Recommendations for preventing fracture in long-term care

Alexandra Papaioannou MD MSc, Nancy Santesso RD PhD, Suzanne N. Morin MD MSc, Sidney Feldman MD, Jonathan D. Adachi MD, Richard Crilly BSc MD, Lora M. Giangregorio PhD, Susan Jaglal PhD, Robert G. Josse MBBS, Sharon Kaasalainen PhD, Paul Katz MD, Andrea Moser MD MSc, Laura Pickard MA, Hope Weiler RD PhD, Susan Whiting PhD, Carly J. Skidmore MSc, Angela M. Cheung MD PhD; for the Scientific Advisory Council of Osteoporosis Canada

This document provides guidance regarding strategies for the prevention of fractures directed toward interprofessional teams caring for frail older adults in long-term care.

Methods

This guideline, which has been endorsed by Osteoporosis Canada, was developed using the Grading of Recommendations Assessment, Development and Evaluation (GRADE) approach[^18] ([www.gradeworkinggroup.org](http://www.gradeworkinggroup.org)), in a process led by a GRADE methodologist (N.S.). The guideline panel comprised the authors, other multidisciplinary health care providers and researchers, and representatives from resident and family councils (see Appendix 1, available at www.cmaj.ca/lookup/suppl/doi:10.1503/cmaj.141331/-/DC1). The panel was first surveyed to prioritize questions and important outcomes. In addition to fractures (hip, vertebral and nonvertebral), the group as a whole identified pain, quality of life, loss of activities of daily living and mobility, death and adverse events requiring medical attention as important outcomes. Family members of residents ranked prevention of pain and maintenance of mobility as most important.

We conducted systematic searches of the literature for published network meta-analyses, system-
atic reviews, randomized controlled trials and non-randomized studies up to June 2013. Details of the synthesis of the evidence, preparation of evidence profiles and evidence-to-decision tables for five key strategies to prevent fracture (vitamin D and calcium intake, osteoporosis medications, hip protectors, exercise and multifactorial interventions) are presented in Appendix 1. Briefly, evidence for relative risks and differences among interventions were converted to absolute effects with 95% confidence intervals (CIs) and were presented in evidence profiles. When data regarding fractures in long-term care were not available, we used data for adults 75 years and older. The overall annual rate of hip, vertebral and other fractures in long-term care residents was 2%;\textsuperscript{2,20–22} for those at high risk, the annual rate of vertebral fracture was 20%,\textsuperscript{23} and annual rates were estimated to be 6% for hip fracture and 6% for all other fractures.

We assessed the quality of the evidence as high, moderate, low or very low according to the GRADE criteria\textsuperscript{18} (Table 1). The evidence-to-decision tables presented to the panel for consideration included a summary of the evidence for benefits and harms, the quality of the evidence, relevant values and preferences of residents and their families, resource use and feasibility. Recommendations were assessed as “strong” or “conditional” (Table 2). Strong recommendations are worded as “we recommend” and conditional recommendations as “we suggest.”

### Recommendations

We developed recommendations for interventions to prevent fracture for two groups: older residents in long-term care who are at high risk of fracture and older residents who are not at high risk of fracture.

Until a method for defining those at high risk of fracture in long-term care has been validated, we propose an adaptation of the definition presented in the 2010 Osteoporosis Canada guideline,\textsuperscript{1} which relies heavily on fracture history (Box 1). If a resident has been identified as having a high risk of fracture and has received osteoporosis treatments before admission to long-term care, that classification may continue to apply at admission.

Screening for vertebral fractures as described in the 2010 guideline\textsuperscript{1} is recommended. If a lateral view is included when chest radiography is ordered, a request can also be made to screen for vertebral fractures.

### Calcium and vitamin D

#### Calcium

*For all residents, we recommend dietary interventions to meet the recommended dietary allowance for calcium (strong recommendation; moderate-quality evidence).*

