture	38 (13.1)
		Older adults	30 (10.4)
		Postmenopausal women with low BMD	27 (9.3)
		Known low BMD	18 (6.2)
		Females with cancer	9(3.1)
		Prior radius fracture	7 (2.4)
		Males with prostate cancer	6(2.1)
		Other	33 (11.4)
	Clinician (n=42)	Primary care physicians	23 (54·8)
		Specialist physicians	4(9.5)
		orthopaedic surgeons	4(9.5)
		Junior doctors	4(9.5)
		Other	7(16.7)
Outcomes n(%)*	Patient level	Fractures	65 (22·5)
		BMD	73 (25.3)
		DXA	87 (30.1)
		Treatment (antiresorptive / anabolic)	113 (39.1
		Vitamin D	38 (13-1)
		Calcium intake (supplement+/- diet)	56 (19.4)
		Osteoporosis knowledge	32 (11.1)
		Osteoporosis self-efficacy	14 (4.8)
		Osteoporosis health beliefs	9(3.1)
	Clinician level	Ordering DXA	21 (7.3)
		Prescribing (antiresorptive / anabolic)	48 (16.6)
		Prescribing Vitamin D	7 (2.4)
		Osteoporosis knowledge	1 (0.3)

 Table 1.

 Summary characteristics of included studies.

Footnote: BMD: bone mineral density; DXA: dual energy X-ray absorptiometry; *n>289 and percentages add to >100% as studies may have more than one outcome.

3.2. MoC classification

The majority of studies used the EPOC delivery arrangement 'coordination of care and management of care processes' (n=177, 61.2%, Table 2), 15 studies compared different delivery arrangements and four studies included more than one subcategory. The most common EPOC implementation strategy was 'interventions targeted at specific types of practice, conditions or settings', observed in 198 studies (68.5%), all of which targeted specific conditions, eight studies compared different implementation strategies, and 25 included more than one subcategory. Classifying MoC by activity, the most common MoC was FLS (n=89, 30.8%), of which the majority (n=54) were classified as a Type A (Table 1). The second most common activity was education (*n*=86, 29.8%), of these 52 targeted patients with eight also sending written communication to a clinician, 24 targeted clinicians only, and 27 targeted both patients and clinicians. In addition, 17 studies included an educational component within another MoC. 32 studies were multi-component (included more than one type of MoC), most commonly screening with education (n=8, 2.8%).

3.3. Study characteristics (Tables 1 and S1)

Most studies were from North America (*n*=123, 42.6%) or Europe (*n*=77, 26.6%) (Table S1). Study designs varied with randomised trials predominating (Table 1), however 30 of these did not report the randomisation method used. All studies targeted a patient population. The median (IQR) number of participants was 210 (87, 667), ranging from 13 to 650,000. While 42 studies targeted clinicians, only 14 (33.3%) of studies reported the number of clinicians involved. The median (IQR) number of clinicians was 57 (24, 327), ranging from 5 to 31,459. The median (IQR) follow-up duration for outcome assessment was 12 (6, 12.5) months, and 130 (45%) of studies had follow-up of ≤ 6 months.

3.4. Implementation characteristics (Table 2)

The majority of studies delivered the MoC in a face-to-face format (n=212, 74.4%), in a medical setting (n=163, 61.7%), with 130 (44.8%) of studies using >one method of delivering care and 34 (11.7%) using >one setting for delivery. Program reach was reported by 120

Table 2.

Summary implementation characteristics of included studies.

	Category	Sub-category	n(%) of studi
EPOC Delivery arrangement n(%)*	How and when care delivered	Group vs individual care	10 (3.4)
	Where care is provided	Outreach services	11 (3.9)
		Site of service delivery	23 (7.9)
	Who provides care	Role expansion or task shifting	21 (7.6)
		Self-management	48 (16.6)
	Coordination of care	Care pathways	17 (5.9)
		Case management	2(0.7)
		Communication between providers	20 (6.2)
		Disease management	27 (9.3)
		Integration	1(0.3)
		Packages of care	110 (37.9)
		Teams	4(1.4)
	Information and communication technology	Health information systems	5(1.7)
		The use of information and communication technology	10 (3.4)
		Telemedicine	1(0.3)
POC implementation strategy <i>n</i> (%)*	Targeted at healthcare workers	Audit and feedback	8(2.8)
	raigerea at neartheare workers	Educational materials	15 (5.2)
		Educational meetings	8(2.8)
		Educational outreach visits, or academic detailing	5 (1·7)
		Clinical Practice Guidelines	5(1.7) 5(1.7)
		Inter-professional education	4(1.4)
		Local consensus processes	14(1.4)
		Local opinion leaders	14(4.8) 1(0.3)
		Patient-mediated interventions	. ,
		Reminders	46 (15·9)
			23 (8·0)
	Tourse data and for the strength of the streng	Tailored interventions	1 (0·3)
	Targeted at specific types of practice, conditions or settings	Health conditions	198 (68.5)
Delivery characteristics n(%)	Contact method (<i>n</i> =285)	Face to face	212 (74.4)
		Written	37 (13)
		Telephone	16 (5.6)
		Electronic	15 (5.3)
		Other	5(1.8)
	Frequency of contact (n=227)	Once	78 (34-4)
		More than once but less than 3 monthly	42 (18.5)
		2-3 monthly	11 (4.8)
		< weekly to monthly	8(3.5)
		Weekly	82 (36-1)
		daily	6(2.6)
	Contact location (n=264)	Medical practice / hospital	163 (61.7)
		University / research facility	12 (4.5)
		Community facility	30 (11.4)
		Home	59 (22.3)
	Group vs individual care (n=260)	Individual	195 (75)
	· · · · · · · · · · · · · · · · · · ·	Group	29 (11.2)
		Both	36 (13.8)
mplementation summary statistics	Reach (<i>n</i> =12), mean (SD)	2011	62,8% (23)
inprementation summary statistics	Fidelity $(n=77)$, mean (SD)		75% (19.2)
	Drop-out (n=155), median (IQR) reganisation of care: $*n > 289$ and percentages add to > 1		15.4% (8

Footnote: EPOC: Effective practice and organisation of care; *n>289 and percentages add to >100% as studies may have more than one classification.

(41.4%) studies, fidelity by 77 (26.6%) studies, and loss to follow-up was reported by 155 (53.6%) studies (Table 2). Frequency of care contact varied between models and within the same model (Table S1). Of primary exercise studies, 62 studies included at least weekly (48 \geq 3 times weekly) contacts, and five were daily. Exercise duration was reported for 64 studies, with a mean (SD) of 53.9 (24.1) min. Education study contact frequency varied with 34 once only, 13 more than once but less than three-monthly, six less than weekly up to monthly, 18 weekly and one daily. The duration of each education session was reported for 30 studies, with a median (IQR) of 52.5 (26.3, 60) min.

3.5. Study outcomes (Table S1)

Overall, 156 (52·2%) of studies reported a significant improvement in one or more of their outcomes (Table S1). The most common outcomes reported were specific osteoporosis treatment rates (antiresorptive / anabolic agents, n=113, 39·1%), followed by DXA rates (*n*=87, 30.1%). Provider outcomes, including prescribing and investigation ordering, were assessed in only 58 (20.1%) studies, of which 18/48 (37.5%) studies reported a significant increase in prescribing rates. Of the MoC reporting treatment rates, the majority used the EPOC delivery arrangement 'coordination of care' (n=80, 70.8%), followed by 'who provides care' (*n*=30, 26.5 %), with the most common subcategory being 'packages of care' (n=47, 41.6%) (Table S1). The most common implementation strategy was 'targeted at healthcare workers' (n=59, 52.2%), followed by 'targeted at a disease' (n=57, 50.4%). Only 38 (33.6%) studies found a significant increase in rates of treatment, including 30 studies classified as 'coordination of care', and using the implementation strategy of 'targeting a disease' in 20 and 'targeting healthcare workers' in 19. Of the MoC reporting DXA rates, most were classified as 'coordination of care' (n=63, 72.4%), followed by 'who provides care' (*n*=25, 28.7%), with the most common subcategory of 'packages of care' (n=40, 46%). The most common implementation strategy was 'targeted at a specific disease' (n=49, 56.3%), followed by 'targeted at healthcare workers' (n=41, 47.1%). Most studies [45 (51.7%)] found a significant increase in DXA completion rates, including 30 studies classified as 'coordination of care', and using the implementation strategy of 'targeting healthcare workers' in 25 and 'targeting a disease' in 21 (Table S1).

3.6. Fracture outcomes (Tables 3, S2, S4)

Fracture outcomes were reported for 66 (22.8%) MoC, for 31 (47.7%) of these fracture was the primary outcome (Tables 3 and S2). Risk of bias assessment was performed for 43 studies (controlled trials, cohort studies and controlled before and after studies), with only 17 (38.6%) graded as high quality.

47 (72.3%) studies had a comparator group for fracture outcomes, of these, 19 (40.4%) found a significant reduction in fractures (Tables 3 and S4). The majority of studies that found a significant fracture reduction had this as a primary outcome (n=16, 84.2%), however only four (21.1%) studies were graded as high quality. Studies that found a significant fracture reduction had median (IQR) follow-up duration of 24 (15, 36.9) months, median (IQR) patient number of 403 (157, 1830), median (IQR) reach of 41.5% (25.5, 61.4) and median (IQR) loss to follow-up of 27.8% (15.8, 30.7). Of the 28 studies which did not find a significant reduction in fractures, 13 (46.4%) were graded as high quality. These studies had a shorter median (IQR) follow-up duration of 12 (12, 25.8) months, median (IQR) patient number of 724 (305, 4326), median (IQR) reach of 67.7% (40.4, 78.3) and median (IQR) loss to follow-up of 14.4% (5.4, 25).

3.7. BMD outcomes

73 (25.3%) MoC reported BMD outcomes, for 66 (90.4%) of these BMD was the primary outcome (Tables 4 and S3). The majority of these were exercise MoC (n=65, 89.0%). 41 (56.2%) studies found a significant improvement in BMD with the MoC. This significant improvement in BMD was seen at the lumbar spine in 27 studies, femoral neck in 18 studies and total hip in 17 studies. 21 studies found an improvement in BMD at >one region of interest.

