Research Paper: Interference of Various Sources of Sensory Feedback on Transition of Relative Phase in Bimanual Coordination on Active and Inactive Woman With Multiple Sclerosis

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**Purpose:** Comparing the effects of manipulating senses on relative phase transition bimanual coordination pattern of active and inactive women with Multiple Sclerosis (MS).

**Methods:** The methodology of this study was repeated-measures design. Study participants comprised 10 active women and 10 inactive women with multiple sclerosis who voluntarily participated in this experiment. The participants (n=20; aged 18-25 years) performed bimanual in-phase and anti-phase movements with their wrists at three different speeds ranging from slow to fast and 5 different sensory conditions, including 1) normal sensory input, 2) masked vision, 3) masked proprioception, 4) masked audition, and 5) full sensory deprivation. Two-way (5 sensory conditions×2 groups) analyses of variance for repeated measures (ANOVA) were performed using SPSS 24.

**Results:** The findings showed that the main effect of the senses and groups was not significant, while the effect of interaction between the senses and groups was significant (P=0.047). Accordingly, there was a significant difference between two groups with regard to the manipulation of proprioception, vision, and audition conditions. Regarding the effect of different senses in both active and inactive women with MS, only a significant effect was observed between two groups in manipulation of proprioception condition, when vision and audition were not manipulated (P=0.004). Also, comparing active and inactive woman with MS in different manipulated sensory conditions did not reveal significant differences.

**Conclusion:** Proprioception in both active and inactive woman with MS has a significant impact on performing bimanual coordination task.

**Keywords:** Relative phase transition, Multiple Sclerosis, Bimanual coordination, Sensory feedback

1. Introduction

Any daily activities require continuous updating of the perception-action cycle to maintain the accuracy of human motor behavior [1]. Among the different sensory modalities, vision, proprioception, and audition have an essential role in planning and guiding movements in time and space [2-4]. Daily motor tasks often demand...
using our hands simultaneously rather than separately, necessitating performance of bimanual movements [5].

For upper limb movements made in the horizontal plane, these principles of coordination are realized in two stable patterns: in-phase (0° relative phase) and anti-phase (180° relative phase). The in-phase coordination mode refers to mirror-symmetrical movements made simultaneously towards and away from the body midline and involves concurrent activation of homologous muscle groups [6-9]. The anti-phase coordination mode refers to movements made, simultaneously, in the same direction from one side of body midline to its other side; resulting in performance of parallel (isodirectional) movement pattern in extrinsic space [6, 10-12]. With increasing movement frequency, in-phase coordination remains stable while anti-phase coordination destabilizes, and if unopposed, eventually results in a spontaneous transition to the in-phase movement pattern [7, 13, 14].

It is generally acknowledged that sensory feedback plays a dominant role in fine-tune movement control. Sensory input for feedback control may include proprioception, vision, and audition. Vision is often regarded as the most important perceptive modality during interaction with environment in daily life. At least for perceiving spatial information, vision dominates other senses [15]. Many motor tasks are impossible, or at least, are much harder to perform without vision, for example, walking on an uneven terrain, hitting a tennis ball, or skiing [15]. Confirmed evidence on the critical role of vision in the coupling of limb movements came from studies using both discrete [16] and cyclical bimanual movements [17]. Specifically, these studies showed that bimanual movements were performed with higher levels of accuracy and stability when visual information on the position of the moving effectors was available as compared to other conditions where visual feedback was absent. Vision, which provides information about target and hand positions, is generally considered to be the main cue leading to sensory motor adaptation, whereas proprioception is thought to be secondary [18-20].

Besides vision, proprioception is also an important source of feedback which is essential for maintaining the required coordination patterns during bimanual movements [16]. Proprioceptive input from the muscle spindles and tendons is crucial for movement control. It allows the central nervous system to monitor the position and speed of the moving limbs and adjust the motor command if necessary. The coordination of ongoing movements uses proprioception in healthy participants [21, 22], while deafferentiated patients exhibit clear coordination deficits [23, 24]. Proprioception, however, cannot fully account for successful performance of a coordination task. For example, coordination deficits in deafferentated patients become apparent only if vision is absent [5, 23-25]. The relative contribution of vision and proprioception to the control of a coordinated movements may depend, nonetheless, on the nature of the task. For example, there is evidence to show that a bimanual circle-drawing task where movements always continue in the same direction with no reversal movements is controlled by proprioceptive feedback [26]. On the other hand, control of bimanual coordination tasks in which the effectors must stop and reverse direction, entails reliance on the use of vision and proprioception [8].

