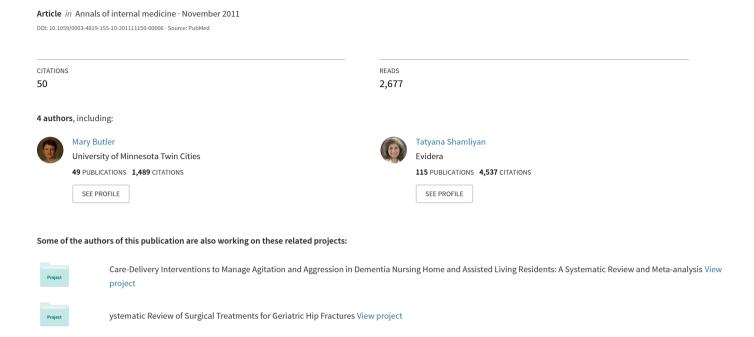
# Whole-Body Vibration Therapy for Osteoporosis: State of the Science



### **Annals of Internal Medicine**

# Whole-Body Vibration Therapy for Osteoporosis: State of the Science

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Clinical guidelines for osteoporosis recommend dietary and pharmacologic interventions and weight-bearing exercise to prevent bone fractures. These interventions sometimes have low adherence and can cause adverse effects. A proposed alternative or adjunctive treatment is whole-body vibration therapy (WBV), in which energy produced by a forced oscillation is transferred to an individual from a mechanical vibration platform. Whole-body vibration platforms are not approved by the U.S. Food and Drug Administration for medical purposes. This review provides a broad overview of important issues related to WBV therapy for prevention and treatment of osteoporosis. Relying on key informants and a search of the gray and published literature from January 2000 to August 2011, the

investigators found that the designs of WBV platforms and protocols for their use vary widely. The optimal target population for the therapy is not defined. Although WBV has some theoretical advantages, key informants have voiced several concerns, including uncertain safety and potential consumer confusion between low-intensity vibration platforms intended for osteoporosis therapy and high-intensity platforms intended for exercise. Finally, the scant literature did not establish whether WBV therapy leads to clinically important increases in bone mineral density or reduces risk for fracture.

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steoporosis contributes to risk for fracture, especially of the wrist, hip, and spine (1). It affects 2% of men and 10% of women older than 50 years in the United States (2). In addition, 49% of older women and 30% of older men in the United States have low bone density or osteopenia (2). Clinical guidelines for osteoporosis recommend dietary and pharmacologic interventions (3-7). Despite their proven effectiveness, these treatments have poor long-term adherence. In addition, drug treatments can have adverse effects, including esophageal irritation, minimal trauma atypical fractures, renal toxicity, and osteonecrosis of the jaw (8-14). Alternative therapies, including weight-bearing exercise, may be safer than drugs and may increase bone mineral density (BMD) (15, 16). Yet, not all older people can undertake high-intensity or weightbearing exercise. Furthermore, although exercise may improve certain outcomes related to bones and fracture risk, it has never been shown to prevent fractures in randomized, controlled trials (17-20). The U.S. Preventive Services Task Force encourages research on alternative osteoporosis interventions that may have better adherence and fewer side effects than available treatments (14, 21).

One proposed alternative is whole-body vibration (WBV) therapy (see Glossary) (22–26), which, like weight-bearing exercise, stimulates muscles and bones (2). In animal studies, vibration increases the anabolic (bone-building) activity of bone tissue, as well as bone area and

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density (27–30); however, the mechanism is not well-understood (31, 32). One hypothesis suggests that vibration signals directly activate mechanosensors in bone cells (33). Whole-body vibration, like weight-bearing exercise (34, 35), may improve muscle strength and power by increasing neuromuscular activation (36–41). In human studies of healthy volunteers, the effects of vibration therapy on muscle strength and performance were similar to those of short-term resistance exercise (34, 35). Whole-body vibration therapy might also improve muscle and bone blood circulation and increase the supply of nutrients needed to build bones (23, 42–46).

This report, commissioned as a technical brief by the Agency for Healthcare Research and Quality (AHRQ) Effective Health Care Program, provides an overview of WBV therapy for the prevention and treatment of osteoporosis. It describes available technology, identifies potential benefits and harms of the therapy, and summarizes published literature relevant to bone outcomes.

#### **METHODS**

#### **Key Informants**

We conducted semistructured interviews with 12 key informants via telephone or in person during December 2010. Key informants included osteoporosis experts, WBV experts, practicing clinicians who use WBV, consumer advocates, potential consumers, and representatives from manufacturers of WBV platforms. We identified key informants via frequently listed and cited authors of relevant literature, Internet searches for persons with potentially relevant viewpoints, and nominations by other key informants. The informants contributed information about the technology, the context in which it is used, and important issues about it that are worth considering.

#### Literature Scan

We searched several databases for published literature from January 2000 to August 2011: MEDLINE via Ovid and PubMed, the Cochrane Library, the Allied and ConWhole-Body Vibration Therapy for Osteoporosis: State of the Science | REVIEW

#### **Key Summary Points**

Whole-body vibration therapy is a proposed intervention for preventing and treating osteoporosis that is not approved by the U.S Food and Drug Administration.

The therapy involves a forced oscillation that transfers energy from a vibration platform to the person on the machine.

The mechanism by which whole-body vibration therapy may increase bone density is not well-understood.

Designs of whole-body vibration platforms and protocols for their use vary widely.

Neither the optimal target population nor the optimal treatment protocol is defined.

Although whole-body vibration therapy has some theoretical advantages and benefits, its efficacy and safety are unknown.

