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How to cite: Crema CMT, Magario LPT, Santos APBC, Riberto M. Treatment of chronic osteoarthritic hip pain with phenol neurolysis of the obturator nerve Quadfen1. Acta Fisiatr. 2018;25(4). DOI: <u>https://doi.org/10.11606/issn.2317-0190.v25i4a163881</u>

## **Article in Press**

GNP 1148 | Original Article

Treatment of chronic osteoarthritic hip pain with phenol neurolysis of the obturator nerve Quadfen 1

- 5 6 Tratamento da dor crônica na osteoartrite do quadril com neurólise do nervo obturador 7 com fenol Quadfen 1
- 8
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- 19 Submitted: 31 December 2018.
- 20 Accepted: 15 January 2019

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22 ABSTRACT

23 The conservative treatment of osteoarthritis (OA) of the hip is essentially symptomatic, seeking 24 to relieve pain and optimize function. Despite presenting great clinical effect, hip arthroplasty 25 may have restrictions, and in these situations, the neurolytic blockade of the anterior branch of 26 the obturator nerve can be a therapeutic alternative, since it stops the afferent pain from the hip 27 joint. **Objective:** To describe the results of treating patients with severe OA of the hip by applying phenol to the obturator nerve. Method: Twelve consecutive patients with OA of the hip, 28 29 refractory to conservative treatment, were recruited to have the obturator nerve localized through electrostimulation and blocked with phenol. They were evaluated in terms of pain 30 intensity via the visual analogue scale (VAS), pressure dolorimetry in the medial and lateral 31 32 gluteus medius, gluteus minimus, and piriformis, and guality of life by the Harris Hip Score (HHS) 33 at baseline (BL) after 1 (M1), 2 (M2), and 6 months (M6). Results: The patients were between 30 and 72 years old with an avarege of  $47.5 \pm 1.7$  years old, 5 of them were women. Three 34 35 patients were excluded for not being able to come for follow up evaluations. VAS values valeded from 8.2  $\pm$  0.9 at BL to 6.6  $\pm$  1.7 at M1, 6.5  $\pm$  1.7 at M2, and 7.3  $\pm$  1 in M6 (p=0.0094). As to the 36 HHS, the values were BL:33.27 ± 2.9; M1:39.2 ± 6.4; M2:40.2 ± 8.1, and M6: 38.8 ± 9.7 in the 37 38 final evaluation (p=0,040). For dolorimetry, non significant variation was BL: 11±5.7, M1: 39 7.9±2.2; M2: 10.9±5.6; M6: 8.1 ± 1.6 (p 0.69). Conclusion: The application of phenol to the 40 anterior branch of the obturator nerve can be an alternative in the treatment of severe OA of the 41 hip in patients with restrictions to Total Hip Replacement (THR), since it reduces pain and 42 improves quality of life.

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- 44 Keywords: Osteoarthritis, Hip, Pain, Rehabilitation, Therapeutics, Phenol, Quality of Life
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- 46 **RESUMO**

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47 O tratamento conservador da osteoartrite (OA) do quadril é essencialmente sintomático, visando alívio da dor e otimização da funcionalidade. Apesar de apresentar grande efeito 48 clínico, a artroplastia quadril pode ter restrições, nestas situações o bloqueio do ramo anterior 49 50 do nervo obturador (RAO) pode ser uma alternativa terapêutica, uma vez que interrompe a 51 aferência dolorosa da articulação do quadril. Objetivo: Descrever os resultados do tratamento 52 de pacientes com OA grave do quadril por meio da aplicação de fenol no nervo obturatório. Método: Nove pacientes com OA de quadril resistente ao tratamento conservador que se 53 apresentaram consecutivamente ao nosso serviço foram submetidos à aplicação de fenol no 54 55 nervo obturatório com localização por meio de eletroestimulação e avaliados quanto a 56 intensidade de dor pela escala visual analógica (EVA), dolorimetria de pressão e qualidade de 57 vida pelo Haris Hip Score (HHS) após 01, 02 e 06 meses. Resultados: Foram selecionados 12 58 pacientes que preenchiam os critérios de inclusão e exclusão, destes, 3 foram excluídos após 59 não conseguirem mais comparecer para as avaliações. Os valores médios de EVA variaram de  $8.2 \pm 0.9$  na medida basal para  $6.6 \pm 1.7$  ao final de um mês,  $6.5 \pm 1.7$  ao final de dois meses 60 61  $e 7,3 \pm 1$  com 06 meses (p=0,0094). Quanto ao HHS, os valores foram  $33,27 \pm 2,9$ ;  $39,2 \pm 6,4$ ; 62 40,2 ± 8,1 e 38,8 ± 9,7, na avaliação final. A última variável analisada foi a Dolorimetria, onde foi realizada a média entre os valores atingidos pela avaliação com o dolorímetro no glúteo 63 64 médio medial, glúteo médio lateral, glúteo mínimo e piriforme. Na avaliação inicial média de 11±5,7, 7,9±2,2 no final do primeiro mês, 10,9±5,6 no segundo e no 6º mês 8,1±1,6. Ao 65 aplicarmos o teste ANOVA, não foi observado diferença estatística (p 0,69). Conclusão: A 66 aplicação de Fenol em RAO pode ser uma alternativa no tratamento da OA de quadril grave 67 em pacientes com restrições à realização de ATQ, pois implica em redução da dor e melhora 68 69 da qualidade de vida.