Another source for sensory feedback which plays an important role in the regulation of coordinated behavior is audition. For instance, top performance in table tennis requires auditory information about the ball bouncing on the table and racket [27]. Perhaps audition could also be taken as a sensory input influencing the coupling of the limbs during bimanual coordination as the sounds produced by the motion may provide feedback about performance. In fact, the association between the auditory and the motor systems has been examined in a recent study during performance of bimanual linear coordination task with a sliding device [28]. Yet, findings from this study suggested that the absence of auditory feedback from the motion of the sliding device was not essential for timing of the coordination patterns. Also auditory is an important source of sensory information for people with visual impairment [29].

Loss of hearing is one of the rare signs in patients with Multiple Sclerosis (MS) and about 6% of patients with MS are experiencing an auditory disorder. Hearing loss is associated with other symptoms such as vision problems, lack of balance control commonly in patients with MS that may be created because of damage in the central nervous system (brain stem). It is believed that hearing loss in patients with MS, is due to the injury and inflammation around the nerve center of the eighth that is the location of the auditory nerve [30]. MS is a disease, which unfortunately has affected many young boys and girls [30] in the present century. The most common age of starting this disease is the young age and approximately two times more prevalent in women than men [31]. Each movement from simple to complex, requires coordination of all parts of the body. Meanwhile the patients with MS because have decreased motor performance compared to healthy subjects and many researchers and physicians are trying to increase the motor performance and motor coordination in these patients.
In the present study, we used a bimanual coordination task consisting of flexion and extension movements with both wrists in either an in-phase or an anti-phase mode. The continuous nature of the bimanual actions in the present study requires the participants to control the limb extensively in an online manner, through visual, proprioception, or audition feedback loops. Assessing the relative phase transitions contribution of each aforementioned sensory source on the strength of coupling between the two wrists in active and inactive woman with MS was our primary goal. It is also of our interest to further insight into the dynamics of bimanual coordination by examining how deprivation of the three sensory sources would affect performance. It is generally acknowledged that one requires proprioceptive and visual information to fine tune motor patterns. Exploring the coordinative behavior at the absence of all sensory sources received less investigation, especially in patients with MS. However, to our knowledge, the technique of sensory feedback transformations has been used mainly in unilateral tasks whereas bimanual tasks have received much less attention. The present experimental design also addressed the question whether in-phase and anti-phase bimanual coordination patterns (which vary from each other by the degrees of their performance stability at increasing movement speeds) are differentially affected by the absence or presence of visual, proprioceptive and or auditory feedbacks in active and inactive woman with MS.

2. Materials and Methods

Participants

The design of this study was quasi-experimental. The statistical population consisted of all active and inactive patients with MS in Tehran City; of them 20 young females were selected by convenience sampling method. The criteria for selection of active patients consist of ability to carry out routine tasks and do exercise three times a week. The study participants were female patients with MS and EDSS (Expanded Disability Status Scale) score between 2 and 5.5 who were referred to MS Society Rehabilitation Clinic of Dr. Shah Begay. All participants were right handed (assessed by the Edinburgh Handedness Inventory) [32] adults, aged 18-26 years with a mean age of 21 years. Also, all had normal vision based on the Snellen chart test. Participants were also asked to complete questionnaires on their health prior to their inclusion in the study. All participants read and signed an informed consent form, which has been approved by the local Ethics Board (Urmia University).

Apparatus

Participants sat on an adjustable chair at a table covered by a white laminated poster board (50 cm deep and 86 cm wide). Wrist movements were permitted in only the extension and flexion orientation from midline. Attached in parallel to the slides were linear potentiometers (Bourns Instruments, Riverside, CA), which encoded the displacement of the handled over a 20-second trial. Data were sampled using a microprocessor (80486) with a sampling rate of 150 Hz. Lab Windows software (National instrument corporation, version 2.2.1) initiated and terminated 20-second trials and also provided data capture and recording of limb position over time.