Studies that found a significant improvement in BMD had median (IQR) follow-up duration of 12 (6, 18) months, median (IQR) patient number of 70 (39, 140), median (IQR) reach of 80·7% (52·6, 89·1) and median (IQR) loss to follow-up of $13\cdot7\%$ (6·2, 22·1). The setting for delivering care was mostly in the community (*n*=10, 24·4%), medical centre (*n*=8, 19·5%) or research facility (*n*=7, 17·1%). Studies that did not find a significant improvement in BMD had median (IQR) follow-up duration of 12 (5·9, 12) months, median (IQR) patient number of 84 (41, 146), median (IQR) reach of 55·8% (50·4, 70·3) and median (IQR) loss to follow-up of 13% (9·1, 26·2). The setting for delivering care for these studies was mostly in the community (*n*=12, 37·5%) or home (*n*=9, 28·1%).

3.8. Gaps in reporting

Only 20 (6.9%) studies reported on consumer satisfaction, seven (2.4%) reported on clinician satisfaction, and 17 (5.9%) reported on cost. Adverse outcomes were reported by 37 (12.8%) of studies and 29 of these were exercise studies. Of these, 17 studies reported musculoskeletal adverse effects, and 16 reported no adverse effects.

4. Discussion

To our knowledge, this is the largest comprehensive review of both primary and secondary MoC for osteoporosis. The most common MoC for osteoporosis were classified as 'coordination of care', with the subcategory of 'packages of care', and used the implementation strategy of 'targeting a specific disease'. The most common activities are FLS and education. Few studies report on implementation characteristics of the model, such as reach, fidelity, and loss to follow-up, which may limit the ability for the MoC to be adapted to other settings and affect the rigour of the results. The majority of models showed an improvement in their primary outcome, although within each outcome, there were mixed results for similar types of models.

It is critical to recognise that implementation characteristics of MoC can influence outcomes [32]. Yet no previous reviews have assessed delivery and implementation characteristics of MoC for osteoporosis, and studies often omit these key details from publications. For example, a FLS may involve face-to-face, telephone or written contact, and may occur on the hospital ward, in a designated clinic or remotely, and each of these approaches may lead to different results. Furthermore, the ability of staff to screen all eligible patients, uptake of FLS by invited patients, fidelity to standardised investigations, and dropout rates, will influence the efficacy of the program. Less than half of included studies reported the reach of the MoC or fidelity to the program, and only half reported loss to follow-up. Where studies have high dropout rates or low reach or fidelity, consumer and clinician feedback may help to explain reasons for this, including the acceptability of the MoC, burden or perceived lack of efficacy, however this was rarely reported by studies. Co-design is now considered standard practice for developing MoC, and consumer and clinician perspectives should be included routinely when reporting MoC [33,34].

We are not the first group to attempt to summarise clinical outcomes of MoC for osteoporosis. Three recent systematic reviews analysed DXA and treatment rates among adults at risk of, or with prior, fragility fracture [15,16,35]. Two included only randomised controlled trials, while one also included quasi-experimental studies with a control group. All used different classification systems for MoC, with one classifying by activities (such as screening, education, feedback) [15], one broadly grouping MoC (FLS, case management, orthopaedic / fracture clinic) [16], and one classifying as structural, healthcare provider- or patient-focussed [35]. Results were mixed. While one study found a significant increase in treatment and DXA rates in a pooled analysis of all types of models [15], another found this benefit for structural and patient-focussed interventions [35], and another only found evidence for benefit in the population who had a prior fracture [16]. In a sub-analysis of studies including only people without prior fracture, the only intervention with benefit was self-scheduling of DXA with education, which increased DXA rates [16]. Several previous reviews have also focussed only on secondary prevention after a fracture [17–19]. One review included only RCTs, while others included additional study types. Again, different classification systems were used to group MoC, with one study not grouping models at all, one classifying models of care as FLS Types A-D, and the other classifying models based on the presence or absence of dedicated personnel, whether BMD was ordered or treatment initiated within model, and whether the model was "intensive" (both of the former criteria) [17–19]. These reviews suggested improvement in treatment rates overall, with a trend towards increased efficacy for more intensive MoC, while results for increased DXA rates were mixed. These mixed results between reviews may relate to inclusion criteria, differences in classifying models of care or implementation characteristics not reported in these reviews. We have attempted to use a validated system for classifying models of care, that can be replicated by other studies, and to include detail on implementation characteristics which may explain differences between trial results.

Although treatment rates are an important outcome for MoC for osteoporosis, it is important to understand that not all patients in primary prevention studies require treatment. The proportion who require treatment will depend on the population and risk of re-fracture, and the success of this treatment depends on patient adherence [36]. Fracture outcomes have been included in two previous reviews, one focussed on secondary prevention after fracture, and the other including both primary and secondary prevention [15,17]. One study including only RCTs performed a meta-analysis of 10 studies, which

Table 3. Summary of studies reporting significant reduction in fractures.

Author (year)	Study design	Type of MoC	Population and sample size (n)	Follow-up months		Delivery of	MoC		EPOC	taxonomy	Clini	cal outcomes	Program reach and loss	Risk of Bias
					Frequency of contact	Contact method	Contact location	Group vs individual care	Delivery arrangement	Implementation strategy	Primary outcome?	Fracture outcomes	to follow-up	
FLS Amphansap (2016)[37] Thailand	Cohort study	FLS type A	>50 yr inpatient with MTF 75	12	More than once, but less than 3monthly	Face to face	Hospital Home	Individual	Packages of care	Targeted at specific health conditions	Fracture	0 (0%) MTF vs 36 (30%) in prior cohort, p<0-001	Reach: not reported Loss to follow-up: intervention 15-7%; control not	+
Bachour (2017)[38] Lebanon	Cohort study	FLS type A	>50 yr ED patient with MTF 250	24	Not reported	Face to face	Hospital	Individual	Packages of care	Targeted at specific health conditions	Fracture	8 (8·2%) total frac- tures vs 18 (18%) in prior cohort, <i>p=</i> 0·004	reported Reach: not reported Loss to follow-up: Intervention 81.7%; Control 23.1%	+
Davidson (2017) [39] Australia	Cohort study	FLS type C	>45 yr inpatient with MTF 140	36	Once	Not reported	Not reported	Individual	Communication between providers	Educational materi- als; Patient-medi- ated interventions	Investigation and treatment	34 (10.5%) MTF vs 25 (19.1) in prior cohort, $p < 0.05$ 13 (8.3%) hip frac- tures vs 16 (23.2%) in prior cohort, p < 0.01	Not reported	+
Huntjens (2011) [40] Netherlands	Cohort study	FLS type A	≥50 yr outpatient or ED patient with non- VF 3255	26	More than once, but less than 3monthly	Face to face	Hospital	Individual	Packages of care	Targeted at specific health conditions	Fracture	89 (6·7%) total frac- tures vs 191 (9·9%) in prior cohort, p=0·001	Reach: 68-4% Loss to follow-up: not reported	+
Inderjeeth (2018) [41] Australia	Cohort study	FLS type A	≥50 yr ED patient with MTF 339	12	Not reported	Face to face	Hospital <i>Home</i>	Individual	Packages of care	Targeted at specific health conditions	Fracture	MTF 17 (8·1%) vs 17 (18·3%) in prior cohort and 8 (17·3%) in usual care, <i>p</i> <0.05 vs prior cohort only	Reach: 64·1% Loss to follow-up: Intervention 16·2%; Usual care 18·2%; Prior cohort 12·4%	++
Lih (2011)[42]	Cohort study	FLS type A	≥45 yr outpatient with non-VF 403	48	More than once, but less than 3monthly	Face to face	Hospital	Individual	Packages of care	Targeted at specific health conditions	Fracture	10 (4.1%) MTF vs 31 (19.7%) in usual care, p < 0.01 1 (0.4%) hip fracture vs 8 (5.1%) in usual care	Reach: 41.5% Loss to follow-up: Intervention 14.6%; Usual care 36.2%	0
Nakayama (2016) [43] Australia	Cohort study	FLS type A	≥50 yr ED patient with MTF 931	36	Not reported	Face to face	Hospital	Individual	Packages of care	Targeted at specific health conditions	Fracture	63 (12·2%) total fractures vs 70 (16·8%) in usual care, <i>p=</i> 0·025	Reach: 20% Loss to follow-up: not reported	+
Van der Kallen (2014)[44] Australia	Cohort study	FLS type A	≥50 yr ED patient with MTF 434	12	More than once, but less than 3monthly	Face to face Telephone	Hospital Home	Individual	Packages of care	Targeted at specific health conditions	Fracture	11 (6-5%) total frac- tures vs 36 (18-6%) in usual care, p<0.001 3 (1.4%) VF vs 4 (1.6%) in usual care	Reach 14% Loss to follow-up: Intervention 27·2%; Usual care 45·5%	+
Wasfie (2019)[45] United States	Cohort study	FLS type A	≥50yr outpatient with VF treated sur- gically 365	26	2-3 monthly	Face to face	Hospital	Individual	Packages of care	Targeted at specific health conditions	Fracture	78 (37%) total frac- tures vs 84 (56%) in prior cohort, <i>p</i>=0.01 46 (22%) VF vs 47 (31%) in prior cohort, <i>p</i>=0.29	Not reported	0
Education Becker (2011)[46], Heinrich (2013) [47] Germany	Controlled before after	Education – patient & cli- nician Exercise	≥65yr in nursing home Clinicians: not reported Patients: 45321	12	Education: not reported Exercise: 60min 2x per wk for 52wk	Face to face Written Video	Home	Group	Disease management	Local opinion leaders	Fracture	331 (2·4%) hip frac- tures vs 917 (2·9%) in usual care, p<0·05	Not reported	+
Pekkarinen (2013) [48] Finland	Non-rando- mised study	Education – patient	60–70 yr post-men- opausal women 2178	120	150min 5x per wk for 1 wk	Face to face Written	Medical Centre	Both	Self-management	Targeted at specific health conditions	Fracture	59 (5-9%) MTF vs 95 (8-1%) in usual care, p=0-045 12 (1-2%) hip	Reach: 39-4% Loss to follow-up: Intervention 28-7%; Control 37-6%	-

