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Characteristics of Paraspinal Muscle Spindle Response to Mechanically-Assisted Spinal Manipulation-A Preliminary Report

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Abstract

Objectives—The purpose of this preliminary study is to determine muscle spindle response characteristics related to the use of two solenoid powered clinical mechanically-assisted manipulation (MAM) devices.

Methods— L_6 muscle spindle afferents with receptive fields in paraspinal muscles were isolated in 6 cats. Neural recordings were made during L_7 MAM thrusts using the Activator V and/or

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Contributorship

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Writing (responsible for writing a substantive part of the manuscript) WRR

<u>Critical review</u> (revised manuscript for intellectual content, this does not relate to spelling and grammar checking) WRR, JWL, RSS, JGP, MRG, MAL

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CONFLICTS OF INTEREST

No conflicts of interest were reported for this study.

Pulstar devices at their 3 lowest force settings. MAM response measures included: a) the time required post-thrust until the 1st action potential, b) differences in mean frequency (MF) and mean instantaneous frequency (MIF) 2s before and after MAM, and c) the time required for muscle spindle discharge (MF & MIF) to return to 95% of baseline following MAM.

Results—Depending on device setting, between 44–80% (Pulstar) and 11–63% (Activator V) of spindle afferents required >6s to return to within 95% of baseline MF values; whereas 66–89% (Pulstar) and 75–100% (Activator V) of spindle responses returned to within 95% of baseline MIF in <6s following MAM. Nonparametric comparisons between the 22 and 44N settings of the Pulstar yielded significant differences for the time required to return to baseline MF and MIF.

Conclusion—Short duration (<10ms) MAM thrusts decrease muscle spindle discharge with a majority of afferents requiring prolonged periods of time (>6s) to return to baseline MF activity. Physiological consequences and/or clinical relevance of described MAM mechanoreceptor responses will require additional investigation.

Keywords

Spinal manipulation; muscle spindles; afferent neurons; spine

Introduction

Low back pain (LBP) continues to be a major societal health problem and prevalence rates are only expected to increase with the aging of the American population.^{1,2} More than 80% of the population will experience an episode of LBP during their lives, with 20 to 40% experiencing a reoccurrence of a LBP episode within 1 year after the initial onset in working populations.^{3,4} LBP can be treated with multiple approaches including non-pharmacological, pharmacological, and surgical interventions. The efficacy, favorable side-effect profile, cost-effectiveness and high patient satisfaction associated with the non-pharmacological approach of spinal manipulation have led to multiple clinical practice guidelines and evidence reports recommending spinal manipulation for certain types of acute and chronic LBP.^{5–11} However, the appropriate use and clinical efficacy of spinal manipulation could be greatly enhanced if underlying peripheral and central neurophysiological mechanisms were clearly established. Accomplishing these objectives will require combined contributions from both basic and clinical investigations.

Spinal manipulation traditionally involves applying a single, short lever, high velocity low amplitude (HVLA) thrust of short duration to a targeted dysfunctional vertebral motion segment and/or joint. HVLA thrusts can be delivered manually with direct hand contact (150ms thrust duration) or using commercially available mechanical-assisted devices (10ms thrust duration).^{12–18} Peak forces achieved during a manipulative thrust can vary dramatically depending upon anatomical location of treatment, localized tissue compliance, clinician and/or patient's physical traits, spinal manipulative technique, mechanical-assisted device used and/or device setting.^{15–25} A distinct advantage mechanically-assisted manipulation (MAM) offers over manual spinal manipulation is that thrust velocity and thrust magnitude can be standardized. MAM is thought to be an effective alternative to manual thrust manipulation based upon both comparable measures of vertebrae

movements^{12,26–30} and similar clinical outcomes reported in a few smaller studies.^{31–34} A recent randomized clinical trial comparing manually-delivered spinal manipulation and MAM indicated differences in short-term clinical efficacy,³⁵ thereby suggesting that distinct physiological mechanisms of action might be responsible for the underlying therapeutic effects of these two types of HVLA spinal manipulation.

