

ORIGINAL RESEARCH

Site-Specific Transmission of a Floor-Based, High-Frequency, Low-Magnitude Vibration Stimulus in Children With Spastic Cerebral Palsy



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Abstract

Objective: To determine the degree to which a high-frequency, low-magnitude vibration signal emitted by a floor-based platform transmits to the distal tibia and distal femur of children with spastic cerebral palsy (CP) during standing.

Design: Cross-sectional study.

Setting: University research laboratory.

Participants: Children with spastic CP who could stand independently (n=18) and typically developing children (n=10) (age range, 4–12y) participated in the study (N=28).

Interventions: Not applicable.

Main Outcome Measures: The vibration signal at the high-frequency, low-magnitude vibration platform (approximately 33Hz and 0.3g), distal tibia, and distal femur was measured using accelerometers. The degree of plantar flexor spasticity was assessed using the Modified Ashworth Scale.

Results: The high-frequency, low-magnitude vibration signal was greater ($P<.001$) at the distal tibia than at the platform in children with CP ($.36\pm.06g$ vs $.29\pm.05g$) and controls ($.40\pm.09g$ vs $.24\pm.07g$). Although the vibration signal was also higher at the distal femur ($.35\pm.09g$, $P<.001$) than at the platform in controls, it was lower in children with CP ($.20\pm.07g$, $P<.001$). The degree of spasticity was negatively related to the vibration signal transmitted to the distal tibia (Spearman $\rho = -.547$) and distal femur (Spearman $\rho = -.566$) in children with CP (both $P<.05$).

Conclusions: A high-frequency, low-magnitude vibration signal from a floor-based platform was amplified at the distal tibia, attenuated at the distal femur, and inversely related to the degree of muscle spasticity in children with spastic CP. Whether this transmission pattern affects the adaptation of the bones of children with CP to high-frequency, low-magnitude vibration requires further investigation.

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Children with physical disabilities, such as cerebral palsy (CP), have reduced muscle¹ and bone^{2,3} mass and quality, especially in the lower extremities. This musculoskeletal deficiency in children with CP is associated with less force-generating capacity of the muscles⁴ and a higher incidence of low-energy fractures in the

lower extremities.⁵ Because children with CP have difficulty participating in physical activities,⁶ leading to reduced mechanical loading on their skeletal system, identifying alternate non-pharmacologic treatments is of interest.⁷

Studies have shown that a floor-based, high-frequency, low-magnitude vibration signal has an anabolic effect on bone in various populations,^{8,9} including children with CP.^{10,11} There are also studies showing no effect of high-frequency, low-magnitude vibration on bone^{12,13} or an inconsistent effect across sites.^{14,15} It is plausible that the effectiveness of high-frequency, low-magnitude vibration is dictated by the degree to which the vibration

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signal is transmitted to a particular bone site, similar to the site-specific effects of exercise.¹⁶ The tibia and femur are of interest in children with CP because they are the most commonly fractured bones, and the distal femur is the most commonly fractured site.⁵ Unfortunately, the transmission of high-frequency, low-magnitude vibration to key bone sites in children with CP has not been studied.

The primary aim of this study was to determine the degree to which a high-frequency, low-magnitude vibration signal emitted by a floor-based platform transmits to the distal tibia and distal femur of children with spastic CP during standing. An amplification of high-magnitude vibration at the ankle and an attenuation at the knee has been observed in typically developing children.¹⁷ Whether a similar profile is exhibited in children with spastic CP using a lower magnitude vibration is unknown. Toe standing, which is common in children with spastic CP, has been shown to attenuate vibration at the ankle and knee.¹⁸ Therefore, we hypothesized that the high-frequency, low-magnitude vibration transmission would be lower in children with spastic CP than typically developing children. We also hypothesized that there would be an inverse relation between the degree of spasticity and vibration transmission in children with CP.