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Author (year)	Study design	Type of MoC	Population and sample size (n)	Follow-up months		Delivery of	f MoC		EPOC t	axonomy	Cli	nical outcomes	Program reach and loss
					Frequency of contact	Contact method	Contact location	Group vs individual care	Delivery arrangement	Implementation strategy	Primary outcome?	Fracture outcomes	to follow-up
Sorbi (2016)[49] Iran	Cohort study	Education - clinician	Orthopedic surgeons ≥60 yr inpatient with MTF Clinicians: 30 Patients: 515	24	15min 2x per wk for 13 wk	Face to face	Hospital	Group	Disease management	Educational materials	Treatment	fractures vs 29 (2.5%) in usual care, p=0.039 0.8 total fractures per person per year vs 1.6 in previous cohort, $p<0.05$	Not reported
Screening Harness (2012) [50] United States	Cohort study	Screening – DXA	≥65 yr female, ≥70 yr male, or ≥50 yr at risk of OP 524612	72	Not reported	Face to face Written	GP practice	Individual	Disease management	Targeted at specific health conditions	Fracture	2595 (1·5%) DR frac- tures vs 6063 (1·7%) in usual care, p<0·05	Not reported
Parsons (2019) [51], Shepstone (2018)[12] United Kingdom	RCT	Screening – DXA, FRAX	524612 70–85 yr women 12483	60	Once	Written	GP practice	Individual	Disease management	Targeted at specific health conditions	Fracture	<i>p</i> <0.03 951 (15-3%) total fractures vs 1002 (16%) in usual care, <i>p</i> =0-183 805 (12-9%) MTF vs 852 (13-6%) in usual care, <i>p</i> =0-178 164 (2-6%) hip frac- tures vs 218 (3-5%) in usual care, <i>p</i> =0-002	Reach: 95.6% Loss to follow-up: Intervention 14.4%; Control 14.8%
Zhumk-hawala (2013)[52] United States	Cohort	Screening – DXA	≥50 yr males w prostate cancer on leuprolide 1482	36	Once	Face to face Written	GP practice	Individual	Disease management	Patient-mediated interventions Reminders	Fracture	18 (1.68%) hip frac- tures vs 17 (4.14%) in usual care, p<0.001	Not reported
Exercise Kemmler (2012, 2014, 2015, 2016, 2016,2017)[53- 58] Germany	Controlled before and after study	Exercise	Post-menopausal women with osteo- penia 137	192	40min 4x per wk for 800 wk	Face to face Written	Home Other not reported	Both	Self-management	Targeted at specific health conditions	Fracture	17 (28-8%) total fractures vs 28 (60-9%) in usual care, p=0-03 13 (22%) MTF vs 24 (52-2%) in usual care, p=0-046	Reach: 53.3% Loss to follow-up: Intervention 31.4%; Control 10.9%
Korpe-lainen (2010)[59] Finland	RCT	Exercise	70–73 yr women with low BMD 160	85 (frac- tures) 72 (BMD)	25min daily	Face to face	Home Other not specified	Both	Group vs individual care	Targeted at specific health conditions	BMD	 cate, <i>p</i>-0 work 17 (20-2%) total fractures vs 23 (30-3%) in usual care, <i>p</i>-0-22 0 hip fractures vs 5 (6-6%) in usual care, <i>p</i>=0.02 1 (1-2%) VF vs 1 (1-3%) in usual care 	Reach: 25.5% Loss to follow-up: Intervention 34.5%; Control 40.8%
OGS Cheung (2018) [60] Hong Kong	Cohort	OGS Specialist review Education – patient Exercise Patient support	≥65 yr w hip frac- ture 153	18	Exercise: 60min weekly Vibration: 20min 3x per wk Education 3- monthly	Face to face	Commu- nity Hospital	Both	Disease management	Targeted at specific health conditions	Fracture	1 (1-3%) total frac- tures vs 8 (10-4%) in usual care, <i>p</i> =0-034	Reach: not reported Loss to follow-up: Intervention 28-3%; Control 25-2%

Footnote: p values are between groups unless otherwise specified. Risk of bias: ++ (high quality), + (acceptable), - (low quality), 0 (unacceptable). MoC: model of care; EPOC: effective practice and organisation of care; FLS: fracture liaison service; yr: year; MTF: minimal trauma fracture; ED: emergency department; VF: vertebral fracture; min: minutes; wk: week; DXA: dual energy X-ray absorptiometry; OP: osteoporosis; GP: general practitioner; DR: distal radius; RCT: randomised controlled trial; BMD: bone mineral density; OGS: orthogeriatric service.

Hospital

Individual

Disease

management

Targeted at specific

health condition

Fractures

Face to face

++

++

n/a

8-6% total fractures,

p<0.001 vs baseline

Reach: not

Loss to follow-up:

reported

10.9%

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Specialist review Gomez

(2019)[61]

Australia

Pre-test

post-test

study

Specialist

review

 ${\geq}65$ yr referred to

falls and fracture

clinic

106

Once

6

Risk of Bias

0

Table 4. Summary of studies reporting significant improvement in BMD.

Author (year)	Study design	Type of MoC	Population and sample size (n)	Follow- up months		Delive	ry of MoC		EPO	IC taxonomy	Clinic	al outcomes	Program reach and loss to follow-up
				montais	Frequency of contact	Contact method	Contact location	Group vs individual care	Delivery arrangement	Implementation strategy	Primary outcome?	BMD change	10110W-up
FLS Chandran (2013)[62] Singapore	Case study	FLS type A	≥50 yr inpatient, outpatient or ED patient with MTF 287	24	More than once, but less than 3monthly	Face to face Telephone	Hospital Home	Individual	Packages of care	Targeted at specific health conditions	Treatment	LS: +4-4%, p <0·01 vs baseline TH +2·7%, p<0·01 vs baseline	Not reported
Eekman 2014) <mark>[63]</mark> Netherlands	Case study	FLS type A	≥50 yr ED patient with MTF 1116	12	2-3 monthly	Face to face Telephone	Hospital Home	Individual	Packages of care	Targeted at specific health conditions	Reasons for not attending FLS and adherence	LS: +3·9%, p<0·001 vs baseline TH: +2·3%, p<0·001 vs baseline	Reach: 50.6% Loss to follow up: 74.9%
Education Hien 2009)[64] Jietnam	Non-rando- mised trial	Education – patient	Postmenopausal women with low calcium intake 140	18	Daily	Face to face Written Video	Home Community	Both	Packages of care	Targeted at specific health condition	Calcium intake	Calcaneal*: 0%; con- trol -0.5%, p<0.05 *calcaneal US	Reach not reported Loss to follow- up: Interven- tion 18-6%; Control 31-7%
Wang (2016)[65] China	RCT	Education — patient Exercise Patient support	Known OP 436	48	Monthly	Face to face Written	Community	Both	Packages of care	Targeted at specific health condition	Multiple out- comes includ- ing BMD	Females: LS: +10.4% vs control +2.19%, p<0.01 FN: +14.1% vs con- trol +2.7%, p<0.01 Males: LS: +10.5% vs control +1.06%, p<0.01 FN: +11.1% vs con- trol +1.14%, p<0.01	Reach: not reported Loss to follow- up: Interven- tion 6.4%; Con- trol 13-8%
xercise boarrage (2018)[66] razil	RCT	Exercise	Postmenopausal women 25	6	30 min 3x per wk for 24 wk	Not reported	Community	Not reported	Site of service delivery	Targeted at specific health condition	BMD	LS +3·7% vs control +0·88%, p<0·01 TF +6·5% vs control -1·38%, p<0·01	Reach: not reported Loss to follow up: 0%
Mayat (2018)[67] audi Arabia	RCT	Exercise Laser Group 1 laser Group 2 exercise Group 3 laser & exercise	Men with low BMD	12	20 min exercise ± 18min laser 3x per wk for 24 wk	Face to face	Not reported	Not reported	Packages of care	Targeted at specific health condition	BMD	1: 50%, p ≤ 01 1: 5: Group 1 - 1%, Group 2 + 10.1%, Group 3 + 13% vs control - 1: 5%, p < 0.001 Group 3 vs control TH: Group 1 0%; Group 2 + 3.3%; Group 3 + 2.2% vs control - 1:1%, p < 0.001 Group 3 vs control or Group 1	Reach: not reported Loss to follow up: Group 1 16%; Group 2 12%; Group 3 12%; Control 20%
Almstedt (2016)[68] Jnited States	Pre-test post-test	Exercise	Female cancer survi- vors 26	7	60 min 3x per wk for 26 wk	Face to face	University	Not reported	Packages of care	Targeted at specific health condition	BMD	LS +2.5% vs base- line, p=0.012 TH +1.7% vs base- line, p=0.048	Reach: not reported Loss to follow up: 23.1%
ngin (2015)[69] urkey	RCT	Exercise	Post-menopausal women with low BMD 44	6	60 min 3x per wk for 24 wk	Face to face	Not reported	Group	Group vs indi- vidual care	Targeted at specific health condition	BMD	LS +6-5% vs control- 3-3%, p<0-001	Reach: not reported Loss to follow up: not reported
Astorino (2013)[70] Jnited States	Pre-test post-test	Exercise	Spinal cord injury 13	6	150 min 2x per wk for 26 wk	Face to face	Rehab centre	Individual	Packages of care	Targeted at specific health condition	BMD	LS: +4-7% vs base- line, p<0.05 TH: -7% vs baseline, p<0.05 EN0% vs baseline	Reach: not reported Loss to follow up: 23.1%

FN -4% vs baseline, p<0.05

(continued on next page)