An auditory metronome (NCH Swift Sound Tone Generator, version 2.01) provided pacing information for bimanual task [28]. In auditory deprivation condition, a with-noise masking stimulus (NCH Swift Sound Tone Generator, version 2.01) was delivered to the subjects’ ears via supra-aural headphones (optimum pro 155 stereo headphone) so that audition about performance from the linear wrist task was masked. We used a self-build tendon vibrator consisted of pager and small vibration motors, which rotated an unbalanced mass attached to the shaft of a small magnet DC motor. This apparatus constitutes very low-cost actuators for inducing tendon vibration. We have also used a Panasonic vibration motor (micro-motor with dimensions 0.59×1.15) with an operating range of 150 Hz. It was reported by Naito et al. [33] that although some qualitative aspects of the illusion were affected by the amplitude of vibration, illusion strength was determined mainly by vibration frequency. The surface area of the vibrator head was adjusted to allow an optimal contact with the skin by adding bars with different profiles. The vibrator was positioned over the wrist tendon near the radiocarpal joint and vibration frequency was set for each participant to generate maximum illusion.

Procedure

The participants were willing to do the task. They were asked to take two handles adhered to the moving slides and put them horizontally in the left-right dimension (wrist extension and flexion). While grasping two handles, the participants produced 00 relative phase (in-phase) and 1800 relative phase (anti-phase) patterns. In the in-phase patterns, the participants were instructed to move their hands toward and then away from each other symmetrically and continuously by activating homologous flexor and extensor muscles of the wrists. In the anti-phase patterns, the limbs moved together in an isodirectional fashion with homologous muscle group
contracting in an alternating fashion. The participants received instructions to keep pace with a metronome by performing a complete cycle of in-out-in handle alternating in time with the beat. The metronome outpaced the assigned speed or frequency of limb movement starting at a slow speed equivalent to a frequency of 58 beat in minute for 20 seconds. After completion of the 20-second trial at slow speed, the same required coordination task was paced at a medium metronome frequency (90 beat per minute), and subsequently at a fast metronome frequency (152 beat per minute).

When the participants entered the laboratory, we obtained consent (and assent, when appropriate). Next, we conducted a handedness inventory after the participants were seated. We encouraged the participants to perform in-phase and anti-phase patterns during experiment. There were 5 counterbalanced conditions: 1) normal sensory condition (vision, proprioception, and audition were available); 2) masked vision (normal proprioception and audition); 3) masked proprioception by tendon vibration (normal vision and audition); 4) masked audition by white noise (normal vision and proprioception); and 5) full sensory deprivation (no vision, masked proprioception, and masked audition).

Data processing

The position signals were smoothed with a symmetrical Bartlett (triangular) filter. Velocity time series were derived from the position signal using a 2-point central difference algorithm and then smoothed with a Bartlett window. The smoothed position and velocity time series were then used to calculate each component of the near-continuous phase state for each trial according to the formula:

$$\phi_R = \tan^{-1}\left(\frac{dXR/dt}{XR}\right)$$

where $\phi_R$ is the phase of the right wrist at each sample, XR is the position of the right wrist rescaled to the interval [-1,1] for each cycle of oscillation, and (dXR/dt) is its normalized instantaneous velocity. The same formula was used to calculate from the position and velocity signals of the left wrist. The relative phase ($\phi$) between the two wrists, was then expressed as:

$$\phi = \phi_R - \phi_L$$

The mean absolute error of relative phase (AE) reflected the spread of relative phase measures around the mean (coordination consistency).

Statistics

$AE$ and $SD\phi$ scores were determined for each trial and subsequently averaged across test trials of each condition. Two-way (5 sensory conditions×2 groups) analyses of variance for repeated measures (ANOVA) were implemented by using Statistical software (version 8.0). The significant results of interest were investigated post hoc by using pairwise comparisons with Bonferroni correction. The level of significance was set to $P=0.05$.