Methods

Participants

Children with spastic CP (age range, 4–12y) at the levels of I to III using the Gross Motor Function Classification System (GMFCS) were recruited from the Nemours AI duPont Hospital for Children, Wilmington, Delaware. Children were excluded if they were unable to stand independently, if they had any metal rods in their thigh or leg, or if they had a surgery involving the musculoskeletal system or any antispastic medication (eg, botulinum toxin) within the last year. Eighteen children with CP were tested. Eight of the children had ≥ 1 previous surgery, including adductor lengthening (n=1), adductor tenotomy (n=1), adductor release (n=1), hamstring lengthening (n=6), tendoachilles lengthening (n=1), gastrocnemius recession/lengthening (n=4), and botulinum toxin (n=8). Thirteen of the children had equinus deformity and walked on their toes. Five of the children were taking antiepileptic medication. Typically developing children (n=10) in the same age range as the children with CP, between the 5th and 95th percentiles for height and body mass and without a history of chronic medication use, were recruited from the Newark, Delaware, community to serve as controls. This study was approved by the Nemours AI duPont Hospital for Children and the University of Delaware Institutional Review Boards. Participants and their parents gave written assent and consent, respectively, before testing.

Study design and procedures

A within-subject and between-group comparison design was used. Anthropometrics, pubertal development, degree of spasticity,

gross motor function, and vibration transmission were assessed during a single visit at the University of Delaware.

Anthropometrics

Height and body mass were measured while the children were wearing minimal clothing and were without shoes or braces. Height was measured to the nearest 0.1cm using a stadiometer (Seca 217^a). Body mass was measured to the nearest 0.1kg using a weighing scale (Detecto D1130^b).

Tanner staging

Tanner staging was conducted by a physician assistant to assess sexual maturity of each participant. The Tanner stage rating scale ranges from I to V, with I indicating no sign of sexual maturity and V indicating full sexual maturity.^{19,20}

Modified Ashworth Scale

Ankle plantar flexor tightness was assessed in children with CP while the participant was lying on a table in a supine position using the Modified Ashworth Scale (MAS). The grading system ranges from 0 to 4, with 0 indicating presence of normative tone and 4 indicating muscle rigidity in flexion/extension.²¹ The grade for each limb was based on an average grade of 3 trials. The reliability of spasticity assessment of the plantar flexors using the MAS was determined by evaluating 12 children with CP (age range, 4–11y) on 2 different days, 1 month apart. Cohen $\kappa = .71$ ($P < .001$) indicates good reliability.

Gross Motor Function Classification System

Gross motor function was assessed using the GMFCS, which ranges from I to V. Only those children who were GMFCS level I (walks without restrictions), II (walks without assistive device; limitations walking outdoors and in the community), or III (walks with assistive mobility device; limitations walking outdoors and in the community)²² participated in our study. The reliability of gross motor function assessment using the GMFCS was determined by evaluating 12 children with CP (age range, 4–11y) on 2 different days, 1 month apart. Cohen $\kappa = .74$ ($P < .001$) indicates good reliability.

Vibration transmission

Participants stood on a high-frequency, low-magnitude vibration platform (Juvent 1000 Motion Therapy System^c) for 3 consecutive conditions (previbration, vibration, postvibration) of 30 seconds per condition. The platform delivered a sinusoidal vertical vibration signal of approximately 0.3g at a frequency of 30 to 37Hz. No vibration was transmitted during the pre- and postvibration conditions, which were immediately before and immediately after the vibration condition, respectively. The platform was divided into right and left halves, and the center of each foot was placed in the center of the respective half. The participants stood on the platform without shoes, socks, or braces. They were instructed to stand on the platform as still as possible in a relaxed position and were encouraged to stand without support. Although all children were able to stand without assistance, poor balance is an issue associated with CP, which can be exacerbated by antiepileptic medications.²³

List of abbreviations:

CP cerebral palsy
GMFCS Gross Motor Function Classification System
MAS Modified Ashworth Scale

Therefore, as a precaution, a standard wooden chair was placed in front of the vibration platform to allow intermittent support, if needed, and a spotter stood on each side of the participant to prevent falls. Only data collected during independent standing was used for the analysis. The same vibration platform was used for all participants.

