

Analysis of the relationship between M wave parameters and pain *

Análise da relação entre parâmetros da onda M e a dor

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SUMMARY

BACKGROUND AND OBJECTIVES: Pain is a subjective and individual sensation causing major discomfort. So, it is necessary to put into practice methods to objectively quantify it. Several studies indicate that evoked potentials (EP) generate responses which may reflect painful processes. Our study has used characteristics extracted from the M wave, which is a type of EP related to neuromuscular response, as possible pain sensation markers.

METHOD: Participated in this study 13 volunteers, being 6 males and 7 females. Five sessions of an experimental protocol were held. In each session, volunteers received electrical stimulations on the abductor hallucis muscle. Initially, 1 mA current with 20 pulses of 2 Hz was applied. Then, the electric current was incremented in 1 mA until volunteer reported maximum pain. For female volunteers, menstrual cycle day in each session was also recorded. Several M wave characteristics were evaluated.

RESULTS: Global minimum of the M wave is correlated to increased electric current amplitude of stimulation, and consequently to more pain felt by the individual. Female participants have reported pain threshold and tolerance with lower electric current intensity as compared to male volunteers.

CONCLUSION: Global minimum of the M wave was related to increased pain and in males pain threshold and tolerance were higher as compared to females. Different menstrual cycle phases have not interfered with evaluations.

Keywords: Electric stimulation, Evoked potentials, Pain, Sensory threshold.

RESUMO

JUSTIFICATIVA E OBJETIVOS: A dor é uma sensação subjetiva e individual que causa muitos desconfortos para quem a sente. Dessa forma, é necessário por em prática métodos que a quantifiquem objetivamente. Vários estudos indicam que os potenciais evocados (PE) produzem respostas que podem refletir os processos dolorosos. Este estudo utilizou características extraídas da onda M, sendo esta um tipo de PE relacionado à resposta neuromuscular, como possíveis marcadores da sensação de dor.

MÉTODO: Treze voluntários, sendo 6 do sexo masculino e 7 do sexo feminino, foram incluídos no estudo. Foram realizadas cinco sessões de um protocolo experimental. Em cada sessão o voluntário recebeu estímulos elétricos no músculo abductor do hálux. Inicialmente foi aplicada uma corrente de 1 mA com 20 pulsos de 2 Hz. Depois a corrente elétrica foi incrementada de 1 mA em 1 mA até que o voluntário relatasse a dor máxima. Nos voluntários do sexo feminino foi registrado também o dia do ciclo menstrual em cada sessão do experimento. Foram avaliadas várias características das ondas M.

RESULTADOS: O mínimo global da onda M é correlacionado com o aumento da amplitude da corrente elétrica do estímulo, conseqüentemente com o aumento da dor sentida pelo sujeito. Os participantes do sexo feminino relataram limiar e tolerância à dor com intensidades

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de corrente elétrica menor quando comparados aos voluntários do sexo masculino.

CONCLUSÃO: O mínimo global da onda M mostrou relação com o aumento da sensação dolorosa sendo que no sexo masculino o limiar e a tolerância à dor foram maiores que no sexo feminino, e as diferentes fases do ciclo menstrual não interferiram nas avaliações.

Descritores: Dor, Estimulação elétrica, Limiar sensorial, Potenciais evocados.

INTRODUCTION

Pain is a subjective and individual sensation, currently measured with subjective scales with potential susceptibility to contamination by several factors external to the immediate pain sensation, such as anxiety, expectations, past experiences, among others, which may contribute to errors and confuse subjective pain variations¹.

For causing so much trouble and being unable to be objectively measured², it is necessary to look for alternatives to objectively quantify pain to use the most adequate treatment according to its real intensity, that is, detecting pain via a mechanism not influenced by emotional factors. Pain intensity evaluation is, in general, done with the help of the visual analog scale (VAS)³, which is also subjective.