This recommendation places a high value on reductions in fractures, mortality and falls and a

| **Table 1:** Quality of evidence: confidence in effect\textsuperscript{18} |
|-----------------|---------------------------------------------------------------|
| **Rating of evidence quality** | **Definition** |
| High            | We are very confident that the true effect lies close to that of the estimate of the effect. |
| Moderate        | We are moderately confident in the effect estimate. The true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different. |
| Low             | Our confidence in the effect estimate is limited. The true effect may be substantially different from the estimate of the effect. |
| Very low        | We have very little confidence in the effect estimate. The true effect is likely to be substantially different from the estimate of effect. |

| **Table 2:** Interpretation of strong and conditional recommendations for fracture prevention\textsuperscript{19} |
|-----------------|---------------------------------------------------------------|
| **Target group** | **Type of recommendation; implication for target group** |
|                 | Strong recommendation ("We recommend …") | Conditional recommendation ("We suggest …") |
| Patients        | Most individuals in this situation would want the recommended course of action, and only a small proportion would not. | The majority of individuals in this situation would want the suggested course of action, but many would not. |
| Clinicians      | Most individuals should receive the intervention. | Clinicians recognize that different choices will be appropriate for each individual patient and that clinicians must help each individual arrive at a management decision consistent with his or her values and preferences. |
lower value on the resources in long-term care that are required to implement interventions to ensure adequate dietary intake of calcium. This recommendation is based on evidence evaluating the effects of calcium supplements, which was used as direct evidence for dietary intake; however, dietary interventions do not have the adverse effects of supplements. For people older than 70 years, the recommended dietary allowance for calcium is 1200 mg daily (3 servings of dairy or dairy equivalents).

For residents at high risk of fractures who cannot meet the recommended dietary allowance for calcium through dietary intake, we recommend daily supplements of calcium up to 500 mg (strong recommendation; moderate-quality evidence).

For residents who are not at high risk of fractures and who cannot meet the recommended dietary allowance for calcium through dietary intake, we suggest daily supplements of calcium up to 500 mg, depending on resources and their (or their carers’) values and preferences (conditional recommendation; moderate-quality evidence).

The recommendation for residents at high risk places a high value on the reduction in hip fractures and the small reductions in vertebral and nonvertebral fractures and in mortality that can be achieved with calcium supplementation. It places a lower value on the small increased risk of gastrointestinal adverse effects that may occur and the resources required in long-term care to provide calcium supplementation.

The recommendation for residents not at high risk is conditional, as there may be little to no benefit of calcium supplementation, and adverse effects of supplementation, such as gastrointestinal and renal adverse effects, may occur. For residents who value avoiding these adverse effects, supplementation may not be a desirable option.

These recommendations apply to supplementation with any calcium compound, including calcium carbonate or citrate. The recommendation to limit supplementation to 500 mg was based on the uncertainty about harms of calcium supplementation in studies of community-dwelling individuals who received calcium supplementation of 1000 mg or more daily. The benefits of calcium supplementation are closely linked to adequate vitamin D intake.

Vitamin D

For residents at high risk of fractures, we recommend daily supplements of 800 IU to 2000 IU vitamin D₃ to meet the recommended dietary allowance, depending on resources and their (or their carers’) values and preferences (conditional recommendation; moderate-quality evidence).

The recommendation for residents at high risk places a high value on reductions in hip fractures, mortality and falls and a lower value on the resources in long-term care that are required to provide vitamin D supplementation. The recommendation for residents not at high risk also places a high value on reduction in falls, as they may lead to serious injuries, fear of falling and burden to staff in long-term care; however, there is some uncertainty about a reduction in falls and little to no reduction in fractures with vitamin D supplementation in this group.

These recommendations apply to supplementation with D₃, as this form may be more accessible because of its lower cost relative to D₂. A dose of about 800 IU reduced fractures in people with normal or low 25-hydroxyvitamin D levels and also increased 25-hydroxyvitamin D levels to normal in those with low levels; therefore, 800 IU is recommended. However, the exact dose may depend on the dosing regimen that is available (e.g., a 1000 IU drop or tablet would be acceptable). The benefits of vitamin D supplementation are closely linked to adequate calcium intake, and therefore recommendations for calcium intake should also be applied. The recommended dietary allowance for vitamin D for people older than 70 years is 800 IU daily, and the tolerable upper intake level is up to 4000 IU.

Summary of the evidence

Overall, there was moderate-quality evidence for benefits and low to very low quality evidence for harms of calcium and vitamin D. We found that vitamin D in addition to calcium probably reduces hip fractures and mortality more than vitamin D alone or calcium alone; for residents at high risk, we estimated 15 fewer hip fractures (95% CI –24 to –5) per 1000; for residents not at high risk, we estimated 5 fewer hip fractures (95% CI –24 to –5) per 1000; for all residents, we estimated 7 fewer deaths (95% CI –14 to –1) per 1000.