Upper barrier totalContatContati LaborationContati LaborationDelorgion <th>Author (year)</th> <th>Study design</th> <th>Type of MoC</th> <th>Population and sample size (n)</th> <th>Follow- up</th> <th></th> <th>Delive</th> <th>Delivery of MoC</th> <th></th> <th>EPO</th> <th>EPOC taxonomy</th> <th>G</th> <th>Clinical outcomes</th> <th>Program reach and loss to</th>	Author (year)	Study design	Type of MoC	Population and sample size (n)	Follow- up		Delive	Delivery of MoC		EPO	EPOC taxonomy	G	Clinical outcomes	Program reach and loss to
101000					months	Frequency of contact	Contact method	Contact location	Group vs individual care	Delivery arrangement	Im plementation strategy	Primary outcome?	BMD change	follow-up
Rt Expension Expen	at (2013)[71] key	RCT	Exercise Group 1 strength exercise Group 2 high-impact exercise	Postmenopausal women with low BMD 42	ى	60 min 3x per wk for 26 wk	Face to face	Hospital	Not reported	Packages of care	Targeted at specific health condition	BMD	LS. Group 1 +1.3%. Group 2 +0.5% vs control-2.5%. p=006 Group 2 vs control FN. Group 1 +1.6%. Group 2 1 22 vs control -1%. p=0.06 Group 2 vs	Reach: not reported Loss to follow- 14%: Group 2 14.3%: Control 14.3%:
1KC1ExerciseEx	cers (2014)[72] ed States	RCT	Exercise Education – Gatient Croup 1: Diet plan Group 2: Exercise Group 3: Diet plan & exercise	≥55 yr. BMI 27-40 and osteoarthritis of knees 392	ő	Exercise: 60 min 3x per Education: 1-2 weekly	Face to face	Community Home	Both	Packages of care	Targeted at specific health condition	BMD	Control 1-0.3% Group 2-0.5% vs Group 2-0.5% vs Group 2-0.4% P=0.47 TH: Group 1-2.5% Group 2-0.2% vs Group 2-0.2% vs Group 2-0.2% vs Group 2-0.2% vs Group 3-1.8% P<0.01 Group 1 vs	Reach: 86-3% Loss to follow- up: Group 1 31%, Group 2 26-4%, Group 2 25%
It Decretes Description Decretes Description Lage of acrivity of activity of act	strom (2012)[73] den	RCT	Exercise	Postmenopausal women with low BMD and DR fracture 112	12	40 min 3.4x wk for 52 wk	Face to face	Community	Not reported	Site of service delivery	Targeted at specific health condition	BMD	Group 3 TH: +0.7% vs control -0.9%, p=0.04	Reach: not reported Loss to follow- up: Interven- tion 20%; Con-
RCT Exercise anomeniation anomeniatio anomeniatio anomeniatio anomeniation anomeniation anomeniation a	lini (2009)[74] li	RCT	Exercise	Postmenopausal women 35	٥	60 min weekly for 24 wk	Face to face	Community	Not reported	Site of service delivery	Targeted at specific health condition	BMD	LS0.1% vs control -1%, <i>p</i> < 0.05 FN: -0.14% vs con- trol -1.6%, <i>p</i> < 0.05	Reach: not reported Loss to follow- up: Interven- tion 13%; Con- trol 16.7%
Pinctio (2016) RCT Exercise towner with low per with per with per with Postmenopausal solut 13 60 min for solut Targeted at specific solut B/D Absolute of representation 2009/71 Eventise per with Bvoner with low per with Postmen with low per with Bvoner with low per with Postmen with low per with Bvoner with low per with Postmen with Post	on (2012) [75] ralia	RCT	Exercise	Postmenopausal women with low BMD 39	12	60 min 3x per wk for 52 wk	Face to face	Community	Not reported	Site of service delivery	Targeted at specific health condition	BMD	LS: -0.3% vs control -0.9% p>0.05 TH: +0.5% vs control -0.9%, p=0.02	Reach : 34.5% Loss to follow- up: Interven- tion 0%; Control
RTExercisePostmenopausal6Exercise:Face to faceNot reportedPackages ofTargeted at specificBMDLS GouptionAntioxidantswomen34wh for 26 wk60 min 3x per60 min 3x perAntioxidantsantioxidantswomen61 min 3x per61 min 3x per61 min 3x per60 min 3x per60 min 3x perAntioxidantsantioxidantswomen61 min 3x perwt for 26 wk61 min 3x per61 min 3x per61 min 3x perAntioxidantsantioxidantsantioxidantsantioxidantsantioxidants61 min 3x per61 min 3x per61 min 3x perGroup 2:froup 3:froup 3:antioxidantsantioxidantsantioxidants61 min 3x per61 min 3x perAntioxidantsfroup 3:froup 3:froup 3:froup 3:froup 3:61 min 3x per61 min 3x per61 min 3x perRTExercise50 yr18free for 80 min 3x perfree for 80 min 3x perfree for 92 min 3x per61 min 3x per61 min 3x perEduction-freerise50 yr18free for 92 min 3x perfree for 92 min 3x per61 min 3x per61 min 3x perEduction-freerise50 yr18free for 92 min 3x perfree for 92 min 3x per61 min 3x per61 min 3x perEduction-freerise50 min 3x perfree for 78 min 3x perfree 10 min 3x per61 min 3x per61 min 3x perFer50 m	.a- Pinheiro (2016) il	RCT	Exercise Group 1: 3x per wk Group 2: 2x per wk	Postmenopausal women with low BMD 60	13	60 min for 56wk	Face to face	Not reported	Not reported	Packages of care	Targeted at specific health condition	BMD	Absolute change not reported. <i>p</i> <0.05 Group 1 vs control for TH, FN <i>p</i> <0.05 Group 1 vs Group 2 for LS, TH	Reach: 96-8% Loss to follow- up: Group 1 0%; Group 2 20%; Control 20%
Exercise ≥60 yr 18 60 min 3x per Face to face Community Both Packages of Tangeted at specific BMD L5:+1-48% Education – wk for 78 wk Telephone care health conditions +0.76%, p=0	ada	RCT	Exercise Antioxidants Group 1: antioxidants Group 2: exercise Group 3: & exercise	Postmenopausal women 34	ى	Exercise: 60 min 3x per wk for 26 wk	Face to face	Not reported	Not reported	Packages of care	Targeted at specific health condition	BMD	La Group 10-1%; group 2-0-1%; Group 3-0-3% vs control -1-5% p<0.05 all groups vs control -1-9%; FN: Group 1-0-9%; Group 2-0-3%; con- rol 40 - 27 - 40 005	Reach: not reported Loss to follow- up: not reported
		RCT	Exercise Education –	≥60 yr	18	60 min 3x per wk for 78 wk	Face to face Telephone	Community	Both	Packages of care	Targeted at specific health conditions	BMD	LS: +1.49% vs control +0.76%, p=0.125	Reach: not specified

Author (year)	Study design	Type of MoC	Population and	Follow-		Deliver	Delivery of MoC		EPO	EPOC taxonomy	Cli	Clinical outcomes	Program reach
			sample size (n)	up months									and loss to follow-up
					Frequency of contact	Contact method	Contact location	Group vs individual care	Delivery arrangement	Implementation strategy	Primary outcome?	BMD change	-
Daly (2019)[78], Gia- noudis (2014)[79] Australia		patient & cli- nician Patient	Clinicians: not reported Patients: 162			Written						TH: $+0.61\%$ vs con- trol $+0.32\%$, $p > 0.05$ FN: $+0.6\%$ vs control $+0.22\%$ $\rightarrow 0.001$	Loss to follow- up: Interven- tion 4.9%; Con-
deMatos (2009)[80] Portugal	Non-rando- mised trial	Exercise	Postmenopausal women with low BMD 59	12	45 min, fre- quency not reported, for 52 wk	Not reported	Not reported	Not reported	Packages of care	Targeted at specific health condition	BMD	-1-3.5%, p<-0-01 LS +1.17% vs control -2.26% p<-0-013 TH: -0-71%; control -0-6%, p>0-05	uor 12.3% Reach: not reported Loss to follow- up: not
El-Kader (2016) 81] Saudi Arabia	RCT	Exercise	COPD on inhaled glucocorticoids 60	Q	30 min 3x per wk for 26 wk	Not reported	Not reported	Not reported	Packages of care	Targeted at specific health condition	BMD	LS: +20.8% vs con- trol -1.1%, <i>p</i> <0.04 DR: +25.8% vs con- trol -1.71%, <i>p</i> <0.05	reported Reach: not reported Loss to follow- up: not
Ekisi (2015)[82] Egypt	RCI	Exercise Group 1: exercise Group 2: electromag- netic field	Postmenopausal women, sedentary 30	m	30 min (elec- tromagnetic field) or 60min (exercise) 3x per wk for 12 wk	Face to face	Hospital	Not reported	Packages of care	Targeted at specific health condition	BMD	LS: exercise 2.18% vs electromagnetic field +29%, p<0.01 FN: exercise +2.7% vs electromagnetic field +16.9%,	reported Reach: not reported Loss to follow- up: 0%
Garcia-Gomariz(2018) [83] Spain	RCT	Exercise	Postmenopausal women 36	24	60 min 2x per wk for 92 wk	Face to face	Hospital	Not reported	Packages of care	Targeted at specific health condition	BMD	p=0.012 LS: +24.3% vs control +14.7% p=0.4 FN: +36.8%; vs con- trol -4.7% p<0.05	Reach: not reported Loss to follow- up: Interven- tion 5-6%; Con-
Hojan (2013, 2013)[84]-[85]	Pre-test post-test	Exercise Phase 1 (control): no exercise Phase 2: aer- obic exercise Phase 3: resistance exercise	Pre-menopausal women with breast cancer receiving endocrine therapy 41	°.	45 min daily for 26 wK for each phase	Written Face 10 Jace	Ноте	Individual	Self- management	Targeted at specific health condition	BMD	6 month change.: Ls: Phase 2 -3.0%; Phase 2 -3.0%; Phase 2 -3.0% control μ -0.05 μ -0.05 phase 2 vs control μ -0.01 control μ seeline TH: Phase 2 -11%; Phase 2 -10%; Control vs baseline TH: Phase 2 -16%; Phase 2 -10%; Phase	rei 5.6% Reach: not Loss to follow- up: 22.6%
Kemmler (2013)[86] Germany	RCT	Exercise	Postmenopausal women 85	12	60 min 3sx per wk for 52 wk	Face to face	Not reported	Group	Group vs indi- vidual care	Targeted at specific health condition	BMD	TH: -6.8% LS: -0.1% vs control -2%, p=0.002 TH: -0.4% vs control -0.8%, p=0.152	Reach: 81% Loss to follow- up: Interven- tion 163%;
Kemmler (2012, 2014, 2015, 2016, 2016,2017) 53- 58 Germany	Controlled before and after study	Exercise	Post-menopausal women with osteo- penia 137	192	40 min 4x per wk for 800 wk	Face to face Written	Home Other not reported	Both	Self- management	Targeted at specific health conditions	BMD	LS: -1.5% vs control -5.8% p-0.001 TH: -5.7% vs control -9.7% p<0.01 FN: -6.5% vs control -6.5% vs control	Control 20-06 Reach: 53-3% Loss to follow- up: Interven- tion 31-4%; Control 10-9%
Kukuljan (2009, 2011) [87]·[88] Australia	RCT	Exercise Fortified milk Group 1:	Older males 180	18	60 min 3x per wk for78 wk	Face to face	Community	Group	Packages of care	Targeted at specific health condition	BMD	-9-001 Absolute change not reported LS: p<0-01, all	Reach: 98.9% Loss to follow- up: Group 1