Proposed physiological mechanisms of action by which spinal manipulation of any type ameliorates musculoskeletal pain include stretch reflex inhibition generated by stretchinduced stimulation of mechanoreceptors or nociceptors in the facet joint capsule and/or the surrounding spinal tissues (muscles, ligaments, discs).^{36–43} In humans, the Hoffmann reflex (H-reflex) and stretch reflexes have been used to provide a neurophysiological index of amotoneuron excitability. A commonality shared by a majority of spinal manipulative studies that investigated H-reflex and stretch reflex responses is the concept that manual therapies mechanistically act by reflexively reducing central α -motoneuron activity via modulating the sensitivity of muscle spindle afferents to muscle stretch.^{37,41–48} Using a computercontrolled feedback motor, we have previously investigated the relationship between various biomechanical parameters of spinal manipulation (thrust duration, thrust magnitude, thrust rate, thrust contact site etc.) and paraspinal muscle spindle responsiveness in both laminectomy-only and acute facet joint fixation animal models.⁴⁹⁻⁵⁵ Time to peak impulse in the majority of these animal studies ranged from 75-150ms which is more closely aligned with manually-delivered spinal manipulation compared to the faster time to peak impulse of 2-7ms reported among MAM devices.^{15,18} This extremely short time to peak impulse with MAM devices may result in unique muscle spindle responses and/or central reflex-related mechanisms as these particular mechanoreceptors are sensitive to the rate of muscle stretch.

To begin to address the neurophysiological impact of these extremely short MAM thrust durations, we recently conducted a pilot study in a single feline preparation to determine the feasibility of recording muscle spindle afferent responses using an instrumented spring-loaded hammer MAM device (Activator IV ; Activator Methods Int. Ltd., Phoenix, AZ).⁵⁶ Despite recording from a limited number of muscle spindle afferents, this pilot study clearly demonstrated that a diverse set of muscle spindle afferent responses are elicited using clinical MAM devices. Post-thrust MAM neural responses ranged from a slight diminution of muscle afferent discharge compared to pre-thrust, to a complete cessation of afferent discharge for nearly 3 seconds.⁵⁶ The purpose of this preliminary study is to begin to characterize the temporal effects of two electromagnetic solenoid-powered MAM devices (Activator Methods Int. Ltd., Phoenix, AZ [Fig. 1A] and Pulstar; Sense Technology Inc., Pittsburgh, PA [Fig. 1B]) on muscle spindle discharge to inform future studies.

Methods

General Surgery

All experiments were reviewed and approved by the Palmer Institutional Animal Care and Use Committee #20120601. Electrophysiological activity in 6 muscle spindle afferent fibers was obtained during MAM delivered to 6 deeply anesthetized male cats weighing between 4.4 and 6.0kg [mean 5.3 (SD 0.6)]. Data collected using these electromagnetic MAM devices were from experimental preparations associated with a separate larger study

investigating the relationship between intervertebral fixation and spinal manipulation as delivered by a computer controlled feedback motor.⁵⁵ The surgical and experimental procedures, including monitoring the preparation's anesthetic level and physiological status as well as methods of isolating paraspinal muscle spindle afferents have been previously described in detail elsewhere^{51,52,55,57,58} and thus are only presented here briefly. Anesthesia was induced with isoflurane and maintained with sodium pentobarbital (35mg/kg, iv). Catheters were placed in the common carotid artery and external jugular vein to monitor blood pressure, introduce fluids and anesthesia. The trachea was intubated, and the cat ventilated mechanically. Arterial gases and body temperature were monitored and maintained within the reference range (pH 7.32–7.43; PCO2, 32–37mmHg; PO2, >85mmHg, 38–39°C). Body temperature was continuously monitored and maintained between 38–39°C using a water-heating pad/radiant heat. Additional Nembutal (5 mg/kg) was administered (iv) upon withdrawal reflex to noxious pinching of the toe pad or when mean arterial pressure increased above 100mmHg.

To better isolate trunk muscle afferents, the right sciatic nerve was cut to reduce afferent input from the hindlimb. The lumbar spine was mechanically secured at the L_4 spinous and the iliac crests. A midline skin incision was made and an L_5 laminectomy was performed to expose the L_6 dorsal rootlets while leaving all intervertebral joints intact. The L6 dorsal rootlets were cut close to the spinal cord and thin filaments were teased using forceps until impulse activity from a single trunk muscle spindle afferent was identified (Fig. 2). Action potentials were recorded using a PC-based data acquisition system (Spike 2; Cambridge Electronic Design, Cambridge, UK). Afferents were identified as muscle spindles based on their increased discharge to succinylcholine (100–200mg/kg intra-arterially; Butler Schein, Dublin, OH), decreased discharge to electrically induced muscle contraction, and/or sustained response to a fast vibratory stimulus. At the end of the experiment, animals were euthanized by an intravenous overdose of pentobarbital.