More research is needed to understand the role of this investigational therapy in preventing and treating osteoporosis.

temporary Medicine Database, CINAHL, the CSA Physical Education Index, Web of Science, the Physiotherapy Evidence Database, and Academic Search Premier. We based our search strategy on relevant Medical Subject Heading terms, text words, and a weighted word-frequency algorithm to identify related articles (Appendix Table 1, available at www.annals.org). We also conducted a gray literature search of federal government Web sites, the U.S. Food and Drug Administration (FDA) Web site, Clinical-Trials.gov, the CSA Physical Education Index, the Web of Science database, Medscape database, and the Internet by using different engines (including Google Scholar, Scirus, and LexisNexis).

Two individuals were involved in the literature review; one served as the primary reviewer, and one provided a quality check. We screened abstracts and full-text articles to identify studies published in English of any sample size and design that involved humans and reported any bone outcome, such as BMD or fracture. We excluded studies that involved children or healthy adults without low BMD or risk for osteoporosis, or patients with primary conditions, such as cerebral palsy, Parkinson disease, multiple sclerosis, cystic fibrosis, and spinal cord injuries. We also excluded studies that examined WBV as an exercise modality without reporting clinical bone measures, as well as market evaluations of vibration platforms. The reviewers read the full text of selected articles and used a standardized data extraction form to collect reported information about study populations, treatment protocols, and outcomes.

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#### **R**FSULTS

### WBV Platform Technology

In WBV therapy, forced oscillations produced by motors beneath a mechanical platform transfer energy to a human body on the machine (32). Available platforms produce sinusoidal-shaped oscillations (32). The International Society of Musculoskeletal and Neuronal Interactions has developed consensus criteria for describing sinusoidal vibrations, including the frequency, amplitude, peak-to-peak displacement, acceleration, and vibration platforms (synchronous, side-alternating, or triplanar) (see Glossary for definitions of the preceding terms) (47). Whole-body vibration platforms are meant to produce sinusoidal oscillations, but the actual oscillations may diverge from a pure sinusoidal shape. The vibrations transmitted to an individual may depend not only on the vibration variables but also on the individual's position on the platform and the rigidity of the platform plates (47).

Acceleration distinguishes acceptable vibration levels from those defined as hazardous by the International Organization for Standardization (ISO). Acceleration also determines whether WBV platforms are considered high-

#### Glossary

Acceleration: Maximal rate of change in velocity during an oscillation cycle. It is a function of the frequency and peak-to-peak displacement (expressed as m/s2) and is often expressed as multiples of Earth's gravity (9.80665 m/s<sup>2</sup>), denoted by the symbol g.

Amplitude: Maximal displacement from equilibrium position (expressed as

Frequency: Repetition rate of the cycles of oscillation (expressed as Hz). High-intensity vibration platforms: Platforms that provide acceleration greater than 1g.

Low-intensity vibration platforms: Platforms that provide acceleration less

Peak-to-peak displacement: Displacement from the lowest to the highest point of the vibrating platform position (expressed as mm).

Side-alternating whole-body vibration platforms: Platforms that use a reciprocating vertical displacement on the left and right sides of a fulcrum. Synchronous whole-body vibration platforms: Platforms on which the left and right feet move up and down simultaneously.

Triplanar whole-body vibration platforms: Platforms that oscillate in 3 planes. These platforms may also be described as "elliptical."

Whole-body vibration therapy: The mechanical repetitive movement, or oscillatory motion, around an equilibrium point. Whole-body vibration therapy involves a forced oscillation in which vibrations generated by motors beneath a platform are transmitted to the person on the machine. Currently available whole-body vibration platforms are intended to produce sinusoidal-shaped oscillations.

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Figure 1. Low-intensity whole-body vibration platform.



The Juvent 1000 platform. Photograph courtesy of Juvent, Dynamic Motion Therapy, Somerset, New Jersey.

intensity (see Glossary) or low-intensity (see Glossary); acceleration is less than 1g for the former and more than 1g for the latter.

The FDA has not approved WBV platforms for medical purposes: No FDA standards regulate their manufacture, and designs of platforms vary widely. Figures 1 and 2 show two examples. Some low-intensity platforms resemble bathroom scales in size and shape, except that they are raised several inches off the ground, whereas some larger high-intensity platforms resemble typical exercise machines. Some platforms have safety features, such as a handrail for balance.

Low-intensity vibration platforms for home use are currently marketed at a retail price of approximately \$1600. Some of these platforms automatically calibrate the treatment to each user's weight and body mass index. Suggested treatment sessions involve standing on the platform for 10 minutes daily. Manufacturers report that home use requires no direct supervision. Newer models are very low in height and offer an optional wheelchair mount (for example, see www.livtherapy.com/products/index.html). Advancements will soon allow individuals with mobility problems to use vibration platforms in a seated or supine position (for example, see http://vibetechglobal.com /prototype.aspx).

High-intensity vibration platforms produce a gravitational force greater than 1g regardless of frequency. Although they are marketed as exercise equipment, these platforms are used in clinical physical therapy or rehabilitation settings, exercise facilities, and the home. No organization provides accreditation or training for vibration therapy use in professional settings. Some exercise facilities provide proprietary training to personal trainers (for example, Power Plate [Irvine, California; www .powerplate.com]). Such training is specific to proper use of the platform in exercise programs, not to osteoporosis prevention or treatment.

#### Potential Benefits and Harms

Whole-body vibration therapy may offer advantages to individuals who cannot or do not want to continue or initiate pharmacologic treatment to increase BMD and those who cannot perform high-impact exercise. In addition, the relative ease of use of vibration therapy may result in better adherence.