(continued on next page)

Program reach and loss to	follow-up	2:2%: Group 2 2:2%: Group 3 2:2%: Control 4:5%	Reach: not reported Loss to follow- up: Group 1 0%; Group 2 42.9%	Reach: not reported Loss to follow- up: Group 2 10%; Group 3 2%; Control 12:5%	Reach: not reported Loss to follow- up: not reported	Reach: 86.6% Loss to follow- up: Group 1 34.8%; Group 2 20.8%; Control 16.7%	Reach: not reported Loss to follow- up: Group 1 70-6%; Group 2 71-4%	Reach: 53.8% Loss to follow- up: Interven- tion 21.9%; Control 22.6%	Reach: not reported Loss to follow- up: Group 1
Clinical outcomes	BMD change	groups increased vs control FN: effect of exer- cise 1.9%, <i>p</i> <0.001	Absolute change not reported p<0.05 Group 2 vs Group 1 at LS and TH	Absolute change not reported LS: p<0.63 all groups improved vs control FN: Group 11.9% higher BMD than control, p=0.01; Group 3.2.3% higher	<i>p</i> < 0.001 LS: +14.9% vs con- trol = 6%, <i>p</i> < 0.001 TH: +5.06% vs con- trol = 8.6%, <i>p</i> =0.026 FN: +10.39% vs con- ent <i>s</i> = 2000.026	Trol 463%, p=0.002 Trol 463%, p=0.002 Group 1 4 16%; Group 2 40 12% to p=0.034 Group 1 vs other groups FN: Group 1 - 1.6% Group 2 -0.46% vs control -0.29%	$\begin{array}{llllllllllllllllllllllllllllllllllll$	p <0.05 LS: -0.52% vs control -1.43% p>0.05 TH: -5% vs control -7.26% p=0.009 FN 44478 vs control	-7.32%, p7.907 LS: Group 1 +5.53% vs Group 2 +3.92%, p<0.001
CI	Primary outcome?		BMD	CIMB	BMD	BMD	НКОСТ	BMD	BMD
EPOC taxonomy	Implementation strategy		Targeted at specific health condition	Targeted at specific health condition	Targeted at specific health condition	Targeted at specific health condition	Targetted at specific health condition	Targeted at specific health condition	Targeted at specific health condition
EPOC	Delivery arrangement		Packages of care	Packages of care	Group vs indi- vidual care	Group vs indi- vidual care	Packages of care	Packages of care	Packages of care
	Group vs individual care		Individual	Not reported	Group	Group	Not reported	Individual	Not reported
Delivery of MoC	Contact location		Home	Not reported	Not reported	University campus	Not reported	Hospital	Not reported
Delivery	Contact method		Face to face	Not reported	Face to face	Face to face	Face to face	Face to face	Face to face
	Frequency of contact		150 min 6x per wk for 24 wk	3 min daily for 52 wk	60 min 3x per wk for 24 wk	60 min 3x per wk for 32 wk	30 min 3x per wk for 52 wk	75 min 3x per wk for 26 wk	35–55min 3x per wk for 43 wk
Follow- up	months		12	12	9	ω	12	Q	10
Population and sample size (n)			Astronauts 35	Postmenopausal women with low BMD 198	Women with low BMD 22	Postmenopausal women 71	Non-ambulatory spi- nal cord injury 20	Bariatric surgery 70	Postmenopausal women with low BMD 64
Type of MoC		Exercise Group 2: Fortified milk Group 3: Exercise & fortified milk	Exercise Alendronate Group 1: exercise Group 2: exercise &	Exercise Group 1: Tai Chi Croup 2: Cal- ciroup 2: Cal- ciroup 3: Tai Croup 3: Tai Croup 3: Tai Chi, Calcium & Vitamin D	Exercise	Exercise Group 1: resistance exercise Group 2: aerobic exercise	Exercise Zoledronic acid Croup 1: exercise acid acid acid exercise exercise	Exercise	Exercise Group 1: land exercise Group 2:
Study design			Controlled before and after study	RC	RCT	RC	لاتا	RCT	RCI
Author (year)			LeBlanc, (2013)[89], Sibonga (2019)[90] United States	Liu (2015)[91] China	Marchese (2012)[92] Italy	Marques (2011)[93] Portugal	Morse (2019) [94] United States	Murai (2019)[95] Brazil	Murtezani (2014)[96] Kosova

Author (year)	Study design	Type of MoC	Population and sample size (n)	Follow- up		Delive	Delivery of MoC		EPOO	EPOC taxonomy	Clini	Clinical outcomes	Program reach and loss to
				months	Frequency of contact	Contact method	Contact location	Group vs individual care	Delivery arrangement	Implementation strategy	Primary outcome?	BMD change	follow-up
Nicholson (2015)[97] Australia	RCT	aquatic exercise Exercise	Postmenopausal women 57	9	50 min 2x per wk for 26 wk	Face to face	Community	Group	Site of service delivery	Targeted at specific health condition	BMD	LS: +1.01% vs con- trol -2.09%, p=0.005 group x time TH: -0.21% vs control -2.99%, p=0.05 FN: +0.11% vs con- FN: +0.11% vs con-	6.1%; Group 2 3.2% Reach: 96.6% Loss to follow- up: Interven- tion 14.3%; Control 10.3%
Saarto (2012) 98] Finland	RG	Exercise	Women with breast cancer 573	12	60 min 3x per wk for 52 wk	Face to face	Ноте	Both	Site of service delivery	Targeted at specific health condition	BMD	trol -1.5% p>0.5 Prencipausal sub- group: LS -1.9%, control -2.3%, p>0.05 FN -0.2% , control -1.4% , p=0.01 Postmenopausal subgroup: LS -1.6%, control -2.1%, p>0.05 FN -1.1%, control	Reach: not reported Loss to follow- up: Interven- tion 7.3%; Con- trol 6.3%
Sen (2020)[99] Turkey	RCF	Exercise Vibration Group 1: High impact exercise Exercise & vibration	40–65 yr postmeno- pausal women with low BMD 49	٥	60 min 3x per wk for 24 wk	Face to face	Research facility	Not reported	Packages of care	Targeted at specific health condition	BMD	-1.1%, p=0.99 LS: Group 1 = 0.7%; Group 2 = 1.9% s control - 1.9%, p=0.005 Group 2 + 3% s control - 1.9%; FN: Group 1 = 1.9%; Group 2 - 45% s control TH: Group 2 - 41.9% s Group 2 + 1.9% s control - 1.3%, p=0.01 Group 2 vs	Reach: Not reported Loss to follow- up: Group 1 15.8%; Group 2 21-1%; Control 10%
Silverman (2009)[100] United States	Non-rando- mised trial	Exercise	Postmenopausal women with BMI 25–40, sedentary 86	9	52 min 3x per wk for 26 wk	Face to face	Community Home	Individual	Packages of care	Targeted at specific health condition	BMD	LS: +0.42% vs control +0.18%, p>0.05 FN: +1.86% vs con- trol -0.87%, p<0.05	Reach: not reported Loss to follow- up: not
Villareal (2017) [101] United States	RCI	Exercise Specialist review Group 1: diet & aerobic exercise & resistance exercise Group 3: diet & croup 3: diet	≥ 65 yr. BMI > 29, sedentary 160	٥	Exercise: 60 min 4x per wk for 26 wk Dietician review: weekly	Face to face	University campus	Both	Packages of care	Targeted at specific health condition	Physical perfor- mance test. Secondary: BMD	LS. Group 1 +0.18%; Group 2 +0.78% Group 3 +0.69% vs portrol +0.88%, portrol +0.88%, TH: Group 1 -2.6%; group 2 -0.57%; Group 3 -1.18, vs p -0.05 Group 1 vs p -0.05 Group 1 vs all other groups	reported reported Loss to follow- Loss for follow- 12.5%, Group 2 12.5%, Control 10%
von Stengel (2011) [102] Germany	RCI	exercise Exercise Vibration Group 1: exercise Group 2: exercise & vibration	Postmenopausal women 151	8	40 min 4x per wk for 78 wk	Face to face	University Home	Both	Packages of care	Targeted at specific health condition	BMD	LS: Group 1 +2.05%; Group 2 +1.49% vs control 10-42%, <i>p</i> =0.05 Group 1 vs Control 1 v TH: Group 1 +0.12%; group 2 +0.12% vs control -0.12%, control -0.12%,	Reach: 80.3% Loss to follow- up: Group 1 10%, Group 2 14%, Control 7.8%
Watson (2015, 2018)[103] [.] [104] Australia	RCT	Exercise	Postmenopausal women 101	∞	30 min 2x per wk for 35 wk	Face to face	University campus	Group	Group vs indi- vidual care	Targeted at specific health condition	BMD	<i>p</i> > 0.05 LS: +29% vs control -1.2% <i>p</i> < 0.001 FN: +0.1% vs control -1.8% <i>p</i> = 0.001	Reach: 48.3% Loss to follow- up: Interven- tion 12.2%; Control 17.3%