Box 1: Factors indicating high risk for fracture*

Residents in long-term care with any one of the following factors:
- prior fracture of the hip or spine OR
- more than one prior fracture† OR
- recent use of systemic glucocorticoids and one prior fracture† OR
- identified as high risk and/or receiving osteoporosis treatment before admission to long-term care

*Adapted from the 2010 Osteoporosis Canada guideline.†Excluding fractures of the hands, feet or ankles.
We found that vitamin D and calcium supplementation likely has little or no effect on vertebral fractures, with only 2 fewer vertebral fractures (95% CI: –44 to 61) per 1000. The effect is similar with vitamin D only, but a reduction may be likely with calcium only (49 fewer fractures [95% CI: –99 to 19] per 1000). Calcium, or vitamin D with or without calcium, probably has little to no effect on the incidence of nonvertebral fractures, quality of life or muscle strength.

The data for falls were not precise (with the CIs including the possibilities of benefit, no effect and harm), and the effects were not consistent when the rate or risk of falls was measured. However, vitamin D and calcium, or vitamin D alone, may reduce falls. This is important because one-third of all falls may result in an injury, and every fifth injurious fall may result in treatment outside the patient’s own setting. We found no data on pain, anxiety, mobility and performance of activities of daily living in relation to calcium and vitamin D.

With respect to minor and major adverse events, vitamin D or calcium supplements probably increase mild or serious gastrointestinal events to a similar extent, by about 8 (95% CI: 0 to 17) per 1000. Gastrointestinal symptoms or difficulties taking tablet may contribute to poor adherence. The evidence suggested slightly more cases of hypercalcemia (5 more cases per 1000, 95% CI: –1 to 18) and renal insufficiency or calculi (3 more cases per 1000, 95% CI: 0 to 6) with vitamin D (D$_3$ or D$_2$). The evidence for more myocardial infarctions with calcium supplementation of 1000 mg or more among community-dwelling individuals is uncertain, as it is not consistent with observed reductions in mortality, and the CIs around the estimates include no effect and the possibility of appreciable harm.

Subgroup analyses from systematic reviews showed that there may be little or no difference in rates of fractures or falls by type of vitamin D (D$_3$ or D$_2$), $^{26,30,38}$ that there may be greater benefits with vitamin D above 792 IU (actual intake in most studies between 792 and 844 IU) but no difference with less than or more than 1000 mg calcium, and that there are inconsistent effects when vitamin D is given in large monthly or annual doses. Analyses did show that vitamin D may have greater effects in reducing falls and fractures among people with low vitamin D status. Also found that about 800 IU daily over several months can increase serum vitamin D levels to “normal” in people with initial vitamin D deficiency (e.g., ≤ 25 nmol/L).

Pharmacologic therapies for those at high risk of fracture

When making recommendations about pharmacologic therapies for those at high risk of fracture, we placed a high value on issues that are prevalent among older persons and that may make it difficult to safely administer these medications and on the immediate risks of fractures and the consequences of falls and fractures, including increased pain, loss of mobility and greater risk of death. We calculated the effects of benefits and harms at one year or more; as such, these recommendations apply to older persons with life expectancy greater than one year.

For residents who are at high risk of fractures, we recommend that alendronate (weekly) or risedronate (weekly or monthly) be used as first-line therapy (strong recommendation; moderate-quality evidence).

The probable reductions in fractures (hip, vertebral and nonvertebral) and mortality with alendronate and risedronate outweigh the low or uncertain risk of harms or adverse effects, such as atypical femoral fractures. Alendronate and risedronate are recommended as first-line therapies because of their low cost relative to other therapies. Tablets of alendronate and risedronate are not to be crushed, and these drugs are to be provided to older persons who can remain upright for 30 minutes after administration. Some formulations must be administered at least 30 minutes before food intake. Other formulations can be taken with food.

For residents who are at high risk of fractures and who have difficulty taking oral medications, alternative first-line therapies are available (see below for recommendations concerning denosumab and zoledronic acid). The product monographs indicate that alendronate and risedronate are not recommended for older persons with severe renal insufficiency (creatinine clearance < 35 mL/min or < 30 mL/min, respectively).

For residents who are at high risk of fractures and who have difficulty taking oral medications, we recommend that zoledronic acid be used as first-line therapy (strong recommendation; moderate-quality evidence).

The probable reductions in fractures (hip, vertebral and nonvertebral) and mortality with zoledronic acid slightly outweigh the uncertain increased risk of musculoskeletal adverse effects (e.g., arthralgia, myalgia) and the higher cost relative to other first-line therapies. This recommendation applies to older persons who have difficulty taking oral medications because of dysphagia, an inability to sit up for 30 minutes, cognitive impairment or intolerance. The product
monograph for zoledronic acid indicates that infusion should be performed over no less than 15 minutes. Health Canada advises that caution is necessary for people who receive other medications that could affect renal function, that creatinine clearance should be monitored before and periodically after treatment, that appropriate hydration (500 mL of water) is necessary before and after treatment, and that this medication should not be given to people with severe renal impairment (creatinine clearance < 30 mL/min).