To quantify the amount of vibration emitted by the platform, a uniaxial accelerometer (model 3711B1110G^d) was secured to the center of the platform using double-sided tape, as per recommendations by Rauch et al.²⁴ and further secured with single-sided tape over the top of the accelerometer. To quantify the transmission of the vibration signal to the distal tibia and distal femur, triaxial accelerometers^d were secured to the skin immediately above the medial malleolus of the distal tibia and at the lateral condyle of the distal femur to the limit of the participant's comfort, using a self-adhesive elastic bandage. The placement of all accelerometers was done by a single research assistant for all participants. All accelerometers were calibrated before each data collection session. The same accelerometers were used at the same sites for all participants. The data collection setup is displayed in figure 1.

A 12-bit analog-to-digital converter data acquisition device (model USB-1208FS^e) was used to collect the vibration data at a sampling frequency of 2kHz using DASyLab version 11.0.^e Spike 2 version 7.10^f was used to analyze the data after it was filtered using custom MATLAB^g coding. Peak-to-peak voltage values (mV) were converted to force (g) per the sensitivity for each respective axis. The resultant values (g) were calculated for the triaxial accelerometers at the distal tibia and distal femur using the following equation: $R = \sqrt{(V_g^2 + ML_g^2 + AP_g^2)}$, with V_g , ML_g , and AP_g representing forces (g) in the vertical, mediolateral, and anteroposterior directions. A 5-second period within each 30-second condition (previbration, vibration, postvibration) was chosen to

calculate amplitude and frequency of high-frequency, low-magnitude vibration signals. Vibration transmission was defined as the difference in high-frequency, low-magnitude vibration signals at the platform versus the distal tibia and the distal femur.

The reliability of acceleration measurement at the platform, distal tibia, and distal femur was determined by assessing 6 children with CP and 9 typically developing children (age range, 4–11y) on 2 different days, 1 month apart. The intraclass correlations of .89, .91, and .95, respectively, in the combined sample (all $P < .05$), indicate excellent reliability.

Statistical analysis

Data were analyzed using SPSS version 22.0.^h Data were assessed for normality using skewness, kurtosis, and the Shapiro-Wilk test. Group differences in physical characteristics were assessed using independent t tests. The chi-square test of independence was used to determine if there were group differences in the Tanner stage. Paired t tests were used to determine if pre- and postvibration signals were different. If they were not different, the previbration condition was used for comparison with the vibration condition. A 2-way analysis of variance (group×site) with repeated measures on site was used to determine if there were group differences and an interaction effect in vibration transmission at the platform, distal tibia, and distal femur. If a group-by-site interaction was detected, simple contrasts were used to identify specific differences. A Bonferroni adjustment was made for multiple comparisons. Spearman ρ correlation analysis was used to determine if there were relations between MAS and the vibration signal transmission. Values are reported as mean \pm SD. The magnitude of effects of group differences was assessed using Cohen d , with 0.2, 0.5, and 0.8 demonstrating small, medium, and large effects.²⁵