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Table 4. (Continued)													
Author (year)	Study design	Study design Type of MoC Population and sample size (n)	Population and sample size (n)	Follow- up		Delive	Delivery of MoC		EPOC	EPOC taxonomy	Clin	Clinical outcomes	Program reach and loss to
				montus	Frequency of contact	Contact method	Contact location	Group vs individual care	Delivery arrangement	Implementation strategy	Primary outcome?	BMD change	dn-wollol
Winters-Stone (2011) [105] United States	RCT	Exercise	≥50yr postmeno- pausal women with breast cancer 106	12	60 min 3x per wk for 52 wk	Face to face Written	University	Both	Group vs indi- vidual care	Targeted at specific health condition	BMD	LS: +0.41% vs con- trol -2.27%, p=0.013 TH: -0.35% vs control -0.83%, p>0.05 FN: -1.37% vs control -0.63% vs control	Reach: not reported Loss to follow- up: Interven- tion 30.8%;
Specialist Review Cheung (2013)[106] Australia	Pre-test post-test	Specialist review	Men with prostate cancer on ADT 113	24	2-3 monthly	Face to face	Hospital	Individual	Packages of care	Clinical practice guidelines	BMD	-2.00%, p.20.00 LS: -1.2% vs baseline, p=0.66 TH: -2.1% vs base- line, p.6.001	Reach: not reported Loss to follow- up: 26.1%

Footnote: p values are between groups unless otherwise specified. MoC: model of care; EPOC: effective practice and organisation of care; BMD: bone mineral density; FLS: fracture liaison service; yr: year; ED: emergency department; MTF:

chronic obstructive pulmonary disease; HRQCT: high resolution quantitative computer tomography; CTI: cortical thickness index; CBV: cortical bone volume.

TH: total hip; US: ultrasound; RCT: randomised controlled

spine; 7

minimal trauma fracture; LS: lumbar

trial; OP: osteoporosis; FN: femoral neck; min: minutes; wk: week; TF: total femur; BMI: body mass index; DR: distal radius; COPD:

demonstrated no fracture reduction overall, or when analysed separately for models grouped by activity [15]. The other study included all study designs, but due to the small number of studies, lack of control group and lack of power, no statement could be made about the efficacy for fracture reduction [17]. It is important to note that we have reported fracture outcomes in any study reporting this, whether or not it was the primary outcome. We would like to highlight that many studies were not powered for fracture outcomes and did not include follow-up of sufficient duration to find a meaningful difference in fracture rates. Many studies also did not include a comparison group due to the study design. Of those that did compare fracture rates, less than half found a significant reduction in fractures, and few of these studies were graded as high quality. As a reduction in fractures is the most important outcome for any osteoporosis MoC, we hope that studies continue to follow up and report on fractures over time. More recently, BMD has been suggested as a surrogate marker for osteoporosis therapeutic trials. Few MoC other than exercise studies have reported this outcome, but it could be considered by investigators in the future.

There are several limitations to our study. The study is descriptive only and does not include comparative statistics due to the broad inclusion criteria in our search. In describing our primary clinical outcomes of fractures and BMD change, we included studies with these as both primary or secondary outcomes. Given this, studies may have been underpowered for these specific outcomes. Strengths of our study include summarising delivery and implementation characteristics of studies, and using the validated EPOC classification system to categorise MoC, which can be applied to a broad variety of different interventions, and reproduced in future studies. We have also included all types of study designs, reflecting the fact that RCTs are not always appropriate for reporting complex interventions, and making this review a comprehensive summary of MoC worldwide.

This comprehensive scoping review in a vital area of rising morbidity and mortality reveals a wide variety of MoC for people with or at risk of osteoporosis. A minority of studies reports delivery and implementation characteristics, and this may influence the efficacy of these models, and the ability to translate them to real-world practice. Results of the MoC demonstrate mixed efficacy for fracture reduction, increases in BMD, and other outcomes such as treatment and DXA rates, and these disparities may be explained by exploring implementation characteristics. We suggest that future studies should include implementation outcomes in their reports, consider a pragmatic trial or effectiveness implementation hybrid trial study design, and report on fractures, or BMD increases as a surrogate marker for this. Lastly, co-design, and the perspectives of clinicians and consumers, is vital to implementation. It is important that researchers recognise this and ensure that these perspectives are included in future studies.

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Data sharing statement

Data dictionary, data collection table, list of excluded studies provided on request to AJ, at alicia.jones@monash.edu.

Declaration of Competing Interest

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AV reports no conflicts of interest.

Supplementary materials

Supplementary material associated with this article can be found in the online version at doi:10.1016/j.eclinm.2021.101022.

References

- Wright NC, Looker AC, Saag KG, et al. The recent prevalence of osteoporosis and low bone mass in the United States based on bone mineral density at the femoral neck or lumbar spine. J Bone Miner Res 2014;29(11):2520–6.
- [2] Watts JJ, Abimanyi-Ochom J, Sanders KM, Osteoporosis costing all Australians a new burden of disease analysis-2012 to 2022. Melbourne, Vic: Osteoporosis Australia; 2013.
- [3] Warriner AH, Patkar NM, Curtis JR, et al. Which fractures are most attributable to osteoporosis? J Clin Epidemiol 2011;64(1):46–53.
- [4] Amin S, Achenbach SJ, Atkinson EJ, Khosla S, Melton 3rd. LJ. Trends in fracture incidence: a population-based study over 20 years. J Bone Miner Res 2014;29 (3):581–9.
- [5] Borgström F, Karlsson L, Ortsäter G, et al. Fragility fractures in Europe: burden, management and opportunities. Arch Osteoporos 2020;15(1):59.
- [6] Johnell O, Kanis JA. An estimate of the worldwide prevalence and disability associated with osteoporotic fractures. Osteoporos Int 2006;17(12):1726–33.
- [7] Odén A, McCloskey EV, Kanis JA, Harvey NC, Johansson H. Burden of high fracture probability worldwide: secular increases 2010–2040. Osteoporos Int 2015;26(9):2243–8.
- [8] Cehic M, Lerner RG, Achten J, Griffin XL, Costa ML, Prieto-Alhambra D. Prescribing and adherence to bone protection medications following hip fracture in the United Kingdom: results from the world hip trauma evaluation (WHiTE) cohort study. Bone Jt J 2019;101-B(11):1402–7.
- [9] Hansen D, Bazell C, Pelizzari P, Pyenson B. Medicare cost of osteoporotic fractures. The clinical and cost burden of an important consequence of osteoporosis. Milliman Research Report For the National Osteoporosis Foundation; 2019. Accessible at: https://assets.milliman.com/ektron/Medicare_cost_of_osteoporotic_fractures.pdf.
- [10] Gillespie CW, Morin PE. Trends and disparities in osteoporosis screening among women in the United States, 2008–2014. Am J Med 2017;130(3):306–16.
- [11] Jones AR, Tay CT, Melder A, Vincent AJ, Teede H. What are models of care? a systematic search and narrative review to guide development of care models for premature ovarian insufficiency. Semin Reprod Med 2021.
- [12] Shepstone L, Lenaghan E, Cooper C, et al. Screening in the community to reduce fractures in older women (SCOOP): a randomised controlled trial. Lancet 2018;391(10122):741–7 (London, England).
- [13] Burch J, Tort S, (on behalf of Cochrane Clinical Answers Editors). What are the effects of professional interventions for general practitioners aimed at improving management of osteoporosis? Cochrane Clinical Answers. 2017.
- [14] Shojaa M, von Stengel S, Kohl M, Schoene D, Kemmler W. Effects of dynamic resistance exercise on bone mineral density in postmenopausal women: a systematic review and meta-analysis with special emphasis on exercise parameters. Osteoporos Int 2020;31(8):1427–44.
- [15] Kastner M, Perrier L, Munce SEP, et al. Complex interventions can increase osteoporosis investigations and treatment: a systematic review and meta-analysis. Osteoporos Int 2018;29(1):5–17.
- [16] Nayak S, Greenspan SL. How can we improve osteoporosis care? a systematic review and meta-analysis of the efficacy of quality improvement strategies for osteoporosis. J Bone Miner Res 2018;33(9):1585–94.
- [17] Ganda K, Puech M, Chen JS, et al. Models of care for the secondary prevention of osteoporotic fractures: a systematic review and meta-analysis. Osteoporos Int 2013;24(2):393–406.
- [18] Sale JEM, Beaton D, Posen J, Elliot-Gibson V, Bogoch E. Systematic review on interventions to improve osteoporosis investigation and treatment in fragility fracture patients. Osteoporos Int 2011;22(7):2067–82.
- [19] Little EA, Eccles MP. A systematic review of the effectiveness of interventions to improve post-fracture investigation and management of patients at risk of osteoporosis. Implement Sci 2010;5:80.
- [20] Black DM, Bauer DC, Vittinghoff E, et al. Treatment-related changes in bone mineral density as a surrogate biomarker for fracture risk reduction: meta-regression analyses of individual patient data from multiple randomised controlled trials. Lancet Diabetes Endocrinol 2020;8(8):672–82.