Therapeutic and occupational vibration exposure presents safety concerns. Vibration has been recognized as an occupational hazard associated with low back pain (48, 49), musculoskeletal problems (50), cardiovascular disorders (51), neurovestibular disorders (52), and the Raynaud syndrome (53). The ISO has defined vibration limits for comfort, performance proficiency, and safety on the basis of their known occupational hazards, and ISO 2631-1 defined high-intensity vibrations (those that produce force greater than 1g) as hazardous regardless of their frequency (54). Safety concerns for vibration as a therapeutic intervention include the possibility of an individual losing contact with the vibration platform and becoming airborne when acceleration exceeds 1g. The resulting impact as the feet return to the platform may be harmful for individuals with fragile bones (32). Vibration may also be harmful to the soft-tissue organs of the head and chest. Furthermore, knee flexion and posture can alter the transmissibility of vibrations to the head and trunk. Thus, an individual's shifting of position on the platform may complicate accurate measurement of vibration in different body parts (32).

Key informants indicated that harms of WBV therapy may include plantar fasciitis, itchy legs, blurred vision, tinnitus, white-finger disease (a secondary form of the Raynaud syndrome), orthostatic hypertension, and aggravation of soft-tissue and joint injuries. Dislocation of an intraocular lens after cataract surgery also raises concerns, particularly because the population using whole-body vibration for osteoporosis prevention and treatment is at greater risk for cataracts (55). In addition, various parts of the body can resonate at different and highly individual frequencies. Inadequate understanding of optimal vibration dosage and transmission to different parts of the body could lead to injuries. Other concerns expressed by key informants in-

cluded loss of balance and falls during platform use and lack of clear distinction between platforms intended for powered exercise and those intended for osteoporosis therapy.

#### Contexts in Which WBV Is Used

Whole-body vibration platforms are used in the home, physical therapy or rehabilitation clinics, and exercise facilities. High-intensity platforms are marketed as powered exercise equipment. Although they may be used for medical purposes, such as muscle or joint rehabilitation, these high-intensity platforms are exempt from FDA premarket notification procedures (56). Manufacturers marketing low-intensity WBV platforms for treatment of osteoporosis or improvement of BMD include disclaimers on their Web sites, specifying that their devices are investigational and that they make no medical claims for osteoporosis (for example, see www.juvent.com and www.marodyne.com /technology). Many of these Web sites summarize or provide links to scientific research for potential consumers to review. Key informants expressed no awareness that any third-party payers cover the costs of WBV therapy; thus, individuals pay out of pocket for clinic sessions or to purchase platforms for home use.

#### **Current Evidence**

Of the 345 citations screened, 12 studies met the inclusion criteria (22, 37, 42, 57-65); these are summarized in Appendix Tables 2 to 7 (available at www.annals.org). The patient populations were individuals who were diagnosed with osteoporosis, had low BMD, or were at risk for low BMD or osteoporosis. Two studies focused on postmenopausal women diagnosed with osteoporosis (57, 63), and 3 focused on children and adolescents with low BMD (37, 58, 64). The remaining 7 studies evaluated individuals at risk for low BMD or osteoporosis; all but one of these evaluated postmenopausal women (22, 42, 59-62). The remaining study included 1 older male participant (65).

Six studies were randomized, controlled trials (22, 42, 60-63); 3 were nonrandomized, controlled trials (37, 57, 59); and 3 were case series (58, 64, 65). Control groups in the trials received no intervention, walking, resistance training or exercise, bisphosphonates, or a placebo device.

The WBV interventions in the studies involved synchronous, side-alternating, or triplanar platforms. Vibration frequencies ranged from 12 to 40 Hz across 11 studies; 1 study did not report frequency (65). Five studies had frequency settings that changed, during either individual sessions or the study period (42, 58-61). The amplitude ranged from 0.7 to 5 mm across the 7 studies that reported it (22, 57-61, 63); the amplitude setting changed during the intervention period in 1 study (60). Acceleration ranged from 0.1g to 10g across 6 studies that reported it (37, 42, 59, 60, 62, 64); 3 had levels less than 1g (37, 62, 64), whereas 3 had levels greater than 1g (42, 59, 60).

Vibration sessions ranged from 15 seconds to 30 minutes and took place 1 to 7 days per week. Studies with acceleration levels less than 1g had more frequent and generally longer sessions than studies with acceleration levels greater than 1g. The duration of the vibration intervention and follow-up ranged from 8 to 72 weeks.

In some studies, participants were instructed to do certain things while on the platform. In 4 studies, participants performed dynamic exercises or extended their lower extremities (58-61); in 4 studies, participants flexed their knees (22, 42, 58, 63); and in 3 studies, participants flexed their knees while performing exercises (59-61). Only 3 studies reported the type of footwear used during sessions (22, 59, 60), and 5 studies stated or illustrated that a support device was available on the platform (57, 58, 62, 64, 65).

Three studies evaluated WBV as an adjunctive therapy; 2 examined it as an adjunct to an exercise program (59, 61), and 1 as an adjunct to drug therapy (63). Two studies required participants to have adequate nutritional intake (22, 61), and 5 studies either provided or advised participants on appropriate levels of vitamin D or calcium (or both) (37, 42, 59, 61, 63).

Of the 11 studies that measured BMD (22, 37, 42, 57, 59-65), 8 did so only with dual-energy x-ray absorptiometry (DXA) (22, 57, 59-63, 65), 2 did so only with computed tomography (42, 64), and 1 used both DXA and computed tomography (37). Location of the BMD mea-

Figure 2. Whole-body vibration platform that produces side-alternating vibration.