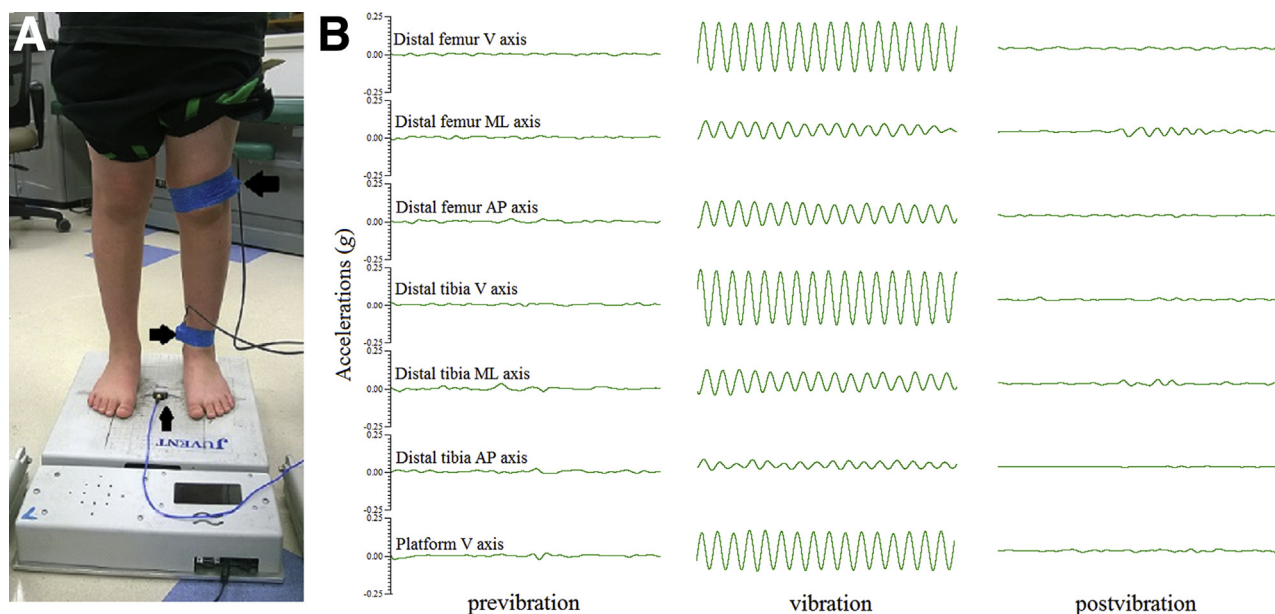


Fig 1 (A) A participant with triaxial accelerometers secured to the distal femur (large arrow) and distal tibia (medium arrow) and a uniaxial accelerometer secured to a platform that emits a high-frequency, low-magnitude vibration signal when turned on (small arrow). (B) Sinusoidal waveforms showing signals at the platform, distal tibia, and distal femur in the V, ML, and AP axes while standing before the vibration platform is turned on (previbration), while it is on (vibration), and after it is turned off (postvibration). Abbreviations: AP, anteroposterior; ML, mediolateral; V, vertical.

Results

Physical characteristics of the participants are reported in table 1. All data were normally distributed except pubic hair and testicular-penile/breast Tanner stage. There were no group differences in age, Tanner stages, body mass, body mass index, or body mass index percentile (all $P > .20$). Children with CP had lower height and height percentile than controls ($P < .05$). In addition, height in children with CP was lower than the 50th age- and sex-based percentile ($P < .001$). Body mass percentile ($P = .064$) was lower in children with CP than controls, and body mass in children with CP was also lower than the 50th age- and sex-based percentile ($P = .059$); however, the differences were marginally insignificant. Height, body mass, and body mass index in controls were not different from the 50th age- and sex-based percentiles ($P > .38$).

High-frequency, low-magnitude vibration data are reported in figure 2. No differences in pre- and postvibration measures were observed at the platform ($d = .18$, $P = .85$), distal tibia ($d = .19$, $P = .805$), or distal femur ($d = .01$, $P = .45$); therefore, the pre-vibration condition was used for comparison against the vibration condition. There was a significant group-by-site effect ($d = 1.49$, $P < .001$). Specifically, in children with CP, the vibration signal was significantly higher at the distal tibia ($d = 1.28$, $P < .001$) and lower at the distal femur ($d = 1.58$, $P < .001$) than at the platform. In controls, the vibration signal was significantly higher at the distal tibia ($d = 2.1$, $P < .001$) and higher at the distal femur ($d = 1.47$, $P < .001$) than at the platform. The same patterns were observed when only those children with CP who had equinus deformity ($n = 13$) were compared with controls ($P < .05$).

Scatterplots of MAS versus high-frequency, low-magnitude vibration transmission, as indicated by the ratio of the vibration signal measured at the site (ie, distal tibia, distal femur) to the vibration signal measured at the platform, in children with CP are depicted in figure 3. MAS was moderately and negatively related to the transmission of vibration from the platform to the distal