- [21] Proctor E, Silmere H, Raghavan R, et al. Outcomes for implementation research: conceptual distinctions, measurement challenges, and research agenda. Adm Policy Ment Health 2011;38(2):65–76.
- [22] Wolfenden L, Foy R, Presseau J, et al. Designing and undertaking randomised implementation trials: guide for researchers. BMJ 2021;372:m3721.
- [23] Pinnock H, Barwick M, Carpenter CR, et al. Standards for reporting implementation studies (StaRI) statement. BMJ 2017;356:i6795.
- [24] Glasgow RE, Vogt TM, Boles SM. Evaluating the public health impact of health promotion interventions: the RE-AIM framework. Am J Public Health 1999;89 (9):1322–7.
- [25] Cooper C, Cole ZA, Holroyd CR, et al. Secular trends in the incidence of hip and other osteoporotic fractures. Osteoporos Int 2011;22(5):1277–88.
- [26] Mithal A, Dhingra V, Lau E. The Asian Audit: Epidemiology, costs and burden of osteoporosis in Asia 2009. Switzerland: International Osteoporosis Foundation; 2009.
- [27] In Peters M, Godfrey C, McInerney P, C BaldiniSoares, Khalil H, Parker D. Chapter 11: scoping reviews. In: Aromataris E, Munn Z, editors. Joanna briggs institute reviewer's manual. The Joanna Briggs Institute; 2017.
- [28] Cochrane Effective Practice and Organisation of Care (EPOC). Describing interventions in EPOC reviews. EPOC resources for review authors 2017 [Available from: epoc.cochrane.org/resources/epoc-resources-review-authors. Accessed 1 April 2020]
- [29] Jessup RL, O'Connor DA, Putrik P, et al. Alternative service models for delivery of healthcare services in high-income countries: a scoping review of systematic reviews. BMJ Open 2019;9(1):e024385.
- [30] Effective Practice and Organisation of Care (EPOC). EPOC taxonomy 2015 [Available from: epoc.cochrane.org/epoc-taxonomy. Accessed 29 Aug 2019]
- [31] Healthcare Improvement Scotland. Scottish intercollegiate guidelines network (SIGN) methodology checklists 2012 [Available from: https://www.sign.ac.uk/ what-we-do/methodology/checklists/. Accessed 5th October 2020]
- [32] Durlak JA, DuPre EP. Implementation matters: a review of research on the influence of implementation on program outcomes and the factors affecting implementation. Am J Community Psychol 2008;41(3-4):327.
- [33] Public Participation Team. Patient and public participation policy. England: NHS; 2017.
- [34] Moore GF, Audrey S, Barker M, et al. Process evaluation of complex interventions: medical research council guidance. BMJ 2015;350 h1258. https://www. bmj.com/content/350/bmj.h1258.
- [35] Martin J, Viprey M, Castagne B, et al. Interventions to improve osteoporosis care: a systematic review and meta-analysis. Osteoporos Int 2020;31(3):429–46.
- [36] Eisman JA, Bogoch ER, Dell R, et al. Making the first fracture the last fracture: ASBMR task force report on secondary fracture prevention. J Bone Miner Res 2012;27(10):2039–46.
- [37] Amphansap T, Stitkitti N, Dumrongwanich P, Evaluation of police general hospital's fracture liaison service (PGH's FLS): the first study of a fracture liaison service in Thailand. Osteoporos Sarcopenia 2016;2(4):238–43.
- [38] Bachour F, Rizkallah M, Sebaaly A, et al. Fracture liaison service: report on the first successful experience from the Middle East. Arch Osteoporos 2017;12 (1):79.
- [39] Davidson E, Seal A, Doyle Z, Fielding K, McGirr J. Prevention of osteoporotic refractures in regional Australia. Aust J Rural Health 2017;25(6):362–8.
- [40] Huntjens KMB, van Geel TCM, Geusens PP, et al. Impact of guideline implementation by a fracture nurse on subsequent fractures and mortality in patients presenting with non-vertebral fractures. Injury 2011;42(Suppl 4):S39–43 0226040, gon.
- [41] Inderjeeth CA, Raymond WD, Briggs AM, Geelhoed E, Oldham D, Mountain D. Implementation of the Western Australian osteoporosis model of care: a fracture liaison service utilising emergency department information systems to identify patients with fragility fracture to improve current practice and reduce re-fracture rates: a 12-month analysis. Osteoporos Int 2018;29(8):1759-70.
- [42] Lih A, Nandapalan H, Kim M, et al. Targeted intervention reduces refracture rates in patients with incident non-vertebral osteoporotic fractures: a 4-year prospective controlled study. Osteoporos Int 2011;22(3):849–58.
- [43] Nakayama A, Major G, Holliday E, Attia J, Bogduk N. Evidence of effectiveness of a fracture liaison service to reduce the re-fracture rate. Osteoporos Int 2016;27 (3):873–9.
- [44] Van der Kallen J, Giles M, Cooper K, et al. A fracture prevention service reduces further fractures two years after incident minimal trauma fracture. Int J Rheum Dis 2014;17(2):195–203.
- [45] Wasfie T, Jackson A, Brock C, Galovska S, McCullough JR, Burgess JA. Does a fracture liaison service program minimize recurrent fragility fractures in the elderly with osteoporotic vertebral compression fractures? Am J Surg 2019;217 (3):557–60.
- [46] Becker C, Cameron ID, Klenk J, et al. Reduction of femoral fractures in long-term care facilities: the Bavarian fracture prevention study. PloS One 2011;6(8): e24311.
- [47] Heinrich S, Rapp K, Stuhldreher N, Rissmann U, Becker C, Konig HH. Cost-effectiveness of a multifactorial fall prevention program in nursing homes. Osteoporos Int 2013;24(4):1215–23.
- [48] Pekkarinen T, Loyttyniemi E, Valimaki M. Hip fracture prevention with a multifactorial educational program in elderly community-dwelling finnish women. Osteoporos Int 2013;24(12):2983–92.
- [49] Sorbi R, Aghamirsalim M, Eslami V, Karimi Dastjerdi MH, Seraj SM. Effect of an educational intervention for surgeons on osteoporosis management at

2-year follow-up in patients with fragility fracture. J Clin Rheumatol 2016;22(4):231-2.

- [50] Harness NG, Funahashi T, Dell R, et al. Distal radius fracture risk reduction with a comprehensive osteoporosis management program. J Hand Surg Am 2012;37 (8):1543–9.
- [51] Parsons CM, Harvey N, Shepstone L, et al. Systematic screening using FRAX leads to increased use of, and adherence to, anti-osteoporosis medications: an analysis of the UK SCOOP trial.. Oxford, United Kingdom: Cooper) NIHR Oxford Biomedical Research Centre, University of Oxford; 2019 Osteoporos Int..
- [52] Zhumkhawala AA, Gleason JM, Cheetham TC, et al. Osteoporosis management program decreases incidence of hip fracture in patients with prostate cancer receiving androgen deprivation therapy. Urology 2013;81(5):1010–5.
- [53] Kemmler W, Bebenek M, Kohl M, von Stengel S. Exercise and fractures in postmenopausal women. Final results of the controlled erlangen fitness and osteoporosis prevention study (EFOPS). Osteoporos Int 2015;26(10):2491–9.
- [54] Kemmler W, Engelke K, von Stengel S. Long-term exercise and bone mineral density changes in postmenopausal women-are there periods of reduced effectiveness? J Bone Miner Res 2016;31(1):215–22.
- [55] Kemmler W, Kohl M, von Stengel S. Long-term effects of exercise in postmenopausal women: 16-year results of the erlangen fitness and osteoporosis prevention study (EFOPS). Menopause 2017;24(1):45–51.
- [56] Kemmler W, von Stengel S. Dose-response effect of exercise frequency on bone mineral density in post-menopausal, osteopenic women. Scand J Med Sci Sport 2014;24(3):526–34.
- [57] Kemmler W, von Stengel S, Bebenek M, Engelke K, Hentschke C, Kalender WA. Exercise and fractures in postmenopausal women: 12-year results of the erlangen fitness and osteoporosis prevention study (EFOPS). Osteoporos Int 2012;23 (4):1267–76.
- [58] Kemmler W, von Stengel S, Kohl M. Exercise frequency and bone mineral density development in exercising postmenopausal osteopenic women. Is there a critical dose of exercise for affecting bone? Results of the erlangen fitness and osteoporosis prevention study. Bone 2016;89:1–6 asr, 8504048.
- [59] Korpelainen R, Keinanen-Kiukaanniemi S, Nieminen P, Heikkinen J, Vaananen K, Korpelainen J. Long-term outcomes of exercise: follow-up of a randomized trial in older women with osteopenia. Arch Intern Med 2010;170(17):1548–56.
- [60] Cheung WH, Shen WY, Dai DLK, et al. Evaluation of a multidisciplinary rehabilitation programme for elderly patients with hip fracture: a prospective cohort study. J Rehabil Med 2018;50(3):285–91.
- [61] Gomez F, Curcio CL, Brennan-Olsen SL, et al. Effects of the falls and fractures clinic as an integrated multidisciplinary model of care in Australia: a pre-post study. BMJ Open 2019;9(7):e027013.
- [62] Chandran M, Tan MZW, Cheen M, Tan SB, Leong M, Lau TC. Secondary prevention of osteoporotic fractures—an "OPTIMAL" model of care from Singapore. Osteoporos Int 2013;24(11):2809–17.
- [63] Eekman DA, van Helden SH, Huisman AM, et al. Optimizing fracture prevention: the fracture liaison service, an observational study. Osteoporos Int 2014;25 (2):701–9.
- [64] Hien VTT, Khan NC, Mai LB, et al. Effect of community-based nutrition education intervention on calcium intake and bone mass in postmenopausal Vietnamese women. Public Health Nutr 2009;12(5):674–9.
- [65] Wang L, Xu X, Hao H, et al. A model of health education and management for osteoporosis prevention. Exp Ther Med 2016;12(6):3797–805.
- [66] Aboarrage AM, Teixeira CVLS, Dos Santos RN, et al. A high-intensity jump-based aquatic exercise program improves bone mineral density and functional fitness in postmenopausal women. Rejuvenation Res 2018;21(6):535–40.
- [67] Alayat MSM, Abdel-Kafy EM, Thabet AAM, Abdel-Malek AS, Ali TH, Header EA. Long-term effect of pulsed Nd-YAG laser combined with exercise on bone mineral density in men with osteopenia or osteoporosis: 1 year of follow-up. Photomed Laser Surg 2018;36(2):105–11.
- [68] Almstedt HC, Grote S, Korte JR, et al. Combined aerobic and resistance training improves bone health of female cancer survivors. Bone Rep. 2016;5:274–9 ((Grote) Center for Nutrition, Healthy Lifestyle, and Disease Prevention, School of Public Health, Loma Linda University, Loma Linda, CA, United States).
- [69] Angin E, Erden Z, Can F. The effects of clinical pilates exercises on bone mineral density, physical performance and quality of life of women with postmenopausal osteoporosis. J Back Musculoskelet Rehabil 2015;28(4):849–58.
- [70] Astorino TA, Harness ET, Witzke KA. Effect of chronic activity-based therapy on bone mineral density and bone turnover in persons with spinal cord injury. Eur J Appl Physiol 2013;113(12):3027–37.
- [71] Basat H, Esmaeilzadeh S, Eskiyurt N. The effects of strengthening and highimpact exercises on bone metabolism and quality of life in postmenopausal women: a randomized controlled trial. J Back Musculoskelet Rehabil 2013;26 (4):427–35.
- [72] Beavers DP, Beavers KM, Loeser RF, et al. The independent and combined effects of intensive weight loss and exercise training on bone mineral density in overweight and obese older adults with osteoarthritis. Osteoarthr Cartil 2014;22 (6):726–33.
- [73] Bergstrom I, Parini P, Gustafsson SA, Andersson G, Brinck J. Physical training increases osteoprotegerin in postmenopausal women. J Bone Miner Metab 2012;30(2):202–7.
- [74] Bocalini DS, Serra AJ, dos Santos L, Murad N, Levy RF. Strength training preserves the bone mineral density of postmenopausal women without hormone replacement therapy. J Aging Health 2009;21(3):519–27.