71 Palavras-chave: Osteoartrite do Quadril, Dor, Reabilitação, Terapêutica, Fenol, Qualidade de
72 Vida
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#### 74 INTRODUCTION

The conservative treatment of osteoarthritis (OA) of the hip is essentially symptomatic, seeking to relieve pain and optimize function.<sup>1</sup> The consensus for therapy consists of nonpharmacological treatments such as exercise, correcting misalignments, and pharmacological treatments such as analgesics, anti-inflammatory drugs, opioids, capsaicin cream, and injections with glucocorticoids and hyaluronic acid.<sup>2</sup>

82 When the clinical treatment fails, a surgical approach with hip arthroplasty may be 83 recommended. The greatest concern is that the surgery is major and can be risky for patients 84 with other diseases. As osteoarthritis of the hip is common in older age groups and in patients 85 with greater body weight, the prevalence of comorbidities that may increase surgical risk is also 86 greater.<sup>3</sup> In these cases the use of a conservative treatment may be advisable, even if less 87 efficient, because of safety issues and morbidity.

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Another problem that afflicts patients in the Brazilian health system is the lack of resources, which results in a lack of surgical material and long waiting lists for surgical intervention. Under these conditions the neurolytic blockade of the obturator nerve can be used as a palliative analgesic therapy.<sup>4</sup>

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Greater duration of the blockade can be achieved if drugs capable of harming the nerve axons
 are used. Phenol has an immediate local anesthetic action, due to its immediate selective effect
 on smaller nerve fibers.<sup>5,6</sup>

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#### 97 **OBJECTIVE**

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99 This study sought to describe the results of treating patients with severe osteoarthritis of the hip, 100 refractory to conservative treatment, for pain relief and quality of life, through the application of

- 101 phenol to the anterior branch of the obturator nerve.
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#### 103 **METHOD**

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105 This study was approved by the research ethics committee of the institution, and all participants 106 signed the free and informed consent form. Consecutive outpatients followed in the HCRP hip 107 clinic were recruited if they met the following inclusion criteria: 108

- 109 Older than 18 years-old.
- 110 Both genders.
- Diagnosed with osteoarthritis of the hip (hip pain associated with at least two of the following:
   VHS<20mm/h, osteophytes in the femur or acetabulum or reduction of articular Reduction of</li>
   joint space in the acetabulum in rx).
- Failure of conservative treatments with therapeutic exercise, modalities, walking aids, remedies (analgesics or anti-inflammatory drugs), or with trigger-point blockades.
- Pain assessed by the Visual Analogue Scale greater than 6 at the time of evaluation.
- Not having any known allergy to phenol or uncontrolled coagulopathy.

119 The exclusion criteria were:

- Having been submitted to total hip replacement (THR) or any other surgical procedure in the lower limbs after inclusion in the study.
- 123 Limited understanding of the procedures or evaluation instruments.
- Difficulty in clinical follow-up or coming in for evaluations for any reason.

In this study, patients were evaluated at the time of recruitment as well as immediately before
the phenolic blockade (baseline - BL), then at 1 month (M1), 2 months (M2), and 6 months (M6)
after the intervention. The following assessment instruments were used:

- The visual analogue scale (VAS), which consists of a straight line of 100 mm anchored at the ends in the expressions 'no pain' and 'worst pain possible' on which the patient indicates the intensity of their pain symptom. The intensity of the pain is considered as the distance in centimeters from the end 'no pain' to the point where the patient has indicated his pain.<sup>7</sup>
- Pressure dolorimetry (PD) is the imposing of progressively more intense pressure at a steady rate of about 1 kgf/cm2/s, on trigger points identified by muscle palpation, with a dynamometer that has a circular, rubberized 1cm<sup>2</sup> standardized end until the patient manifests pain.<sup>7</sup>
- The functional questionnaire Harris Hip Score (HHS) consists of four items: pain in the affected joint, function, presence or absence of deformity, and the range of motion of this joint. The function is assessed by asking about the patient's activities of daily living and gait, whether the patient presents with claudication or needs external support, and the maximum distance he can walk. The score may range from 0 to 100, where the lower the score, the greater the functional impairment of the patient.<sup>8</sup>
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The treatment was performed with application of phenol in the anterior branch of the obturator nerve. A Braun Stimuplex HNS 12 electrostimulator was used along with a Stimuplex A 0.8X75mm needle which was inserted two fingers from the inguinal region, just below the ventral muscle of the adductor longus, with the tip of the needle pointing to the antero superior iliac spine on the same side.