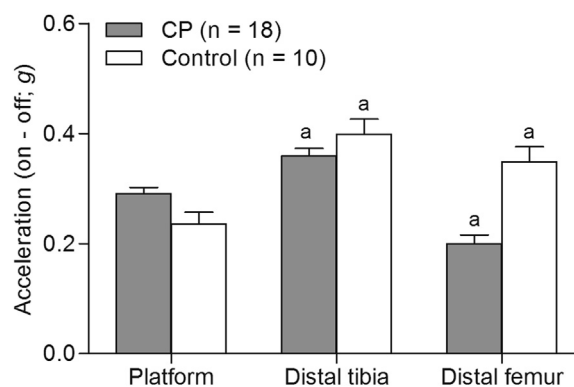


Fig 2 Bar graph showing transmission of the high-frequency, low-magnitude vibration signal from the vibration platform to the distal tibia and distal femur. The vibration signals are presented as values in the on condition (vibration) minus the off condition (previbration). ^aDifferent from the vibration signal values at the platform ($P < .05$).

tibia (Spearman $\rho = -.547$, $P = .019$) and from the platform to the distal femur (Spearman $\rho = -.566$, $P = .014$).

Discussion

This is the first study to investigate the transmission of a high-frequency, low-magnitude vibration signal from a floor-based platform (ie, transmits $< 1g$) across the lower extremity of ambulatory children with spastic CP. Relative to the signal generated at the platform, the signal at the distal tibia was amplified in children with CP and in typically developing children. The signal was also amplified at the distal femur in typically developing children, but it was dampened in children with CP to approximately 70% of the signal emitted at the platform.

One factor that may have contributed to the different pattern of high-frequency, low-magnitude vibration signal transmission in children with CP versus typically developing children is the spasticity in children with CP. There was an inverse relation between MAS and the degree of vibration transmission to the distal femur. The reason for this relation is unclear, but it may be related to the effect of spasticity on posture. Some of the children with CP had equinus deformity ($n = 13$ of 18). Therefore, they had difficulty limiting knee and ankle plantar flexion and had difficulty or were unable to place their feet completely flat on the vibration platform, which are common postures in children with spastic CP.²⁶ Previous studies have shown that increased knee flexion and toe standing leads to attenuation of a vibration signal at the knee.^{18,27} However, the pattern of the results in the present study was the same when only those with equinus deformity were included in the statistical analysis, which suggests that other factors may have contributed to the dampened transmission of vibration to the distal femur. Another factor that may have contributed to the dampened transmission of vibration to the distal femur includes the unequal weight distribution in the lower extremities, creating variable degrees of muscle tension or muscular force production. Increased muscular force production has been shown to dampen vibration.²⁸ The loss of the vibration signal at the distal femur may also be related to mechanical filtering

Table 1 Physical characteristics of children with CP and typically developing children (controls)

Characteristic	CP (n=18)	Controls (n=10)	P	d
Age (y)	8.0±2.5	8.7±1.7	.413	0.33
Tanner stage (I/II/III)				
Pubic hair	13/4/1	9/1/0	.261	0.47
Testicular-penile/breast	16/2/0	8/1/1	.323	0.42
Height (m)	1.20±0.12*	1.32±0.07	.011	1.11
Height (percentile)	21±24* [†]	56±32	.003	1.32
Body mass (kg)	25.7±9.3	29.8±6.7	.231	0.49
Body mass (percentile)	35±32	58±27	.064	0.77
BMI (kg/m ²)	17.3±3.8	16.9±2.9	.786	0.11
BMI (percentile)	52±37	48±33	.791	0.11
GMFCS (I/II/III)	10/7/1	NA		
MAS (1/1.5/2/3)	6/7/1/4	NA		

NOTE. Values are means ± SD or as otherwise indicated. Abbreviations: BMI, body mass index; NA, not applicable.

* Group difference ($P < .05$).

[†] Different from the 50th age- and sex-based percentiles ($P < .001$).