For residents who are at high risk of fractures and who have difficulty taking oral medications, we recommend that denosumab be used as first-line therapy (strong recommendation; moderate-quality evidence).

The reductions in fractures (hip, vertebral and nonvertebral) and mortality with denosumab slightly outweigh the small and uncertain risk of serious infections and greater cost relative to other first-line therapies. This recommendation applies to older persons who have difficulty taking oral medications because of dysphagia, an inability to sit up for 30 minutes, cognitive impairment or intolerance. Although denosumab may be prescribed for residents with renal impairment, the product monograph for denosumab indicates that in clinical studies, patients with renal impairment (creatinine clearance < 30 mL/min) and those receiving dialysis were at greater risk of hypocalcemia than those without renal impairment.

For residents who are at high risk of fractures, we suggest teriparatide (conditional recommendation; moderate-quality evidence).

Although the benefits of teriparatide (in particular for vertebral fractures) probably outweigh potential harms of treatment, the cost of therapy restricts access to this medication, and there may be a higher burden because of the need for daily injections. A low value was placed on the uncertain effect of teriparatide on back pain because of past treatment, and after treatment, and that this medication may include a small reduction in back pain related to past and future vertebral fractures for teriparatide, but there was little to no effect on quality of life for other therapies.

There was low to very low quality evidence for very small risks of serious events such as osteonecrosis of the jaw and atypical fractures or delayed healing (estimated at < 1 per 10000 community-dwelling older people). It is unclear whether these risks would be higher among long-term care residents. The evidence for risk of atrial fibrillation with bisphosphonates and cerebrovascular or cardiovascular events with raloxifene was also of low quality; these results were imprecise and include the possibility of small to no increases in these events. Venous thromboembolism may increase with raloxifene (12 more [95% CI 7 to 19] per 1000), musculoskeletal events may increase with zoledronic acid (146 more [95% CI 125 to 169] per 1000), and serious infections may increase with denosumab (8 more [95% CI 0 to 18] per 1000). Randomized controlled trials and pharmacovigilance for bisphosphonates and raloxifene showed little to no effect of these drugs on serious gastrointestinal events.

We estimated that direct drug costs were worth the overall beneficial consequences of most drugs, with the exception of etidronate and raloxifene. The costs of zoledronic acid, denosumab and teriparatide were also high relative to those of other therapies.
Hip protectors

For residents who are mobile and at high risk of fractures, we recommend hip protectors (strong recommendation; moderate-quality evidence).

For residents who are mobile but not at high risk of fractures, we suggest hip protectors, depending on resources available and the residents’ values and preferences (conditional recommendation; moderate-quality evidence).

These recommendations place a high value on avoiding the serious consequences of hip fractures in mobile residents, including pain, loss of mobility and death. A lower value was placed on the cost or burden for an individual or for the long-term care home’s resources. Given the small reductions in hip fractures achieved with hip protectors, older persons who are not at high risk may choose alternative options to prevent hip fractures. It is recognized that adherence to a recommendation to wear hip protectors may be challenging, and therefore strategies to improve adherence may be needed. This recommendation applies to hard and soft hip protectors, and the choice between them may depend on preference.

Summary of the evidence

Moderate-quality evidence from systematic reviews showed a relative risk reduction in hip fractures of 18% (95% CI 0% to 33%) among older persons wearing hip protectors in institutional settings.58 Over one year, four fewer hip fractures (95% CI –7 to 0) per 1000 older persons wearing hip protectors may be likely, and among older persons at higher risk, 11 fewer (95% CI –20 to 0) per 1000. However, one more pelvic fracture (95% CI 0 to 4) per 1000 older persons not at high risk and eight more (95% CI –3 to 30) per 1000 older persons at high risk may be likely. Moderate-quality evidence also showed that there is probably little or no difference in the frequency of falls or adverse events requiring medical attention, and that minor adverse events, such as skin irritation, occurred in less than 2% of people wearing hip protectors (soft or hard). The effect on quality of life and mortality is uncertain, and data for pain, anxiety, mobility and performance of activities of daily living were not available. Adherence to hip protector use varied across studies, from 24% to 80%. The impact of adherence on the effects of hip protectors is unclear, but the effects observed may represent the true effects when this strategy is implemented.