Mechanically-Assisted Manipulation

Both the Activator V and Pulstar are commercially available hand-held clinical MAM devices. The Activator V has four progressive thrust magnitude settings and requires that the device tip to completely retracted during pre-load (~2.9N) before a single mechanical impulse can delivered at any device setting. When tested against a stiff spinal tissue analog (258.07 N/mm), Activator V delivered mean peak forces of 52, 68, 112, and 155N at settings 1-4 respectively in hand-held operations.¹⁸ Device testing on more compliant spinal tissue analog (30.22 N/mm) resulted in mean peak forces of 35, 63, 102N at settings 1-3 respectively.¹⁸ The Pulstar device has 7 settings and can be used to deliver either single or multiple high velocity mechanical impulses (22-155N) once the pre-defined preload value of ~15N has been successfully applied.^{13,14} The three lowest settings on the Pulstar device deliver approximate peak forces of 22, 44, and 67N respectively. In the present study, only single impulses were delivered using the Pulstar device. All MAM thrusts were delivered perpendicularly onto the exposed fascia overlying the cat's L₇ spinous process as previously described.⁵⁶ The L₇ spinous was chosen to receive MAM due to the potential increased risk for repetitive L_6 manipulations to tear the L_6 afferent fiber off the recording electrode. Typically, three HVLA thrusts were delivered per afferent at each of the three lowest settings

of the respective MAM devices (18 thrusts total) with 5 minute intervals between thrusts. Use of MAM devices and device settings (thrust magnitudes) were randomized to minimize any possible ordering effects. At times, not all MAM test protocols were completed before the neural recording was lost. Four afferents received L_7 HVLA thrusts from both MAM devices while two afferents received L_7 HVLA thrusts from only the Pulstar device.

Data Analysis

Muscle spindle discharge was quantified as mean frequency (MF) and mean instantaneous frequency (MIF). MIF was calculated by taking the reciprocal of the time interval between successive action potentials. Baseline MF/MIF activity measures were obtained during a 2 second period that immediately preceded MAM device contact and/or soft tissue preload as in previous studies 51-53,55 (Fig. 2). The following three response measures were determined: 1) the time required post-thrust until the 1st action potential, 2) MF and MIF differences between 2s post-MAM thrust and 2s of baseline activity, and 3) the time required for spindle discharge to return to 95% of baseline MF/MIF for a minimum period of 1s following MAM (Fig. 2). The descriptive data presented in Tables 1–4 are mean responses or time durations from 8 to 18 individual MAM thrusts delivered at a given device setting. Statistical comparisons were made between device settings of a given MAM device using the mean afferent responses to the MAM thrusts delivered at each device setting. For the Pulstar device, nonparametric comparisons of magnitude settings were performed on ranked data to minimize the effect of increased variability coupled with smaller sample size using the Wilcoxon Signed Ranks Test. Statistical comparisons for the Activator V device were not performed due to fewer afferents tested with this device. Statistical significance was set at P < 0.05 and data variance is shown as standard error of the mean (SEM) unless otherwise noted.

Results

Responses to MAM from 6 muscle spindle afferents with receptive fields located in paraspinal muscles (longissimus or multifidus) were recorded in 6 cats. Mean muscle spindle response characteristics for individual MAM thrust settings are presented in Tables 1–4. Note that applied forces, pulse widths, device preload settings, etc. are not equivalent between devices thus direct comparisons between MAM devices are inappropriate.