The Osci Health platform. Image courtesy of Health Mark, Acworth,

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surements included the femoral neck, lumbar spine (L1 to L4), total body, total hip, trochanter, forearm, and tibia. Overall, the trials found statistically nonsignificant or small increases in BMD of 1.5% to 4.3% at specific sites in treatment groups compared with controls. Statistically significant increases in BMD were found at the femoral neck in 2 studies (22, 57), total hip in 1 study (60), and lumbar spine in 1 study (57). One study found statistically nonsignificant increases in BMD for treatment participants compared with controls in an intention-to-treat analysis, but it also found that participants in the 3 highest quartiles of treatment adherence had statistically significant increases in BMD of the lumbar spine compared with controls or those in the lowest quartile of adherence (37). One study reported a statistically significant decrease in BMD at the forearm (radius, 33%) for treatment participants compared with controls (59).

Two studies reported data on fractures (58, 63). One of these found no new thoracic or lumbar vertebral fractures or nonvertebral osteoporotic fractures of the hip, wrist, or shoulder in any treatment or control participant over the 12 months of the intervention (63). Two studies reported minor harms from vibration (42, 58). Eleven studies evaluated other outcomes, including bone mineral content; bone turnover markers; falls; balance; mobility; back pain; postural control; bone area; muscle force, strength, power, mass, and area; fat mass; adherence to the study protocol; and efficacy of device use (22, 37, 42, 57-64).

#### **DISCUSSION**

We found scant evidence on the benefits and harms of WBV for the prevention and treatment of osteoporosis. Key informants unanimously urged caution in making claims about this intervention because of inadequate evidence and knowledge about the appropriate target population, optimal protocol features, safety of platforms, and long-term beneficial or harmful effects. Several key informants indicated that WBV therapy should not replace osteoporosis therapies but warrants evaluation as a possible adjunctive therapy.

We are not certain which groups might benefit from or be more susceptible to harms from WBV. Neither the literature nor our key informants offered clear guidance on the optimal target population. Also lacking was evidence for how this therapy might affect individuals with different levels of risk for or severity of osteoporosis. Potential harms are unknown for individuals with health issues, such as heart or musculoskeletal problems, because most published studies excluded them.

The 12 published studies that we reviewed included diverse treatment protocols, reflecting uncertainty about the most efficacious platform type and settings (frequency, amplitude, and acceleration), and session length and frequency. Few studies systematically assessed potential harms or clinically important outcomes, such as fractures. Trials had small samples and short follow-up, and they generally reported statistically nonsignificant or small increases in BMD that were of uncertain clinical significance.

Several key informants indicated that bone outcomes, including fractures, are of the most interest for future studies that evaluate the effectiveness of WBV therapy. Because fractures are rare outcomes, and changes in bone outcomes occur slowly, measuring these would require a large number of participants and long follow-up. Measurement techniques also influence the detectability of changes in bone outcomes. For instance, peripheral quantitative computed tomography measures volumetric BMD, whereas DXA measures areal BMD, and the former technique may offer better accuracy and diagnostic information than DXA (66). Many key informants also suggested muscle strength and balance as potentially valuable intermediate outcomes, because improvements in these functions might prevent fractures and falls. Finally, quality-of-life measures could provide valuable information, but like fracture, these would require long-term studies.

Informants noted adherence and access as additional key considerations. Both the site (clinic or home) and the frequency of sessions may affect long-term adherence. The 8 studies that measured adherence found a wide range, raising questions about long-term adherence for various treatment protocols (22, 37, 42, 58, 59, 61, 62, 64). Site and frequency of sessions may also affect treatment access. Currently, third-party payers do not cover WBV devices, so consumers must pay out of pocket for sessions in a clinical setting or to purchase a platform. Out-of-pocket costs may affect whether, where, and how often a consumer uses a WBV platform.

Finally, safety features must be considered. Individuals using WBV platforms may be at risk for falls because of balance problems or disorientation. Decreased blood pressure may occur during platform use. Not all platforms provide safety features, such as handrails, to address these issues. Direct-to-consumer marketing also causes problems; consumers may not be able to clearly distinguish lowintensity platforms intended for osteoporosis therapy from platforms intended for high-intensity exercise.

Our review has limitations. We relied on a limited number of key informants. We included only studies published in English that reported a bone outcome and that involved participants with either risk for or diagnosed osteoporosis or low BMD. Finally, we did not formally evaluate each study's quality or risk of bias.

In summary, WBV therapy for the prevention and treatment of osteoporosis is an investigational therapy. We know little about its benefits and harms. Although several clinical trials of this therapy are completed or ongoing (67– 69), much additional research is needed to fully understand the role of this therapy.

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#### Appendix Table 1. Search Strategy

#### Preliminary literature search

MEDLINE, Cochrane Library, CINAHL, CSA Physical Education Index, Web of Science, PEDro, and Academic Search Premier databases using the keywords "whole body vibration," "vibration," and "osteoporosis."