Exercise

For residents who are not at high risk of fractures, we suggest balance, strength and functional training exercises to prevent falls (conditional recommendation; moderate-quality evidence).

This recommendation places a high value on the probable small reduction in falls that is achieved with exercise, as falls may lead to serious injuries. It also places a high value on the other benefits that exercise could provide and a lower value on the uncertain costs to implement exercise interventions in long-term care settings.

For residents who are at high risk of fractures, we suggest balance, strength and functional training exercises only when such exercises are part of a multifactorial intervention to prevent falls (conditional recommendation; low-quality evidence).

This recommendation places a high value on avoiding the small increase in falls that may occur among individuals at high risk of falls who participate in exercises, such as balance, strength and functional training. Some older persons may value exercising despite the potential risk of falls. When exercise is made available to residents, it should be provided as part of a multifactorial intervention to prevent falls (including a review of medications [e.g., using the Beers criteria59], assessment of environmental hazards or use of assistive devices) or with other interventions to prevent fractures (including vitamin D and calcium supplementation, hip protectors and pharmacologic therapies).

Summary of the evidence

These recommendations are based on systematically reviewed evidence that is of moderate to low quality.31 This review included subgroup analyses for older persons in high-level care and intermediate-level care facilities, which were used to inform the recommendations for those at high risk and not at high risk of fractures, respectively. Most studies did not measure fractures, quality of life, mobility or pain. Instead, the risk of falls was used to inform this recommendation. Costs were not reviewed.

Subgroup analyses for high-level versus intermediate-level care among older residents at high risk of fractures suggested increases in the number of falls (870 more [95% CI –210 to 2370] per 1000 older people) and the number of older persons falling (85 more [95% CI –20 to 210] per 1000). Among older residents not at high risk of fractures, the analyses suggested reductions in the number of falls (660 fewer [95% CI –1290 to 390] per 1000 older persons) and the number of older persons falling (20 fewer [95% CI –115 to 105] per 1000). These results were from studies that evaluated balance training (such as tai chi), strength training and functional training. One study measured hip fractures, but the results were uncertain because there were very few events. A systematic review of exercise as part of a multifactorial intervention to prevent falls showed that the multifactorial intervention might reduce falls and the number of hip fractures.31
Multifactorial interventions

For all residents, we suggest multifactorial interventions that are individually tailored to reduce the risk of falls and fractures (conditional recommendation; low-quality evidence).

Multifactorial interventions are defined as any combination of interventions to reduce falls that are tailored to an individual’s risk. These interventions may include medication reviews (e.g., using the Beers criteria), assessment of environmental hazards, use of assistive devices, exercise, management of urinary incontinence and educational interventions directed to staff. This recommendation is conditional because of the low-quality evidence for important but small benefits and the unknown and potentially greater costs to implement multifactorial interventions in long-term care. A high value was placed on the small reductions in falls that may occur, as falls may lead to serious injuries. We have not suggested which interventions should be part of a multifactorial intervention, as it is unclear which combination of strategies provides benefit. It will be important to consider the resident’s level of fracture risk and tailor strategies accordingly.

Summary of the evidence

The evidence from a systematic review of interventions to prevent falls in older people in care facilities was of low quality because of the risk of bias of the included studies and the moderate to high inconsistency of effects across studies, which could not be explained by the level of care, cognition or combination of interventions. Most studies did not measure fractures, quality of life, mobility or pain; therefore, the risk of falls was used to inform this recommendation. Overall, the systematic review suggested reductions in the number of falls (660 fewer falls [95% CI –1230 to 120] per 1000 older persons per year) and the number of residents who fell (55 fewer residents falling [95% CI –115 to 10] per 1000) with the application of multifactorial interventions. There was low quality of evidence for a reduced risk of hip fractures (10 fewer [95% CI –14 to 1] per 1000). There were insufficient data to explore the effects of different combinations of interventions, or specific interventions, and their human and financial costs.

Implementation

An overview of the implementation of these recommendations appears in Figure 1. As advocated in the American Geriatrics Society’s “Guiding Principles for the Care of Older Adults with Multimorbidity,” the preferences of residents and their caregivers must be incorporated into care. Tools that will enable organizational leaders and care teams to incorporate the recommendations into their organizational policies, structures and care processes are being developed in partnership with stakeholders (Osteoporosis Canada, www.osteoporosis.ca; Ontario Osteoporosis Strategy for Long-Term Care, www.osteoporosislongtermcare.ca). The recommendations will be reviewed as new treatment options or new evidence becomes available that changes the effects described in this guideline.