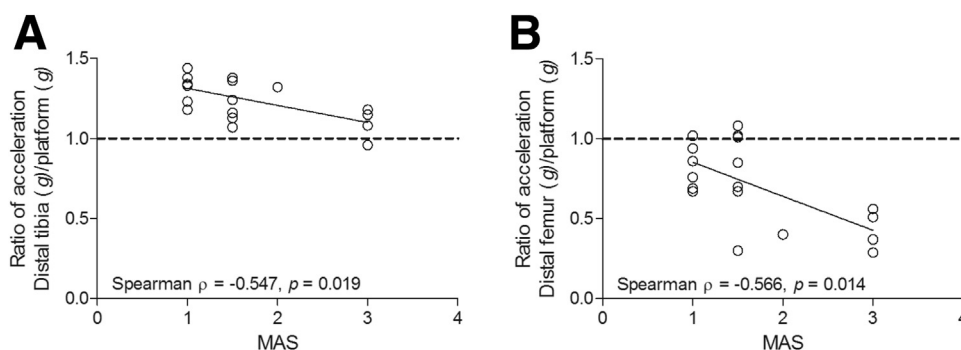


Fig 3 Scatterplots show the relation between spasticity estimated using the MAS and the transmission of the high-frequency, low-magnitude vibration signals emitted from the vibration platform to the (A) distal tibia and (B) distal femur in children with CP ($n=18$). The transmission of the vibration signal was expressed as the ratio of the acceleration measured at the bone site divided by the acceleration measured at the vibration platform. A value of 1 (dotted line) indicates 100% transmission. Values >1 indicate an amplification of the vibration signal, and values <1 indicate a loss of the vibration signal.

through altered soft tissue.²⁸ Individuals with CP have a high concentration of fat within²⁹ and surrounding³⁰ their musculature, which might lead to attenuated vibration signals at the distal femur. However, the effect of leg adiposity on high-frequency, low-magnitude vibration transmission has not been investigated. Foot position on the vibration platform is another element that could affect vibration transmission and local site-specific resonance frequency.²⁴ For example, standing in the step position (ie, standing with one foot behind the other) has been shown to give rise to local resonance at 12.5 to 25Hz, leading to amplified vibration transmission at the lateral epicondyle of the femur.¹⁸ The potential effect of foot position was minimized in this study by having all participants place their feet in the same place at the center of the platform.

The amplified vibration signal at the distal tibia in children with CP and in typically developing children is consistent with previous findings in adults.^{17,18,31} It is also consistent with a study by Bressel et al,¹⁷ who reported an amplification of vibration signal to the distal tibia in typically developing children during standing while using a higher magnitude vibration signal (approximately 2.15–5.15g) than used in the present study (approximately 0.3g) and not considering a high-frequency, low-magnitude vibration (ie, not $<1g$). The amplification of the vibration signal can be attributed to many factors, chief being the resonant frequency. Resonance is the propensity of a structure to oscillate at a greater amplitude at some specific frequencies or over a range of frequencies. Previous studies have determined that the resonance frequency of the ankle lies between 10 and 63Hz in healthy adults^{18,31} and between 28 and 33Hz in children.¹⁷ In the current study, a mean frequency of 33Hz was used. Another factor that may contribute to the amplified vibration signal at the distal tibia is the lack of natural shock absorbers to attenuate the vibration signal at the foot.³² The effect of an amplified signal on bone is unknown and requires further investigation.

The vibration signal at the distal femur of typically developing children was higher than the signal measured at the vibration platform (see fig 2). This is inconsistent with previous studies that have examined vibration transmission to the knee and showed a dampening of the signal.^{17,18,31} However, previous studies have assessed vibration signals^{18,31} that were higher in magnitude or were at different sites (eg, tibial tuberosity).^{17,31} In the present

study, we assessed the transmission of a very low magnitude vibration signal (approximately 0.3g) at the lateral femoral condyle. There is evidence that the resonance frequency is different along the length of a bone,³³ which may partially account for our different results.

One strength of our study was the homogeneous nature of the participants. All the children with CP were able to stand independently, and all had spasticity in their leg muscles. The pattern of findings remained the same when typically developing children were matched to a subgroup of 10 children with CP for age, sex, and race. Moreover, typically developing children were not different from the 50th age- and sex-based percentiles for height, body mass, and body mass index. Another strength of this study is that the transmission of the high-frequency, low-magnitude vibration signal was evaluated at key bone sites. More than 80% of all fractures in children with CP occur in the lower extremities, with almost half of all fractures occurring at the distal femur.⁵

Study limitations

One of the limitations of this study is the lack of kinematic data. We did not collect any data related to posture of our participants. Skin-mounted accelerometers can overestimate the acceleration signal by approximately 10%.³⁴ However, even when a 10% correction was made (not reported), the pattern of the findings remained the same.