#### Ovid Technologies Email Service, 1950 to week 4 of August 2010

Search for: 18 not 19 Results: 120

Ovid MEDLINE	Citations Retrieved, n
1. exp Vibration/tu [Therapeutic Use]	511
2. whole body.mp.	39 402
3. 1 and 2	71
4. exp Muscle Strength/	10 075
5. exp "Recovery of Function"/	19 156
6. 4 or 5	28 640
7. 1 and 6	27
8. 3 or 7	85
9. wbv.mp	309
10. 1 and 9	36
11. 8 or 10	85
12. exp Muscle, Skeletal/	165 830
13. 1 and 12	65
14. 11 or 13	114
15. exp Physical Therapy Modalities/	99 346
16. 1 and 15	206
17. 14 or 16	278
18. limit 17 to (English language and humans and yr="2000-Current")	127
19. limit 18 to (case reports or editorial)	7
20. 18 not 19	120
21. exp Osteoporosis/rh, th [Rehabilitation, Therapy]	2609
22. 1 and 21	14
ubMed search strings	
Search "Vibration/therapeutic use" [MAJR] Limits: Humans, Randomized Controlled Trial, English	68
Search "Vibration/therapeutic use" [MAJR] Limits: Humans, Journal Article, English	1287
Search "Vibration" [Mesh] Limits: Humans, Journal	6541

#### Cochrane Library

Article, English

Journal Article, English Search vibration AND osteoporosis

Whole body vibration for preventing and treating osteoporosis (Protocol)

71

119

Search vibration AND osteoporosis Limits: Humans,

#### CINAHL

212 references retrieved

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Appendix Table	2. Studies of Who	e-Body Vibration for Osteoporosi	Appendix Table 2. Studies of Whole-Body Vibration for Osteoporosis: Inclusion Criteria, Study Sample Characteristics, and Receipt of Previous Osteoporosis Therapy	tics, and Rece	ipt of Previou	s Osteoporosis	Гһегару
Study, Year (Reference)	Country; Study Design	Patient Inclusion Criteria	Patient Exclusion Criteria	Men Included	Minorities Included	Patients With Comorbid Conditions Included	Previous Treatment for Osteoporosis
Gusi et al, 2006 (22)	Spain; RCT	Women; ≥5 y from last menstruation; adequate nutritional status according to World Health Organization (determined by questionnaire); nonsmoker; consumed ≤4 alcoholic beverages per week; could follow the protocol; no disease or medication known to affect bone medaciation known to affect bone metabolism or muscle strength	Acute hernia; thrombosis; any pharmacologic intervention for osteopenia in the previous 6 mo; any history of severe musculoskeletal problems; engaged in high-impact activity at least twice a week (any weight-bearing activity or exercise more intense than brisk walking)	° Ž	χ Z	ž	No; diet/calcium intake unknown
Ruan et al, 2008 (57)	China; controlled trial	Women with osteoporosis; postmenopausal, without ppical menopausal symptoms; age =80 y; willing to participate as volunteers	Blood pressure >160/110 mm Hg despite medical thrapy, ystolic blood pressure <90 mm Hg, heart disease or cerebrovascular disease; epilepsy, thrombosis or a history of thrombosis in the past 6 mo; body implants or heart stents; lumbar disc herniation or spondyolisthesis; spinal nerve canal stenosis or oppression; poor health and symptoms of imbalance or vertigo; treatment with drugs for osteoporosis or other agents affecting bone metabolism; not recovered from surgery; not recovered from joint injuries, fractures, or muscle strain	S.	X Z	X.	No; prior physical activity and dieVcalcium intake unknown
Semler et al, 2008 (58)	Germany; case series	Motor-impaired children with osteogenesis imperfecta	¥Z.	Yes	Z Z	Z.	Yes—bisphosphonates; diet/calcium intake unknown
Bemben et al, 2010 (59)	United States; controlled trial	Healthy women volunteers aged 55–75  Yi = 54 postmenopausal; not taking HRT; previous HRT users had not received it for at least 1 y; had not participated in a weight-training program for at least 1 y; medically stable, ambulatory, and capable of undergoing physical strength testing and training; were able to give written informed consent and adhere to the protocols	Diagnosed osteoporosis or a BMD site with a T-score less than -2.5; physical disabilities that prevented persons from being strength-tested and -trained, including orthopedic or arthritic problems; heart problems (e.g., congestive heart failure and arrhythmias), chronic high blood pressure, or receipt of fa-blockers; current smokers or past smokers in the previous 15 y; current smokers or past smokers in the previous 15 y; current diagnosis or a history of renal disease, chronic digestive or eating disorders, theumatoid arthritis, or uncontrolled thyroid disease; use of medications that affect bone dersity (e.g., steroid hormones, calcitonin, or corticosteroids); use of medications for osteoporosis treatment, including bisphosphonates, SERMs, or parathyroid hormone	2	Z	<del>Z</del>	No; baseline calcium and physical activity recorded
Verschueren et al, 2004 (60)	Belgium; RCT	Women aged 60–70 y; noninstitutionalized; free of diseases or medications known to affect bone metabolism or muscle strength	Total body BMD T-score less than –2.5	o N	N N	X.	No; prior physical activity and diet/calcium intake unknown
von Stengel et al, 2011 (61)	Germany; RCT	Postmenopausal women aged ≥65 y living independently in the community were contacted by mail (mailing lists were obtained from the Siemans Health Insurance Company database)	Diseases or medication affecting bone metabolism; diseases or medication affecting neuromuscular performance and falls; implants of the lower extremity or of the spine, eye diseases affecting the retina; low physical capacity (<50 W)	ON.	Z Z	¥	No: baseline calcium, vitamin D., and physical activity recorded