Several studies report the importance of identifying pain intensity, however they use subjective and not always reliable methods for such measurements, since most methods require some voluntary answer from the patient who, sometimes, is unable to react or to understand what should be done⁴⁻⁶.

Pain may be evaluated with tools such as electromyography (EMG), evoked potentials analysis and electrical stimulation⁷. Electrical stimulation is widely used in physical therapies⁸. A study carried out to assess the level of sensory discomfort caused by low and medium frequency currents in electrical stimulation of femoral quadriceps muscles in healthy women makes clear the presence of pain during electrical stimulation sessions⁹.

This study aimed at using electrical stimulation as pain stimulation means and at assessing the possible correlation between M wave parameters and pain sensations reported by volunteers and evaluated with VAS.

METHOD

After the approval of the Research Ethics Commit-

tee, Federal University of Uberlândia (UFU), protocol 036/09, this study was carried out with 13 healthy individuals, being 6 males and 7 females, aged between 18 and 30 years, students of UFU courses. Data were collected in the Laboratory of Biomedical Engineering (Biolab) of the university.

Inclusion criteria were no history of surgery, dominant lower limb pain or injury. Exclusion criteria were presence of central or peripheral neurological disorders and rheumatic impairment; use of pacemaker or heart problems; obesity; use of drugs changing motor control and peripheral sensitivity, such as benzodiazepines, opioids, anti-histaminics, anticonvulsants and antidepressants; lower limb amputation and diabetes mellitus.

Every volunteer participated in five experimental sessions. All sessions were carried out in the same period of the day due to possible Circadian rhythm influences. During the experiments, volunteers remained comfortably in the supine position in a reclining chair, with plantar ankle flexion and foot inversion. Dominant foot was used. Before positioning the electrodes, skin was cleaned with alcohol and, when needed, it was shaved. Reference electrode of the electrical stimulation was fixed at the lateral foot margin and electrical stimulation electrode was placed at the medial foot margin, at the motor point of the abductor hallucis muscle.

To detect the motor point of the abductor hallucis muscle a pen-shaped electrode was used which was slid over the medial foot margin simultaneously with electrical pulses emission. The point with the maximum mechanical response with minimum current was considered the motor point of the abductor hallucis muscle (Figure 1).

EMG electrodes were fixed between the motor point and the distal muscle tendon, with the reference electrode positioned on the medial malleolus of the same limb. Electromyographical signal capturing electrode and the reference electrode were fixed with tapes specific for this purpose and an electroslag paste was used between the electrodes and the skin (Figure 2).

To ensure that EMG and electrical stimulation electrodes were positioned at the same place during the five days of test, each electrode received a staining layer which remains on the skin for approximately one week, popularly known as henna, because medial malleolus is very protuberant.

Tolerance level of research subjects was assessed by gradually increasing stimulation intensity until the max-



Figure 1 – Detection of abductor hallucis muscle motor point (medial margin).