Conclusions

The results from this study suggest that the high-frequency, low-magnitude vibration signal from a floor-based platform is amplified at the distal tibia but dampened at the distal femur in children with CP. The dampening of vibration is related to the degree of spasticity, with greater spasticity associated with less signal transmission. Future studies are needed to uncover the specific mechanisms underlying the relation between spasticity and vibration signal transmission. Studies are also needed to determine if the potential vibration-induced benefits to bone mass and architecture in children with CP are influenced by the degree of vibration transmission.

Suppliers

- a. Seca 217; Seca Medical Measuring Systems and Scales.
- b. Detecto D1130; Detecto.
- c. Juvent 1000 Motion Therapy System; Juvent.
- d. model 3711B1110G; PCB Piezotronics.
- e. Measurement Computing.
- f. Spike 2 version 7.10; Cambridge Electronic Design.
- g. MATLAB; MathWorks.
- h. SPSS version 22.0; IBM.

Keywords

Bone and bones; Cerebral palsy; Muscle spasticity; Rehabilitation; Vibration

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References

1. Elder GC, Kirk J, Stewart G, et al. Contributing factors to muscle weakness in children with cerebral palsy. *Dev Med Child Neurol* 2003;45:542-50.
2. Modlesky CM, Kanoff SA, Johnson DL, Subramanian P, Miller F. Evaluation of the femoral midshaft in children with cerebral palsy using magnetic resonance imaging. *Osteoporos Int* 2009;20:609-15.
3. Modlesky CM, Whitney DG, Singh H, Barbe MF, Kirby JT, Miller F. Underdevelopment of trabecular bone microarchitecture in the distal femur of nonambulatory children with cerebral palsy becomes more pronounced with distance from the growth plate. *Osteoporos Int* 2015; 26:505-12.
4. Riad J, Haglund-Akerlind Y, Miller F. Power generation in children with spastic hemiplegic cerebral palsy. *Gait Posture* 2008;27:641-7.
5. Presedo A, Dabney KW, Miller F. Fractures in patients with cerebral palsy. *J Pediatr Orthoped* 2007;27:147-53.
6. Carlon SL, Taylor NF, Dodd KJ, Shields N. Differences in habitual physical activity levels of young people with cerebral palsy and their typically developing peers: a systematic review. *Disabil Rehabil* 2013; 35:647-55.
7. Houlihan CM, Stevenson RD. Bone density in cerebral palsy. *Phys Med Rehabil Clin N Am* 2009;20:493-508.
8. Rubin C, Recker R, Cullen D, Ryaby J, McCabe J, McLeod K. Prevention of postmenopausal bone loss by a low-magnitude, high-frequency mechanical stimuli: a clinical trial assessing compliance, efficacy, and safety. *J Bone Miner Res* 2004;19:343-51.
9. Gilsanz V, Wren TA, Sanchez M, Dorey F, Judex S, Rubin C. Low-level, high-frequency mechanical signals enhance musculoskeletal development of young women with low BMD. *J Bone Miner Res* 2006;21:1464-74.
10. Wren TA, Lee DC, Hara R, et al. Effect of high-frequency, low-magnitude vibration on bone and muscle in children with cerebral palsy. *J Pediatr Orthoped* 2010;30:732-8.
11. Ward K, Alsop C, Caulton J, Rubin C, Adams J, Mughal Z. Low magnitude mechanical loading is osteogenic in children with disabling conditions. *J Bone Miner Res* 2004;19:360-9.
12. Slatkowska L, Alibhai SM, Beyene J, Hu HX, Demaras A, Cheung AM. Effect of 12 months of whole-body vibration therapy on bone density and structure in postmenopausal women a randomized trial. *Ann Intern Med* 2011;155:668-79. W205.
13. Leung KS, Li CY, Tse YK, et al. Effects of 18-month low-magnitude high-frequency vibration on fall rate and fracture risks in 710 community elderly-a cluster-randomized controlled trial. *Osteoporos Int* 2014;25:1785-95.