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Appendix Table 2—Continued	2—Continued						
Study, Year (Reference)	Country: Study Design	Patient Inclusion Criteria	Patient Exclusion Criteria	Men Included	Minorities Included	Patients With Comorbid Conditions Included	Previous Treatment for Osteoporosis
Rubin et al, 2004 (62)	United States; RCT	Normal nutritional status as determined by questionnaire; stable weight maintenance (no elective weight loss or diet); estimated daily calcium intake ≥500 mg; could follow the protocol for daily use of the device and understand and provide informed consent, body mass >45 kg and <84 kg (owing to design constraints of the oscillating device)	Any pharmacologic intervention for osteopenia in the previous 6 mo; any use of steroids; current smoker, excessive alcohol consumption (>2 drinks/d), evidence of osteomalacia; Paget disease; osteogenesis imperfecta; gastrointestinal disease, history of cancer, prolonged immobilization of the axial or appendicular skeleton in the past 3; evidence of spondyloarthosis, thyrotoxicosis, psychomotor disturbances, thyperparathyroidism, renal or hepatic disease, and chronic diseases known to affect the musculoskeletal system, or engaged in high-impact activity at least 3 times/wy.	°Z	Ϋ́ Z	ž	No; minimum calcium intake
Russo et al, 2003 (42)	Italy; RCT	Women belonging to a hospital volunteers association, =1 y postmenopausal; not affected by conditions that contraindicated the vibration training still eligible if receiving HRT	Women with metabolic bone disorders	O <sub>Z</sub>	N N	R	NR; minimum calcium and vitamin D intake
lwamoto et al, 2005 (63)	Japan; RCT	Hospitalized postmenopausal women aged 55–89 with osteoporosis (BMD T-score <70% or 70%–80% with a history of osteoporotic fractures); chronic back pain that did not require bed rest treatment; no history of HRT or receipt of medication that affects bone metabolism; no receipt of such medication as NSAIDs to relieve chronic back pain; instructed to take 800 mg of calcium daily in food	Osteoarthritis of the knee; moderate to severe spondylosis or degenerative disc disease of the thoracia and lumbar spine, musculoskeletal diseases other than osteoporosis that cause back pain; arthroplasty of the knee or hip joint	<del>2</del>	α Z	ž	No; all participants had low physical activity, diet/calcium intake unknown
Gilsanz et al, 2006 (37)	United States; controlled trial	Healthy white females aged 15–20 y who had perviously had at least 1 fracture; completed puberty (ranner stage V of sexual development); of the 150 potential candidates, the 50 females with the lowest CT values for vertebral cancellous BMD (~1 SD below mean peak BMD values) were invited to participate in the intervention phase	Any underlying disease or chronic illness, had been till for >2 wk in the previous 6 mo, had been admitted to the hospital at any time in the previous 3 y; receipt of any medications, including oral contraceptives; pregnant, had ever been pregnant, or had had menses for >4 consecutive mo or 2 cycle lengths after establishing regular cycles; epiphyses of the phalanges and the metacarpels had not fused completely	<u>S</u>	°Z	2	No; baseline physical activity and calcium intake recorded
Pitukcheewanont and Safani, 2006 (64)	United States; case series	Girls diagnosed with endocrine disorders of various causes and low bone density. Tanner stage 1 or 11 of sexual development	Any medication known to affect bone density	ON N	N R	Z.	No; physical activity and diet/calcium intake unknown
Ezenwa et al, 2008 (65)	United States; case series	Age ≥65 y; able to go from sitting to standing without assistance, walk up and down 3 steps; ambulate 50 ft with or without a cane and without shortness of breath or chest pain	Any medical condition that might affect BMD (e.g., bone cancer, ESRD, long-term steroid use); other neurologic conditions affecting balance and strength (e.g., history of stroke, Parkinson disease, vertigo)	Yes	۳ ک	ZZ Z	Z.

BMD = bone mineral density; CT = computed tomography; ESRD = end-stage renal disease; HRT = hormone replacement therapy; NA = not available; NR = not reported; NSAID = nonsteroidal anti-inflammatory drug; RCT = randomized, controlled trial; SERM = selective estrogen receptor modulator.

# Appendix Table 3. Studies of WBV for Osteoporosis: Setting, Participants' Living Arrangements, and WBV Platforms and Manufacturers

Study, Year (Reference)	Setting of Intervention	Living Arrangement of Participants	WBV Platform Type	WBV Platform Model and Manufacturer
Gusi et al, 2006 (22)	Clinic (assumed)	Community	Side-alternating	Galileo 2000, Novotec, Pforzheim, Germany
Ruan et al, 2008 (57)	Clinic	Community (campus of Beijing Institute of Technology)	Synchronous	ZD-10, Beijing Maidakang Medical Equipment Company, Beijing, China
Semler et al, 2008 (58)	Home	Community	Side-alternating	Cologne Standing and Walking Trainer System, Galileo (modified tilt-table combined with the Galileo WBV system)
Bemben et al, 2010 (59)	Clinic	Community	Triplanar	Power Plate North America, Northbrook, Illinois
Verschueren et al, 2004 (60)	Clinic	Community	Not reported	Power Plate, Amsterdam, the Netherlands
von Stengel et al, 2011 (61)	Clinic and home	Community	Synchronous	Vibrafit, Solms, Germany
Rubin et al, 2004 (62)	Home	Community	Synchronous	Model LA18-18, BEI, San Marcos, California
Russo et al, 2003 (42)	Clinic	Community	Side-alternating	Galileo 2000
Iwamoto et al, 2005 (63)	Clinic	Community	Side-alternating	Galileo
Gilsanz et al, 2006 (37)	Home	Community	Synchronous	Not reported explicitly
Pitukcheewanont and Safani, 2006 (64)	Clinic	Community	Synchronous	Model LA18-18, BEI
Ezenwa et al, 2008 (65)	Clinic	Community	Not reported	Developed for study

WBV = whole-body vibration.