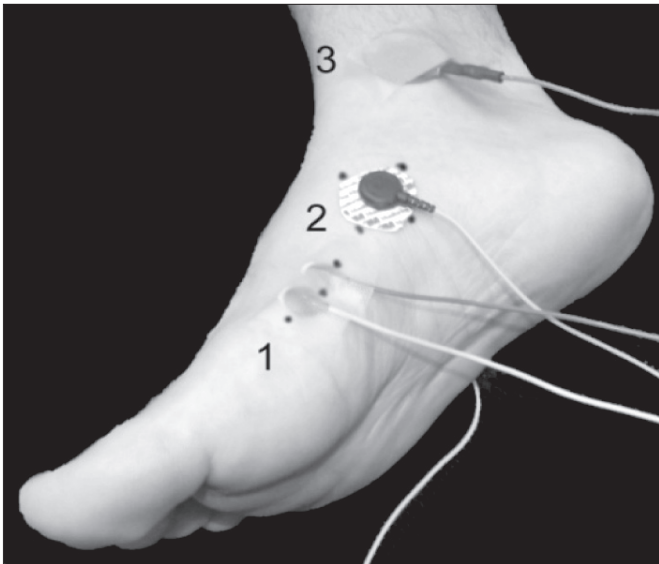


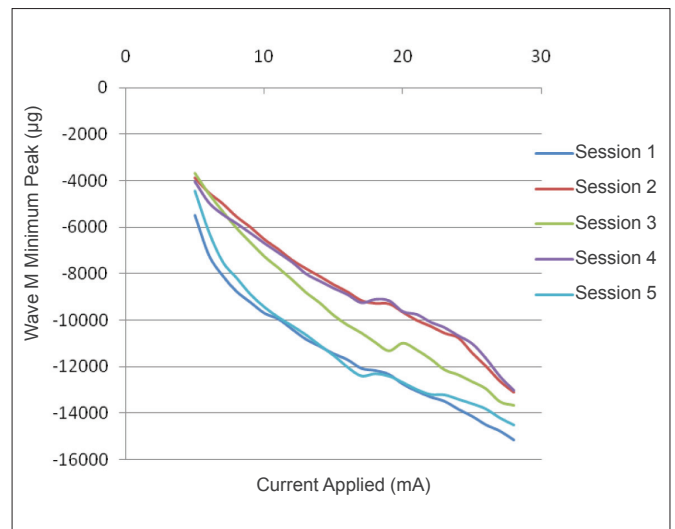
Figure 2 – Positioning of electrodes on foot medial margin. (1) EMG electrodes, (2) electrical stimulation electrode, (3) EMG reference electrode.

imum supportable limit using VAS, and the volunteer has verbally indicated pain perception from zero to 10, being zero no pain and 10 maximum tolerable pain. Volunteers received stimulations of 20 pulses of 2 Hz and 0.3 ms as from 1 mA. After each series of 20 pulses, volunteers have reported pain intensity perceived according to VAS and made observations they considered necessary for the researcher. After volunteers' report, other 20 pulses were emitted with 1 mA increments and so on, until volunteers reported pain intensity = 10. During each 20 pulses session, the researcher has recorded pain perceived by the volunteer on a table according to VAS and also reported observations. The researcher

has also recorded birth date, period of the day in which the experiment was carried out, experiment day, gender and menstrual cycle period for females. Electrical stimulation and electromyographic data were recorded with the Neuropack S1 MEB-9400 equipment, from Nihon Kohden, Japan. M wave data generated by the Neuropack equipment were stored in text format (.txt) and analyzed by customized programs developed by MatLab (MathWorks). From these programs, it was possible to estimate the following M wave parameters: minimum and maximum peak and time when they occurred. Each parameter was correlated to pain sensation reported in VAS. Preliminary evaluations have shown that only the minimum peak, that is, M wave global minimum, was correlated to pain sensation and so, only results with correlation with pain sensation were considered.

RESULTS

Graph 3 shows the relationship between applied current value (in mA) in the X-axis and minimum M wave peak value (in μV) in the Y-axis. Tables 1 and 2 show values of applied current in mA when each subject has reported the presence of pain sensation. The leftmost column describes the general mean of all volunteers described in each one of the tables, with their respective standard deviation. Tables 3 and 4 show values where each subject has reported tolerance to pain in each session and table 5 shows the menstrual cycle day in which female volunteers were submitted to tests, considering day one the day in which the volunteer menstruated and that all volunteers have a menstrual cycle of 28 days.



Graph 3 – Relationship between M wave minimum peak and applied current.

Table 1 – Current values in mA in which volunteers have reached pain threshold and mean and standard deviation of male subjects in each test.

Subjects	Session 1	Session 2	Session 3	Session 4	Session 5	Mean ± SD	General mean ± SD
1	15	17	19	20	22	18.6 ± 2.7	
2	17	19	19	18	21	18.8 ± 1.4	
3	27	19	26	30	30	26.4 ± 4.5	16.8 ± 5.7
4	9	9	11	10	12	10.2 ± 1.3	
5	13	12	14	13	13	13 ± 0.7	
6	17	13	13	14	14	14.2 ± 1.6	

Table 2 – Current values in mA in which volunteers have reached pain threshold and mean and standard deviation of female subjects in each test.