14. Lam TP, Ng BK, Cheung LW, Lee KM, Qin L, Cheng JC. Effect of whole body vibration (WBV) therapy on bone density and bone quality in osteopenic girls with adolescent idiopathic scoliosis: a randomized, controlled trial. *Osteoporos Int* 2013;24: 1623-36.
15. Judex S, Lei X, Han D, Rubin C. Low-magnitude mechanical signals that stimulate bone formation in the ovariectomized rat are dependent on the applied frequency but not on the strain magnitude. *J Biomech* 2007;40:1333-9.
16. Haapasalo H, Kannus P, Sievanen H, Heinonen A, Oja P, Vuori I. Long-term unilateral loading and bone-mineral density and content in female squash players. *Calcif Tissue Int* 1994;54:249-55.
17. Bressel E, Smith G, Branscomb J. Transmission of whole body vibration in children while standing. *Clin Biomech* 2010;25:181-6.
18. Harazin B, Grzesik J. The transmission of vertical whole-body vibration to the body segments of standing subjects. *J Sound Vib* 1998; 215:775-87.
19. Marshall WA, Tanner JM. Variations in pattern of pubertal changes in girls. *Arch Dis Child* 1969;44:291-303.
20. Marshall WA, Tanner JM. Variations in pattern of pubertal changes in boys. *Arch Dis Child* 1970;45:13-23.
21. Ashworth B. Preliminary trial of carisoprodol in multiple sclerosis. *Practitioner* 1964;192:540-2.
22. Palisano R, Rosenbaum P, Walter S, Russell D, Wood E, Galuppi B. Development and reliability of a system to classify gross motor function in children with cerebral palsy. *Dev Med Child Neurol* 1997; 39:214-23.
23. van Gaalen J, Kerstens FG, Maas RP, Harmark L, van de Warrenburg BP. Drug-induced cerebellar ataxia: a systematic review. *CNS Drugs* 2014;28:1139-53.
24. Rauch F, Sievanen H, Boonen S, et al. Reporting whole-body vibration intervention studies: recommendations of the International Society of Musculoskeletal and Neuronal Interactions. *J Musculoskelet Neuronal Interact* 2010;10:193-8.
25. Cohen J. *Statistical power analysis for behavioral sciences*. 2nd ed. Hillsdale: Erlbaum; 1987.
26. Lidbeck CM, Gutierrez-Farewik EM, Brostrom E, Bartonek A. Postural orientation during standing in children with bilateral cerebral palsy. *Pediatr PhysTher* 2014;26:223-9.
27. Matsumoto Y, Griffin MJ. Dynamic response of the standing human body exposed to vertical vibration: influence of posture and vibration magnitude. *J Sound Vib* 1998;212:85-107.
28. Wakeling JM, Nigg BM. Modification of soft tissue vibrations in the leg by muscular activity. *J Appl Physiol* 2001;90:412-20.
29. Noble JJ, Charles-Edwards GD, Keevil SF, Lewis AP, Gough M, Shortland AP. Intramuscular fat in ambulant young adults with bilateral spastic cerebral palsy. *BMC Musculoskelet Disord* 2014;15:236.
30. Johnson DL, Miller F, Subramanian P, Modlesky CM. Adipose tissue infiltration of skeletal muscle in children with cerebral palsy. *J Pediatr* 2009;154:715-20.
31. Kiiski J, Heinonen A, Jaervinen TL, Kannus P, Sievanen H. Transmission of vertical whole body vibration to the human body. *J Bone Miner Res* 2008;23:1318-25.
32. Voloshin A, Wosk J. An in vivo study of low back pain and shock absorption in the human locomotor system. *J Biomech* 1982;15: 21-7.
33. Zhao LM, Dodge T, Nemani A, Yokota H. Resonance in the mouse tibia as a predictor of frequencies and locations of loading-induced bone formation. *Biomech Model Mechanobiol* 2014;13:141-51.
34. Kim W, Voloshin AS, Johnson SH, Simkin A. Measurement of the impulsive bone motion by skin-mounted accelerometers. *J Biomech Eng* 1993;115:47-52.