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During certain exercises

8

2 2 R

Bend and straighten knees while on platform During certain exercises

Extend Lower Extremities While on Platform

	Flex Knees	Platform	Yes	° N	Yes	During certain exercises	During certain exercises	During certain exercises	o N	Yes	Yes	NR	Z Z
	Changes in Platform Settings	During Intervention Period	O Z	OZ.	Yes: changes in frequency and tilting angle	Yes: changes in frequency, acceleration, session length, and sets	Yes: changes in duration of session, number of series of 1 exercise, number of different exercises, amplitude, and frequency	Yes: changes in frequency and exercise intensity	No	Yes: change in frequency and session length	No	No	No
Icipants	Changes ii	During Session	O Z	ON	Yes: changes in frequency	O <sub>N</sub>	O Z	No	No N	No	No	No	No
		Session Rest Periods	1-min vibration, 1-min rest	ON.	3-min vibration, 3-min rest	15-s rest between sets	Yes	1-min break with stretching between exercises	At least 10 h between 2 daily sessions	1- or 2-min vibration (3 sets), 1-min rest between	No	No	O Z
	Characteristics	Session Length	First 2 wk, 3 min; last 6 wk, 6 min	10 min	9 min twice daily	15- to 60-s sessions with 1–3 sets	30 min, which included warming up and cooling down	6 min	10 min, 2 times per day	First 1 mo, 3 min; last 5 mo, 6 min	4 min	10 min	30 min
	WBV Intervention Characteristics	Acceleration	<u>م</u> 2	N N	χ α	2.16–2.8g (acceleration magnitude)	2.28–5.09g (peak acceleration)	N N	0.2g (peak to peak)	0.1–10g (acceleration)	Z Z	0.3g (peak to peak)	0.3g (vertical acceleration)
		Amplitude	3 mm (vertical amplitude)	5 mm (amplitude)	1–2 mm (amplitude)	2–4 mm (peak to peak)	1.7–2.5 mm (amplitude)	1.7 mm (amplitude)	NR	N N	0.7–4.2 mm (upward and downward)	NR	N N
		Frequency	12.6 Hz	30 Hz	15–25 Hz	30–40 Hz	35-40 Hz	25–35 Hz	30 Hz	12–28 Hz	20 Hz	30 Hz	30 Hz
	Study, Year		Gusi et al, 2006 (22)	Ruan et al, 2008 (57)	Semler et al, 2008 (58)	Bemben et al, 2010 (59)	Verschueren et al, 2004 (60)	von Stengel et al, 2011 (61)	Rubin et al, 2004 (62)	Russo et al, 2003 (42)	Iwamoto et al, 2005 (63)	Gilsanz et al, 2006 (37)	Pitukcheewanont and Safani, 2006 (64)

NR = not reported; WBV = whole-body vibration.

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5. Studies of	
Appendix Table	

Study, Year (Reference)	Footwear Worn on Platform	Support Device on Platform	Intervention Frequency	Intervention Duration, wk	Combination of Treatments	Concomitant Treatments for Osteoporosis	Calcium Supplementation	Length of Follow-up, wk
Gusi et al, 2006 (22)	Barefoot	Z Z	3 d/wk; ≥1 d of rest between sessions	32	No, but WBV program included warm-up with 5 min of bicycling and 5 min of stretching	No medications at start	° Z	32
Ruan et al, 2008 (57)	Z Z	Yes	5 times/wk	24	No	No bone medications at start	No	24
Semler et al, 2008 (58)	X X	Yes: patients were strapped to the tilt-table	7 d/wk	24	No	Yes: medications and physiotherapy continued	No	24
Bemben et al, 2010 (59)	Shoes while standing; also sat on platform	NR T	3 d/wk	32	Yes: WBV (which included dynamic movements) and resistance training	No bone medications at start	No, but participants were instructed to increase calcium intake if less than 1500 mg/d	32
Verschueren et al, 2004 (60)	Shoes	NR N	3 d/wk; ≥1 day of rest between sessions	24	No, but WBV program included exercise on platform, warm-up, and cool-down	No bone medications at start	o Z	24
von Stengel et al, 2011 (61)	Z Z	Z Z	2 clinical, 2 home	72	Yes: WBV and training	No bone medications at start	Yes, for participants who needed it to ensure 1500 mg of calcium and 400 IU of vitamin D daily	72
Rubin et al, 2004 (62)	W Z	Yes	7 d/wk	48	No	No bone medications at start	No	48
Russo et al, 2003 (42)	NR	N.	2 d/wk	24	O N	NR if medications taken	Yes: all participants received 1 g of calcium and 0.25 μg of vitamin D daily	24
Iwamoto et al, 2005 (63)	<u>«</u> ک	N N	1 d/wk	48	Yes: WBV and alendronate (5 mg/d)	Yes: medication	No, but participants were instructed to get 800 mg in food daily	48
Gilsanz et al, 2006 (37)	N N	NR	7 d/wk	48	0 Z	No medications at start	Yes: all participants took a 500-mg calcium tablet daily	48
Pitukcheewanont and Safani, 2006 (64)	W Z	Yes	3 d/wk	∞	N <sub>o</sub>	No bone medications at start	No	<b>∞</b>
Ezenwa et al, 2008 (65)	W.Z.	Yes	3 times/wk	20	No	NR if medications taken	ON	20