Subjects	Session 1	Session 2	Session 3	Session 4	Session 5	Mean ± SD	General mean ± SD
7	14	14	14	16	17	15 ± 1.4	
8	11	11	15	16	14	13.4 ± 2.3	
9	15	16	13	22	22	17.6 ± 4.1	14.9 ± 2.8
10	11	14	10	10	12	11.4 ± 1.6	
11	15	11	11	11	10	11.6 ± 1.9	
12	15	18	17	15	21	17.2 ± 2.4	
13	17	18	19	19	18	18.2 ± 0.8	

Table 3 – Stimulation current values in mA in which volunteers have reached pain tolerance reported by each male subject with individual and general mean and standard deviation.

Subjects	Session 1	Session 2	Session 3	Session 4	Session 5	Mean ± SD	General mean ± SD
1	30	30	33	33	38	32.8 ± 3.2	
2	39	44	43	43	49	43.6 ± 3.5	
3	-	-	-	-	-	-	36.5 ± 7.4
4	21	29	26	24	31	26.2 ± 3.9	
5	44	41	47	45	42	43.8 ± 2.3	
6	35	39	35	35	38	36.4 ± 1.9	

Table 4 – Stimulation current values in mA in which volunteers have reached pain tolerance reported by each female subject with individual and general mean and standard deviation.

Subjects	Session 1	Session 2	Session 3	Session 4	Session 5	Mean ± SD	General mean ± SD
7	27	30	30	33	33	30.6 ± 2.5	
8	48	-	-	-	-	48 ± 0	
9	32	35	33	39	41	36 ± 3.8	
10	27	28	27	29	31	28.4 ± 1.6	31.2 ± 5.2
11	23	24	24	22	23	23.2 ± 0.8	
12	27	33	34	30	34	31.6 ± 3	
13	26	43	38	42	40	37.8 ± 6.8	

Table 5 – Menstrual cycle day in which volunteers were submitted to tests.

Subjects	Session 1	Session 2	Session 3	Session 4	Session 5
7	Day 7	Day 10	Day 11	Day 12	Day 14
8	Day 20	Day 21	Day 22	Day 23	Day 24
9	-	-	-	-	-
10	Day 2	Day 3	Day 6	Day 9	Day 10
11	Day 14	Day 15	Day 17	Day 18	Day 24
12	Day 2	Day 3	Day 4	Day 5	Day 7
13	Day 18	Day 19	Day 25	Day 26	Day 28

DISCUSSION

There is major correlation between M wave minimum peak decrease and increased intensity of the current applied to individuals, both males and females. Results were as expected, showing a strong correlation between analyzed M waves parameter, minimum peak and increased current, consequently with increased pain.

A study using as pain stimulation hypertonic saline solution injection has not shown M wave changes with regard to conduction velocity, spectral amplitude and content¹⁰, leading the author to conclude that saline solution injection does not change muscle fibers membrane properties and that muscle activity decrease during voluntary contractions was probably due to a central, spinal or supraspinal phenomenon.

In our study, the pain induction protocol has shown that M waves had their amplitudes changed. Some authors understand that a change in M wave amplitude is related to changes in pH, temperature and muscle fibers diameter^{10,11}. M wave valley is exactly the point where there is major change in membrane patency and fibers contract due to a fast ions inflow caused by this patency change. This contraction is closely related to pain perception, which is the focus of the study. Such factor clearly shows the high correlation between minimum peak of this wave and the stimulating electrical current.

Pain threshold and its means in male subjects in each test (Table 1) allow us to observe that in subjects 2, 4, 5 and 6 values were maintained close in all tests, with a maximum standard deviation of 1.6 without major variations.