Activator V Device

An example of a raw recording from a single afferent to MAM using the Activator at the 2nd device setting is shown in Figure 2. Progressive increases in Activator V MAM thrust forces typically resulted in longer mean delays until the 1st post-thrust action potential (Table 1). Compared to the 2s of baseline activity, MAM resulted in mean decreases in spindle discharge for both MF and MIF during the 2s post-MAM thrust. Progressive increases in MAM thrust forces did not necessarily result in greater changes in MF/MIF between 2s after MAM and 2s baseline measures. The greatest mean change in MF and MIF occurred at the 2nd device setting on the Activator V (Table 1). Regarding the time required for MF/MIF to return to 95% of baseline following Activator MAM (Table 2), 25–45% of the afferents tested returned to baseline MF within 2s post-MAM and no afferent required longer than 8s

to return to baseline MF at device setting 1. However at ActivatorV device settings 2 and 3, over 33% of afferents required >10s thus creating a distinct separation of those afferents requiring shorter (<2s) or longer (>10s) durations to return to baseline MF. Regardless of device setting, the vast majority of afferents (67–89%) tested with the Activator V returned to baseline MIF within 2s and no afferent required a period greater than 8s (Table 2).

Pulstar Device

An example of 3 raw recordings from a single muscle spindle afferent to MAM using the Pulstar device at settings of 22, 44, and 67N are shown in Figure 3. The time required until the first action potential was similar at 44 and 67N. Compared to the 2s of baseline activity, Pulstar MAM resulted in mean decreases in spindle discharge for both MF and MIF during the 2s post-MAM thrust at all thrust magnitudes (Table 3). Regarding the time required for MF/MIF to return to 95% of baseline following MAM (Table 4), it is interesting to note that over 33% of afferents tested required more than 10s to return to baseline MF at all Pulstar device settings tested. The delay in return to baseline MIF with the Pulstar was broadly distributed across post-MAM time intervals at each of the three device settings tested (Table 4). Nonparametric comparisons between outcomes among the different Pulstar device settings yielded significant differences between 22 and 44N settings for the time required to return to baseline MF was 8.4s (1.9) and 14.8s (2.6) for the 22 and 44N setting respectively. The mean time required to return to baseline MIF was 3.4s (0.3) and 5.2s (0.8) for the 22 and 44N Pulstar device settings.

Discussion

Basic investigations related to neurophysiological changes associated with MAM are important in order to establish underlying mechanisms responsible for spinal manipulation efficacy and to optimize treatment regimens for the 40–60% of chiropractic and/or other manual therapy clinicians in the United States, Canada, Britain, Belgium, New Zealand and Australia who use these mechanical manipulation devices in some capacity of patient care.^{59–64} Despite recent strides being made in determining both the mechanical characteristics and/or physiological effects of MAM,^{13,15,17,18,25,56,65–74} to our knowledge, this study is the first to record muscle spindle response evoked using solenoid powered hand-held clinical MAM device technology and only the second to investigate muscle spindle response using any type of MAM device.

This study confirms findings of a previous report using a spring-loaded MAM device (Activator IV) demonstrating that a range of muscle spindle responses are elicited by MAM.⁵⁶ The combined tissue preload and extremely short MAM thrust durations elicited high frequency discharge from paraspinal muscle spindle afferents followed by a decrease in post-MAM activity. The progressive increase in applied forces typically increased the time required until the 1st action potential (Tables 1, 3). However, this relationship between applied force and delayed neural response did not necessarily prove to be the case with regard to the amount of time required to return to 95% of baseline MF or MIF. Nonparametric analyses of afferent response to Pulstar thrusts indicated significant

differences between 22 and 44N thrust settings with greater time being required to MF/MIF to return to 95% baseline levels for the 44N MAM thrust but not between the 44 and 67N thrust settings. There appeared to be distinct subpopulations of afferent responses with some returning to baseline activity levels relatively rapidly (<4s) while others required much longer periods (>10s) (Tables 2, 4). It is not known whether these differences are due to inherit differences related to muscle spindle intrafusal fibers types such as bag₁, bag₂ and/or chain fibers (see Reed et al.⁷⁵ for greater detail) or to other biological factors.