NR = not reported; WBV = whole-body vibration

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Ruan et al, 2006 (22)  Reported rate  Reported high adherence 2008 (58)  Semler et al, 2008 (58)  Been self-report)  Bemben et al, 2010 (59)  Verschueren et al, 2004 (60)  Von Stengel et al, 2011 (61)  Russo et al, 2003 (42)  Ves: reported rate and assessed outcomes by adherence  Russo et al, 2003 (42)  NB  Seported rate and assessed outcomes by adherence  Russo et al, 2003 (42)  NB	ate		Number of Comparators	parators	Measured	חואום ואוכמסית	
,	ate		Intention-to-Treat Analysis	Completed	Measured	Technique	Site
		Walking program training group	18	41	Yes	DXA	Right proximal femur (femoral neck, trochanter, and Ward triangle); lumbar spine
\$\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\		Control group (no program)	50	43	Yes	Dual-energy bone densitometers	Lumbar L2-L4; femoral neck
\$	adherence report (may have oort)	· «	<b>₹</b> Z	Ϋ́ Y	No	₹ Z	VΑ
Z	ate	Resistance training group and control group (no program)	Unknown	22 and 12	Yes	DXA	Total body; AP lumbar spine L1–L4; dual proximal femur (femoral neck, trochanter, total hip); 33% radius of forearm
» » » <u> </u>		Resistance training group and control group (no program)	22 and 23	22 and 23	Yes	DXA	Total hip; total body; lumbar spine
× × <del>Z</del>	ate	Exercise training group and wellness program control group	50 and 51	47 and 48	Yes	DXA	Lumbar spine L1–L4; proximal femur
	ate and comes by	Placebo device control group	37	28	Yes	DXA	Proximal right and left femora: neck, trochanter, lumbar spine, distal one third of nondominant radius
	ate	Control group (no program)	17	41	Yes	рост	Tibia: trabecular and cortical
		Alendronate only (bisphosphonate) control group	25	25	Yes	DXA	Lumbar spine L1–L4
Gilsanz et al, Yes: reported rate and 2006 (37) assessed outcomes by adherence	ate and comes by	Control group (no program)	25	24	Yes	CT; DXA	Lumbar spine L1–L3; total body
Pitukcheewanont and Yes: reported all Safani, 2006 (64) participants completed study	all completed	NA N	NA	<b>∀</b> Z	Yes	J	L1–L3 of lower axial spine (cancellous); femurs (cortical)
Ezenwa et al, NR 2008 (65)		NA	NA	۷ ۷	Yes	DXA	Lumbar spine L1–L4; total hip; femoral neck; trochanter; forearm

AP = anteroposterior, BMD = bone mineral density; CT = computed tomography; DXA = dual-energy x-ray absorptiometry; NA = not applicable; NR = not reported; pQCT = peripheral quantitative computed tomography.

Appendix Table 7. Studies of Whole-Body Vibration for Osteoporosis: Bone Mineral Content, Fractures, Quality of Life, Harms, and Other Outcomes

			ssessment of Motor igle to calculate ce and measure iscle force	one turnover markers from blood samples (CTX for bone resorption and bone ALP for bone formation); muscle strength	one turnover markers from blood samples (serum osteocalcin for bone formation and CTX for bone resorption); muscle strength (isometric, dynamic); fat mass and muscle mass;		of device use; bone rption through mples		X, serum ALP, horus; falls		iscle mass; bone ALP	
Other Outcomes	Balance; BMI	Chronic back pain	Mobility (with Brief Assessment of Motor Function); tilting angle to calculate ground reaction force and measure improvement in muscle force	Bone turnover markers from blood samples (CTX for bone resorption bone ALP for bone formation); n strength	Bone turnover markers from blood samples (serum osteocalcin for bone formation and CTX for bone resorption); muscle strength (isometri dynamic); fat mass and muscle mass; postural control	Falls	Adherence; efficacy of device use; bone formation and resorption through serum and urine samples	Muscle power	Back pain; urinary NTX, serum ALP, calcium, and phosphorus; falls	Muscle area	Fat mass; femoral muscle mass; bone area; bone-specific ALP	٩
Harms	None reported	None reported	Yes: some patients reported itching after vibration session; 1 patient reported localized pain at the end of an intramedullary rod that had been dislocated before the vibration intervention; 1 patient withdrew from the study after dislocation of a telescopic rod (which had happened before in this patient)	None reported	None reported	None reported	None reported	Yes: transient lower-leg itching and erythema in 6 of 17 patients; moderate knee pain in 2 overweight participants with preexisting knee osteoarthritis	None reported	None reported	None reported	None reported
Quality of Life Measures	۷ ۷	Y Y	₹ Z	Α	₹ Z	NA V	₹ Z	Υ V	Y Y	NA V	<b>∢</b> Z	₹ Z
Quality of Life	° Z	No	9 Z	° N	O <sub>N</sub>	No No	°Z	<u>8</u>	No	No No	o N	o N
Measure of Fracture	₹Z	Ϋ́	Count	Y X	₹ Z	NA V	۲ ۲	NA	Radiography	NA	₹ Z	<b>∢</b> Z
Fractures	° Z	No	≺es	o N	O Z	No	o N	ON.	Yes	No	o N	N <sub>O</sub>
Bone Mineral Content Measures	<b>∢</b> Z	ΝΑ	<b>﴿</b> 2	NA	<b>∢</b> Z	NA	۷ ۲	Y.	V V	DXA	<b>∀</b> Z	<b>∢</b> Z
Bone Mineral Content	ON.	No	2	ON.	<u> </u>	No No	O Z	No.	No No	Yes	O <sub>N</sub>	O <sub>N</sub>
Study, Year B( (Reference) C	Gusi et al, N 2006 (22)		Semler et al, N 2008 (58)	Bemben et al, N 2010 (59)	Verschueren et al, N 2004 (60)	von Stengel et al, N 2011 (61)	Rubin et al, N 2004 (62)	Russo et al, N 2003 (42)	Iwamoto et al, N 2005 (63)	Gilsanz et al, Ye 2006 (37)	Pitukcheewanont N and Safani, 2006 (64)	Ezenwa et al, N 2008 (65)

ALP = alkaline phosphatase; BMI = body mass index; CTX = C-terminal tetopeptide of type I collagen; DXA = dual-energy x-ray absorptiometry; NA = not applicable; NTX = N-terminal telopeptide of type I collagen.