This brings more reliability to the method used because it shows that even in different days, the subjects had the same perception of the stimulation and reported pain with levels very close to the current, if not equal. Table 1 also shows that in most male subjects pain threshold was reported with a lower stimulation current in the first session with regard to subsequent sessions. This may be explained by subject's pain expectations because all of them have reported at the end of the first session that pain perceived was lower than pain expected before the experiment.

With regard to females, subjects 8, 9 and 12 (Table 2) had standard deviation higher than 2 with regard to pain threshold mean. As with males, pain threshold reported during the first session was with low currents

as compared to other sessions. Only subject 11 has reported pain perception at 15 mA in the first session, at 11 mA in the second, third and fourth sessions and at 10 mA in the last session.

In comparing males and females, it was observed that mean pain threshold value was higher for males, 16.8 and 14.9, respectively. These results are in line with a study aiming at assessing physiological and psychological measurements related to gender in experimental pain reports¹², study where females have reported lower pain thresholds and tolerance. Our study supports the literature review^{13,18} which has concluded that further studies are needed to differentiate pain sensation between males and females.

The analysis of pain tolerance between males and females (Tables 3 and 4) shows that, as in pain threshold, in the first session, with exception of subject 5, all subjects have reported such threshold with a lower current value as compared to other sessions. Subject 3 could not be evaluated because in all sessions he reached 50 mA reporting VAS values below 10. With regard to females, current value needed to reach maximum pain in the first session was not higher than in other sessions.

In comparing males and females, it was noticed that current value needed to reach maximum pain value was higher for males (36.5) as compared to females (31.2). These data are also in line with a study showing significant difference in pain tolerance between males and females¹².

Table 5 shows menstrual cycle phase of each volunteer during each test. First menstrual cycle day was considered the day in which menstruation has started. Except to volunteer 3, all volunteers have reported the cycle phase they were in. When women are in the pre-menstrual cycle, they feel abdominal and muscle pain and so we have supposed that volunteers could be more susceptible to pain sensation during this period, but this was not confirmed^{14,15}.

The most rigorous tool about pre-menstrual parameters, the Daily Symptom Report (DSR), states that pre-menstruation presents symptoms 6 days before menstruation¹⁴. So, subjects 9 (tests 4 and 5), 11 (test 5) and 13 (tests 3 4 and 5) were in the pre-menstrual cycle during the tests. There has been no difference when tests performed in the pre-menstrual cycle were compared to other menstrual cycle periods of volunteer females.

Several studies along time have tried to correlate dif-

ferent factors to pain, aiming at finding a way to quantify this sensation. Biopotentials may be very helpful for this task because they are processes not sensitive to personal changes, such as past experiences, mood status, etc. Such changes may impair studies evaluating pain only through subjective scales.

Our study has chosen the M wave because it is an evoked potential free from personal changes, thus being able to use physiological parameters to measure pain. Our protocol has shown that M wave parameter with higher correlation with the stimulation is its lower peak, or valley. So, our study has correlated M wave valley to pain inducing electrical stimulation. This correlation was very consistent in all volunteers, indicating that we have followed the right path.

The study has also observed that females had a lower pain threshold as compared to males and that pain tolerance in females was also lower, showing higher pain sensitivity of females.

A major contribution of this study was the development and validation of a new pain induction protocol. Another major contribution was the finding of a major M wave parameter which may be correlated to pain. Showing the difference in pain perception between males and females was also a major aspect of this study which should be taken into consideration. Further studies should be carried out to expand the validity of our results: using the same protocol with a higher number of subjects for further validation of our data; assessment of other M wave characteristics as possible pain sensation markers; statistical analysis to assess the relevance of pain threshold differences between males and females; using this protocol to compare healthy subjects to those with chronic pain.

CONCLUSION

M wave global minimum has shown relationship with pain sensation increase and in males pain threshold and tolerance were higher than in females. Different menstrual cycle phases have not interfered with evaluations.

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