In the present study we provide post-MAM data regarding changes in both MF and MIF (Tables 1–4), while in the past 51,54 we have primarily focused our attention on MIF changes during the thrust phase (time to peak force) of the spinal manipulation. Due to the lack of tissue preload standardization and hand-held vs computer controlled feedback motor clinical delivery methods, we have chosen to focus on the post-thrust MAM muscle spindle response instead of the spindle response elicited during the thrust. Since muscle spindle afferents typically discharge tonically while at rest, we thought providing the time required to return to 95% baseline activity for both MF and MIF might provide a more complete picture of post-thrust MAM response. As clearly illustrated in Fig. 2, MF and MIF recovery do not always occur within close proximity, but generally speaking, these two response measures typically occurred within 0.5-2s of each other. Comparisons for the Pulstar device indicated significant increases for both MF and MIF when comparing time to return to baseline spindle activity following MAM thrusts delivered at the 22 and 44N device settings. We are uncertain as to how this prolonged delay in return to baseline discharge following MAM affects central interneuron, gamma and/or alpha-motoneuron activity and whether or not these delay differences contribute to the physiological mechanisms or clinical efficacy of spinal manipulation. Answers to these questions will require additional basic and clinical investigation using MAM devices.

Limitations

Preload forces and preload durations were not standardized and MAM devices were not rigidly secured or instrumented to record the actual forces delivered. Instead, MAM devices were hand-held as typically used in clinical settings and care was taken to apply the minimum preload necessary to trigger the respective MAM device. While we tested the 3 lowest settings on each clinical device (with peak forces as low as 22N), peak forces were not scaled to feline bodyweight per se. While aspects of a scaling requirement have validity, determining the best scaling parameters is far from clear. Localized thrust forces are quickly dampened by surrounding viscoelastic in vivo tissues and biological cell response does not scale with body weight.^{18,76} Compared to direct load cell MAM testing, recent evidence using spinal tissue analogs demonstrate that applied MAM peak forces are substantially lower (up to 59%) based on localized viscoelastic tissue compliance combined with additional compliance from the hand/wrist/forearm when devices are hand-held.¹⁸ It should be noted that L_7 MAM thrust procedures were not intense enough to tear nearby L_6 finely teased afferent fibers from the recording electrode and no tissue damage or spinal fractures occurred with multiple MAM thrusts. While future studies should consider scaling of applied MAM peak forces, this study highlights the need to determine the actual forces being transmitted through the spine and surrounding tissues in vivo and suggests that these

applied forces are likely substantially lower than those reported in the literature following testing on highly idealized beam structures.¹⁸

Conclusions

This study marks the first time that paraspinal muscle spindle response characteristics related to the delivery of extremely short duration (<10ms) solenoid powered MAM thrusts has been reported. Overall, MAM thrusts resulted decreased muscle spindle discharge with a majority of afferents requiring prolonged periods of time (>6s) to return to pre-thrust levels of MF/MIF activity. This neurophysiological effect of MAM far outlasts the 2–7ms stimulus duration, but determination as to how this prolonged delay in return to baseline activity levels impacts spinal or supraspinal mechanosensory processing or clinical outcomes will require additional investigation.

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This study found that short duration (<10ms) mechanically-assisted lumbar manipulative thrusts decreases muscle spindle discharge relative to baseline activity,

It required prolonged periods (>6s) for paraspinal muscle spindles to return to their prethrust resting discharge activity levels following short duration (<10ms) mechanicallyassisted spinal manipulative thrusts.

This study suggests that subpopulations of muscle spindle afferents respond differently to short duration mechanically-assisted spinal manipulative thrusts.

These findings may become important to patient care if future studies show that mechanosensory response is a mechanism that contributes to the clinical efficacy of spinal manipulation.

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Figure 1. Photographs of the Activator V (A), and Pulstar (B) mechanical-assisted manipulation devices.

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Figure 2.

Example of a paraspinal muscle spindle response to an Activator V mechanically-assisted manipulation (MAM) at magnitude setting of 2. Note the increase in discharge during preload and thrust delivery followed by a period of discharge cessation and subsequent return to resting discharge frequency. Time required for spindle discharge to return to 95% of baseline MIF/MF for a minimum period of 1s was 4.5 and 10.5s respectively.



Figure 3.

Three recordings from the same muscle spindle afferent in response to mechanically-assisted manipulation (MAM) using the Pulstar at the 22, 44 and 67N device settings (A–C respectively). Mean discharge frequency was the greatest during the 22N thrust (A). Peak mean discharge frequencies were similar for the 44N (B) and 67N (C) MAM thrust. The 67N thrust exhibited a more pronounced decrease in post-MAM discharge. The secondary

peak in mean frequency discharge with the 67N thrust may have been the result of the increased forces and/or recoil contact due to the device being hand-held.