Other guidelines

These recommendations developed using the GRADE approach are consistent with recommendations for fracture prevention in residential aged care facilities in Australia and with the expert consensus recommendations of AMDA – The Society for Post-Acute and Long-Term Care Medicine (formerly known as the American Medical Directors Association). However, osteoporosis therapies available for residents at high risk of fracture vary among countries.

The optimal dosage of vitamin D supplementation for older adults has been a topic of debate within both the American Geriatrics Society and the US Institute of Medicine. Although there is agreement about evidence that a dose of 800 IU will reduce fractures, there is less agreement about serum levels required for optimal bone health and the dosage required to achieve that level for the greatest proportion of older adults. Given the challenge of meeting vitamin D requirements from dietary sources and sunlight and evidence regarding the safety of vitamin D, Osteoporosis Canada has endorsed the position that individuals at high risk of vitamin D deficiency can safely take a supplement up to 2000 IU daily.

Gaps in knowledge

These recommendations were developed after consideration of the best available evidence. In most instances, the evidence was assessed to be of moderate or low quality, and further research may therefore change the estimates of effects and our confidence in those estimates. In particular, the evidence regarding fracture reduction achieved with pharmacologic therapies was based on effects observed among 100,000 postmenopausal, primarily weight-bearing women at high risk of fracture and was therefore assessed as indirect and downgraded because it may not be directly applicable to residents in long-term care. Further research is urgently needed to determine the efficacy and safety of osteoporosis therapies when administered to residents with limited mobility and multiple comorbidities. Future research initia-
Figure 1: Fracture prevention for residents of long-term care facilities. CrCl = creatinine clearance, IU = international units, IV = intravenous.
Guidelines


8. Crilly RG, Tanner DA, Kloseck M, et al. Hip fractures in long-term care and ultimately to maximize opportunities for quality living among long-term care residents. This guideline provides recommendations for the use of both nonpharmacologic and pharmacologic approaches to reduce fractures while considering residents’ multimorbidities and life expectancy.

References


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Conclusion

The goals of fracture prevention are to prevent pain, loss of mobility, serious injury and transfers to acute care and ultimately to maximize opportunities for quality living among long-term care residents. This guideline provides recommendations for the use of both nonpharmacologic and pharmacologic approaches to reduce fractures while considering residents’ multimorbidities and life expectancy.


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Affiliations: Department of Medicine (Papaioannou, Adachi, Pickard), Department of Clinical Epidemiology and Biostatistics (Papaioannou, Santesso) and School of Nursing (Kaasalainen), Faculty of Health Sciences, McMaster University, Hamilton, Ont.; Geriatric Education and Research in Aging Sciences Centre (Papaioannou, Pickard, Skadimore), St. Peter’s Hospital, Hamilton, Ont.; Department of Medicine (Morin), McGill University Health Centre, Montréal, Qué.; Montreal General Hospital (Morin), Montréal, Qué.; Department of Medicine (Feldman, Josse, Moser, Cheung) and Department of Physical Therapy (Jaglal), University of Toronto, Toronto, Ont.; Baycrest Geriatric Health Care System (Feldman, Katz, Moser), Toronto, Ont.; St. Joseph’s Healthcare (Adachi), Hamilton, Ont.; Division of Geriatric Medicine, Department of Medicine (Crilly), University of Western Ontario, London, Ont.; Department of Kinesiology (Giangregorio), University of Waterloo, Waterloo, Ont.; Institute for Clinical Evaluative Sciences (Jaglal), Toronto, Ont.; Li Ka Shing Knowledge Institute (Josse), St Michael’s Hospital, Toronto, Ont.; School of Dietetics and Human Nutrition (Weiler), McGill University, Montréal, Qué.; Department of Nutrition and Dietetics (Whiting), University of Saskatchewan, Saskatoon, Sask.; Toronto General Hospital (Cheung), Toronto, Ont.

Contributors: Alexandra Papaioannou and Nancy Santesso led the research group responsible for reviewing and synthesizing the evidence, developing the recommendations and drafting the manuscript. Nancy Santesso oversaw the application of the GRADE approach to guideline development and facilitated the guideline panel. All authors contributed to the discussion of issues related to care of older people and/or osteoporosis care, exercise and nutrition. All authors reviewed the final recommendations, and all reviewed and approved the final manuscript and agree to act as guarantors of the work.

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