- [75] Bolton KL, Egerton T, Wark J, et al. Effects of exercise on bone density and falls risk factors in post-menopausal women with osteopenia: a randomised controlled trial. J Sci Med Sport 2012;15(2):102–9.
- [76] Borba-Pinheiro CJ, Dantas EHM, Vale RGDS, et al. Resistance training programs on bone related variables and functional independence of postmenopausal women in pharmacological treatment: a randomized controlled trial. Arch Gerontol Geriatr 2016;65:36–44 8214379, 7ax.
- [77] Chuin A, Labonte M, Tessier D, et al. Effect of antioxidants combined to resistance training on BMD in elderly women: a pilot study. Osteoporos Int 2009;20 (7):1253–8.
- [78] Daly RM, Gianoudis J, Kersh ME, et al. Effects of a 12-month supervised, community-based, multimodal exercise program followed by a 6-month research-topractice transition on bone mineral density, trabecular microarchitecture, and physical function in older adults: a randomized controlled trial. J Bone Miner Res 2020;35(3):419–29 ((Hill) School of Physiotherapy and Exercise Science, Curtin University, Perth, Australia).
- [79] Gianoudis J, Bailey CA, Ebeling PR, et al. Effects of a targeted multimodal exercise program incorporating high-speed power training on falls and fracture risk factors in older adults: a community-based randomized controlled trial. J Bone Miner Res 2014;29(1):182–91.
- [80] de Matos O, Lopes da Silva DJ, Martinez de Oliveira J, Castelo-Branco C. Effect of specific exercise training on bone mineral density in women with postmenopausal osteopenia or osteoporosis. Gynecol Endocrinol 2009;25(9):616–20.
- [81] El-Kader SMA, Al-Jiffri OH, Al-Jiffri HO. Aerobic exercise training modulates bone mineral status in patients with chronic obstructive pulmonary disease. Eur J Gen Med 2016;13(3):51–4.
- [82] Elsisi HFEM, Mousa GSM, Eldesoky MTM. Electromagnetic field versus circuit weight training on bone mineral density in elderly women. Clin Interv Aging 2015;10:539–47 101273480.
- [83] Garcia-Gomariz C, Blasco JM, Macian-Romero C, Guillem-Hernandez E, Igual-Camacho C. Effect of 2 years of endurance and high-impact training on preventing osteoporosis in postmenopausal women: randomized clinical trial. Menopause 2018;25(3):301–6.
- [84] Hojan K, Milecki P, Leszczynski P. The impact of aerobic exercises on bone mineral density in breast cancer women during endocrine therapy. Pol Orthop Traumatol 2013;78:47–51 101593867.
- [85] Hojan K, Milecki P, Molinska-Glura M, Roszak A, Leszczynski P. Effect of physical activity on bone strength and body composition in breast cancer premenopausal women during endocrine therapy. Eur J Phys Rehabil Med 2013;49(3):331–9.
- [86] Kemmler W, Bebenek M, von Stengel S, Engelke K, Kalender WA. Effect of blockperiodized exercise training on bone and coronary heart disease risk factors in early post-menopausal women: a randomized controlled study. Scand J Med Sci Sport 2013;23(1):121–9.
- [87] Kukuljan S, Nowson CA, Bass SL, et al. Effects of a multi-component exercise program and calcium-vitamin-D 3-fortified milk on bone mineral density in older men: a randomised controlled trial. Osteoporos Int 2009;20(7):1241–51.
- [88] Kukuljan S, Nowson CA, Sanders KM, et al. Independent and combined effects of calcium-vitamin D3 and exercise on bone structure and strength in older men: an 18-month factorial design randomized controlled trial. J Clin Endocrinol Metab 2011;96(4):955–63.
- [89] LeBlanc A, Matsumoto T, Jones J, et al. Bisphosphonates as a supplement to exercise to protect bone during long-duration spaceflight. Osteoporos Int 2013;24 (7):2105–14.
- [90] Sibonga J, Matsumoto T, Jones J, et al. Resistive exercise in astronauts on prolonged spaceflights provides partial protection against spaceflight-induced bone loss. Bone, 2019;128:112037.
- [91] Liu BX, Chen SP, Li YD, et al. The effect of the modified eighth section of eightsection brocade on osteoporosis in postmenopausal women: a prospective randomized trial. Medicine 2015;94(25):e991.
- [92] Marchese D, D'Andrea M, Ventura V, et al. Effects of a weight-bearing exercise training on bone mineral density and neuromuscular function of osteopenic women. Eur J Inflamm 2012;10(3):427–35.
- [93] Marques EA, Wanderley F, Machado L, et al. Effects of resistance and aerobic exercise on physical function, bone mineral density, OPG and RANKL in older women. Exp Gerontol 2011;46(7):524–32.
- [94] Morse LR, Troy KL, Fang Y, et al. Combination therapy with zoledronic acid and FES-row training mitigates bone loss in paralyzed legs: results of a randomized comparative clinical trial. JBMR Plus 2019;3(5):e10167.
- [95] Murai IH, Roschel H, Dantas WS, et al. Exercise mitigates bone loss in women with severe obesity after roux-en-y gastric bypass: a randomized controlled trial. J Clin Endocrinol Metab 2019;104(10):4639–50.
- [96] Murtezani A, Nevzati A, Ibraimi Z, Sllamniku S, Meka VS, Abazi N. The effect of land versus aquatic exercise program on bone mineral density and physical function in postmenopausal women with osteoporosis: a randomized controlled trial. Ortop Traumatol Rehabil 2014;16(3):319–25.
- [97] Nicholson VP, McKean MR, Slater GJ, Kerr A, Burkett BJ. Low-load very high-repetition resistance training attenuates bone loss at the lumbar spine in active post-menopausal women. Calcif Tissue Int 2015;96(6):490–9.
- [98] Saarto T, Sievanen H, Kellokumpu-Lehtinen P, et al. Effect of supervised and home exercise training on bone mineral density among breast cancer patients. a 12-month randomised controlled trial. Osteoporos Int 2012;23(5):1601–12.

- [99] Sen EI, Esmaeilzadeh S, Eskiyurt N. Effects of whole-body vibration and high impact exercises on the bone metabolism and functional mobility in postmenopausal women. J Bone Miner Metab 2020;38(3):392–404.
- [100] Silverman NE, Nicklas BJ, Ryan AS. Addition of aerobic exercise to a weight loss program increases BMD, with an associated reduction in inflammation in overweight postmenopausal women. Calcif Tissue Int 2009;84(4):257–65.
- [101] Villareal DT, Aguirre L, Gurney AB, et al. Aerobic or resistance exercise, or both, in dieting obese older adults. N Engl J Med 2017;376(20):1943–55.
- [102] von Stengel S, Kemmler W, Engelke K, Kalender WA. Effects of whole body vibration on bone mineral density and falls: results of the randomized controlled ELVIS study with postmenopausal women. Osteoporos Int 2011;22 (1):317–25.
- [103] Watson SL, Weeks BK, Weis LJ, Horan SA, Beck BR. Heavy resistance training is safe and improves bone, function, and stature in postmenopausal women with

low to very low bone mass: novel early findings from the LIFTMOR trial. Osteoporos Int 2015;26(12):2889–94.

- [104] Watson SL, Weeks BK, Weis LJ, Harding AT, Horan SA, Beck BR. High-intensity resistance and impact training improves bone mineral density and physical function in postmenopausal women with osteopenia and osteoporosis: the LIFT-MOR randomized controlled trial. J Bone Miner Res 2018;33(2):211–20.
- [105] Winters-Stone KM, Dobek J, Nail L, et al. Strength training stops bone loss and builds muscle in postmenopausal breast cancer survivors: a randomized, controlled trial. Breast Cancer Res Treat 2011;127(2):447–56.
- [106] Cheung AS, Pattison D, Bretherton I, et al. Cardiovascular risk and bone loss in men undergoing androgen deprivation therapy for non-metastatic prostate cancer: implementation of standardized management guidelines. Andrology 2013;1(4):583–9.