Neural Response Characteristics Following Activator V MAM

Shows the time required until the first action potential and the mean differences between mean frequency (MF) and mean instantaneous frequency (MIF) 2s after Activator V mechanical-assisted manipulation (MAM) and 2s baseline measures. All Activator V MAM thrusts resulted in a decrease in spindle discharge relative to baseline activity.

Г

Time (s) to 1 st Actio MAM	n Potential Af	ter		Change from Basel MAM	line During 2	s After	
Force Setting 3 (n=8)	Setting 1 (n=9)	Setti (n=	ng 2 12)	Force Setting 3 (n=8)	Setting 1 (n=9)	Setti (n=	ng 2 [2)
Moon Time (SEM)	054 (0.01)	10 07 391	W W LUS	Mean MF (SEM)	-2.2 (0.6)	-6.0 (2.2)	-3.9 (1.8)
	(10.0) +00.	(7.0) 004.	(+.0) /00.	Mean MIF (SEM)	-2.1 (0.5)	-6.6 (2.5)	-4.0 (1.9)

Time Required to Return to 95% of Baseline MF & MIF After Activator VTM MAM

return to MF/MIF baseline. The reported values are the percentage of afferents returning to MF/MIF baseline discharge for the respective time period. Activator V mechanical-assisted manipulation (MAM). Different subpopulations of spindle afferents appear to exist according to the time required to Time required to return to 95% of baseline mean frequency (MF) and mean instantaneous frequency (MIF) discharge for a minimum of 1s following

	Mean	Frequency	(MF)	Mean Instan	taneous Frequ	ancy (MIF)
Force Time (s)	Setting 1 (n=9)	Setting 2 (n=12)	Setting 3 (n=8)	Setting 1 (n=9)	Setting 2 (n=12)	Setting 3 (n=8)
0 - 1.99	45%	33%	25%	%68	%19	75%
2 – 3.99	33%	%8	%0	11%	8%	0%0
4 – 5.99	11%	8%	12%	%0	25%	0%0
6 - 7.99	11%	%0	25%	%0	%0	25%
8 - 10	%0	8%	0%	%0	%0	%0
>10	%0	43%	38%	%0	%0	%0

Table 3

Neural Response Characteristics Following Pulstar MAM

Shows the time required until the first action potential and mean differences between mean frequency (MF) and mean instantaneous frequency (MIF) 2s after Pulstar mechanical-assisted manipulation (MAM) and 2s baseline measures. All Pulstar MAM thrusts resulted in a decrease in spindle discharge relative to baseline activity.

Г

Time (s) to	o 1 st Action F MAM	otential Afte	r	Change fr	om Baseline l MAM	During 2s Aft	er
Force 67N (n=15)	22N (n=18)	=u) 74	N 18)	Force 67N (n=15)	22N (n=18)	44 (n=	N 18)
Mana Time (SEM)	00000000	(1.0).080	10,190	Mean MF (SEM)	-5.1 (0.6)	-4.9 (0.9)	-4.3 (1.3)
	(6.0) 677.	(1.0) 607.	(1.0) +02.	Mean MIF (SEM)	-4.1 (0.5)	-4.3 (1.0)	-3.4 (1.5)

Time Required to Return to 95% of Baseline MF & MIF After Pulstar MAM

Pulstar mechanical-assisted manipulation (MAM). Different subpopulations of spindle afferents appear to exist according to the time required to return to Time required to return to 95% of baseline mean frequency (MF) and mean instantaneous frequency (MIF) discharge for a minimum of 1s following MF/MIF baseline. The reported values are the percentage of afferents returning to MF/MIF baseline discharge for the respective time period.

	Mean	Frequency	7 (MF)	Mean Instan	taneous Frequ	iency (MIF)
Force Time (s)	22N (n=18)	44N (n=18)	67N (n=15)	22N (n=18)	44N (n=18)	67N (n=15)
0 - 1.99	11%	%0	13%	39%	22%	13%
2 - 3.99	17%	11%	%L	22%	33%	33%
4 – 5.99	28%	11%	%0	28%	17%	20%
6 - 7.99	2%	17%	20%	%0	11%	7%
8 -10	5%	5%	13%	5%	6%	%0
>10	34%	26%	47%	6%	11%	27%