3. Results

Sensory condition and group on coordination consistency

The main effect for group ($P=0.123$) and sensory condition ($P=0.156$) did not reach the level of significance. However, interactions between sensory conditions and groups for the AE scores were visible in relative phase transition (Table 1). The results of the 5×2 ANOVA for the relative phase transition revealed a significant interaction effect for sensory condition and group ($F_{4,56}=325.13$, $P=0.001$). This issue showed a significant difference between two groups in sensory condition of the vision, audition, and proprioception masked. To study the mutual impact, a 2-way analysis of variance test was utilized to investigate different sensory conditions between two groups (Figure 1). Moreover, 2-way replicated measures analysis of variance tests was done to assess differences between each experimental condition in either group separately. In order to do so, the adjustment coefficient alpha was used. Table 2 presents the results of follow-up. As shown in Table 2, the only significant effect was observed between the two groups in a state of proprioception sense manipulated along with normal vision and audition ($P=0.004$). Figure 2 shows situation of the trial between the two groups.

Figure 2 shows that in every sensory condition, MS active group have been better than MS inactive one. Only in the full sensory deprivation condition, MS inactive group slightly performed better than MS active group.

4. Discussion

The experiment was designed to investigate whether sensory information contributed by proprioception, vision, and audition is parametrically redundant or distinct. MS active and inactive subjects performed coordination
The results of analysis of variance with repeated measures to investigate the relative phase transition between active and inactive groups with MS

<table>
<thead>
<tr>
<th>Statistical Indicators</th>
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<td>543.2</td>
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<td></td>
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Table 1. Results of ANOVA test to determine the effect of various senses in both active and inactive female patients with MS

<table>
<thead>
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Figure 1. Histograms showing the average frequency relative phase transfer (AE scores) in two groups and five sensory conditions performed bimanual coordination tasks with higher mean relative phase error scores and poorer coordination consistency as compared to other conditions when proprioception was not manipulated. Stated differently, the reliance on proprioception in performance was evident not only during the performance of the “non-stable” anti-phase patterns but also in the “stable” in-phase patterns, suggesting that bimanual coupling appears to be controlled, for the main part, by proprioceptive information. These findings are largely supported by previous

Table 2. Results of ANOVA test to determine the effect of various senses in both active and inactive female patients with MS

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Our results agree with positive effects of exercise on proprioception. In Swanik [35] study, 6-week plyometric training (exercise that use muscle stretching and contraction for improving muscle function) improved proprioception of the shoulder in women swimmers. In Sharbatoghli [36] study, by using rhythmic exercises (rhythmic stabilization, tilt board) on lumbosacral area, a significant improvement was reported in proprioception. According to conducted research, exercise improves proprioception. The present study was in agreement with Swanik [35] and Sharbatoghli [36] studies, which reported that exercise therapy program can improve proprioception and reduce error angles (that is criteria for evaluating proprioception error) at least in healthy young women [37].

A possible mechanism to improve proprioception due to exercise is increasing attention. Attention is a neuropsychological process by which the central nervous system affects received information. Probably, proprioceptive training increases proprioception signs (first, at the conscious level and then at the automatic level) in the brain. Other possible mechanisms to explain the improvement in proprioception as a result of exercise, can be attributed to the activated pathway, increasing the number of synapses, and higher plasticity seen in the sensory area. However, these possible mechanisms in changing proprioception have not been justified yet. Studies have also shown that muscle spindle output can be increased voluntarily by changing the tone that eventually increase accuracy [38]. Myers et al. [39] also believe that the specific rehabilitation techniques are effective in improving motor sensory system, mechanical retrieval receptor afferents of the nervous system and facilitate the afferents aid as a compensatory mechanism that caused by defects in proprioception [39]. However, the discussed mechanism can only be seen in patients with lesions but not in normal subjects [37].

In this study, the complicated data from the wrist joints were altered by means of tendon vibration. It was shown that movement patterns with vibrations produce larger mean relative phase error than those without vibrations. Vibratory stimuli activate muscle spindles, through increase in their discharge levels. The extra stimulation of these receptors creates an artificial input and introduces a discordant source of afferent messages. Considering the convergence of different sources of proprioceptive information, whether from an external perturbation (vibration) or an active movement, one can hypothesize that the available afferent inflow is distorted and does not correspond to the actual situation at wrists. Interference prevails, leading to less accurate coordination patterns and an increase of relative phase errors. Our findings were, consistent with the findings reported in a study by Baldissera et al. and Serrien et al. [10, 17] indicating the need for attention. The present study also shows that the triceps and anterior deltoid muscle vibration lead to reducing the amplitude of the drawing circles. The vibration of the wrist flexor tendon destroys coordination between the two hands [10, 17, 34]. This result shows that proprioceptive information plays an essential role in the central nervous system for keeping posture [17]. The present study agrees with Grillo et al. [28] and Rocha et

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**Figure 2.** Histograms showing the average frequency relative phase transfer (AE scores) in the MS active and inactive groups in five different sensory conditions

- **Normal sensory**
- **Masked proprioception**
- **Masked audition**
- **Masked vision**
- **Full sensory deprivation**

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Our findings were, nonetheless, in contrast to the findings of Grillo et al. [28]. This may be attributed to differences in procedure applied in two studies to deprive vision stimuli. First, in Grillo et al. [28] study combination of the vision and audition was not under control. Therefore findings from this study lack the ability to determine the extent by which participants integrate the two sensory modalities (i.e. vision and audition). Second, vision in Grillo et al. [28] study was manipulated by switching off the lights in the room, which may have allowed participants to use some visual input whereas in our study the occultation of vision was complete. Participants in our study were blindfolded, allowing complete preclusion of visual feedback. Where the sensory input from the hand was biased during ipsilateral hand and foot coordination at different levels of speed.

Our study observations also suggested that afferent information was elaborated differently during in-phase and anti-phase movements. For the in-phase movements (isodirectional movements), inferences are matched by a feedback mechanism automatically, whereas for anti-phase movements (non-isodirectional movements), the afferent signals rely more strongly on a controlling process requiring abundant purposeful demands. It is important to note that Baldissera et al. [10] examined the effect of sensory bias on the coordination of ipsilateral hand and foot movements whereas the task/effectors tested in our study were bimanual coordination with the wrists. Nonetheless, trends in behavior that emerged for both studies were similar. Specifically, both studies showed that the influence of sensory bias on relative phase error was more pronounced during execution of the difficult tasks than the easy ones. Taken together, our study findings and that of Baldissera et al. [10] suggest that processing of sensory information in general and afferent input in particular are essential for successful performance of coordinated movements irrespective of the task and/or effector combinations.

The availability of visual feedback also influenced the performance of subjects in the present study. This finding was in agreement with the observations reported in the studies of Cardoso de Oliveira and Barthelemy [16], showing that the presence of visual information enabled stable in-phase movements, meanwhile the stability of the anti-phase movements. The latter finding suggests that visual monitoring influences the production of both coordination modes in a different way. It might be interpreted as follows. During in-phase coordination, the extremities of both hands are in central vision and when the reversal occurs in a flexed position. This position may allow for an adequate calibration in terms of spatiotemporal dimensions. During anti-phase coordination, the bimanual phase offset never permits this optimal position because one hand will always be out of view [17]. In contrast to vision and proprioception the presence or absence of audition has no significant influence on performance. Our findings were in contrast to those of Ghez et al. [41] that used discrete auditory cues for audition feedback. This contrast was related to the method by which auditory feedback was presented to the participants. When we experimentally manipulated auditory feedback, for example by delaying audition, motor performance was significantly influenced. Yet, by obstructing task sound with headphone like in the study of Grillo et al. [28] and in the present study, coordination performance was not affected. These findings replicated those from previous research [7, 42]. These findings can be explained based on ideomotor theories of action control [43]. The vital presumption of this method is that motor actions are cognitively indicated by their sensory impacts, i.e. by the codes of the impressive impacts that progressively follow certain motor actions. Consequently a motor action can solely be achieved by recollecting the codes of the sensory ramifications that normally come along with this action, and serve to mentally represent it. Put it in a different way, there is no other way to produce a motor action other than by anticipating its sensory ramifications.

The important implication of this approach is that all the constraints of motor control that we know of, such as complexity effects, stimulus-response compatibility, and or symmetry tendencies in bimanual coordination, do not arise because of inherent constraints in the structure and/or functions of the motor system, but due to constraints in the representation of the perceptual differences of to-be-produced motor actions [44]. More studies should be conducted to further explore this hypothesis.

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Conflict of Interest

The authors declared no conflicts of interest.
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