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An electrical current of 2mA and 1 Hz helped to locate the nerve, which resulted in rhythmic contraction of the adductor muscles concomitant in the same frequency. Maintenance of contractions after decrease of current intensity to 1mA confirmed the tip of the needle positioned in close contact with the obturator nerve. Nerve block consisted in the injection of 2.5ml of 6% phenol. After the application, the patient was also evaluated on whether or not he needed gait support, to continue the therapeutic measures in use, and was counselled as to analgesics and maintaining physiotherapy, if necessary.

Quantitative variables were assessed for normality by the Kolmogorov-Smirnoff and described by the mean and standard deviation. Categorical variables were grouped into categories and expressed in percentages. Any incomplete data from clinical follow-up spreadsheets were filled in a conservative manner with the LOCF technique (last observation carried forward). The clinical measures of the effect of therapeutic intervention were analyzed by ANOVA for repeated measures and, where significant, there was a post hoc comparison between moments of evaluation by the paired t test. The significance level was 0.05.

### 169 **RESULTS**

Twelve patients were selected who met the inclusion and exclusion criteria and agreed to participate. The patients were between 30 and 72 years of age, with an average of 47.45±1,7; time with the disease between 2 and 16 years, with failures in conservative treatment using analgesics such as dipyrone, paracetamol, tramadol and codeine, as well as physiotherapy with THR recommended. One month after the procedure one patient could no longer participate in the study, followed by 2 other patients after the third evaluation, due to transportation problems.

Among the 12 patients at the beginning of the study, 5 were female and 7 males. After the application, all patients reported improvement of symptoms. Only one patient reported loss of motor strength in the adductor muscles: a foreseeable side effect of the procedure.

Concerning pain intensity, in the initial assessment, prior to the application of phenol to the anterior branch of the obturator nerve, the patients had a mean of  $8.2 \pm 0.9$ . with a drop to  $6.6 \pm$ 1.7 and remained stable with a slight increase in the sixth month (7.3 $\pm$ 1). A p-value of 0.009 was found when the ANOVA test was applied. When the results were analyzed in relation to the HHS, a baseline mean of  $33.27\pm2.9$  was found in the HHS, with improvement of the score after the first and second month, and a slight decrease in the sixth month (38.8 $\pm$ 9.7) and p-value of 0.040.

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The last variable analyzed was for dolorimetry, where the mean of the values was calculated for the assessment of pain in the medial and lateral gluteus medius, gluteus minimus, and piriformis. In the baseline evaluation they showed a mean of  $11\pm5.6$  and a worsening of the score in the evaluation after 1 month (7.9±2.2) and slight improvement afterwards (8.1±1.6 in the sixth month). When applying the ANOVA test, there was no statistical difference observed (p 0.69) (Table 1).

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**Table 1.** Clinical evolution of patients with hip osteoarthritis in a 6 months period following the
 application of phenol to the anterior branch of the obturator nerve

	Baseline	Month 1	Month 2	Month 6	Р
N	12	12	11	9	
EVA	$8,2 \pm 0,9$	6,6 ± 1,7*	6,5 ± 1,7*	7,3 ± 1,0* §	0,009
HHS	$33,0 \pm 2,9$	39,2 ± 6,4*	40,2 ± 8,1*	38,8 ± 9,7*	0,04
DP	11,0 ± 5,7	7,9 ± 2,2	10,9 ± 5,6	8,1 ± 1,6	0,694

Legend — VAS: visual analogue scale; HHS: Harris Hip Score. PD: pressure dolorimetry: \* difference in relation to the baseline value p<0.05; §: difference between 2 and 6 months p<0.05

#### 203 DISCUSSION

Nerve blockades are therapeutic strategies in the treatment of chronic pain. Nerve blockades can be applied when a patient presents pain in joints or even muscles. The alleged therapeutic mechanism is the interruption the nociceptive path from their origin, blocking the conduction of nerve impulses.<sup>9</sup> The analgesic effect of these procedures may be permanent or temporary by interrupting the vicious cycle of pain.

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Phenol has been used in the treatment of pain since 1936 as a neurolytic agent applied via 211 intrathecal or epidural routes, or directly on the nerves.<sup>6,10</sup> It causes nerve destruction by 212 inducing protein precipitation. There is a loss of cellular elements, separation of the myelin 213 sheath from the axon and axonal edema. The substance takes immediate effect by its selective 214 effect on small nerve fibers.<sup>11</sup> Because of these properties, phenol was chosen for the neurolysis 215 216 of the anterior branch of the obturator nerve. In a study conducted by Monagle & Ee<sup>6</sup> a 3ml intraarticular injection was applied to the hip, with improvement of pain and mobility, and after 1 year 217 218 the patient was sent for a repeat procedure. 219

Silva et al.<sup>4</sup> blocked the obturator nerve with 10ml of 0.25% bupivacaine and obtained a 100% improvement in the pain and function of one patient after the procedure, who reported independence for all activities, with the analgesic effect lasting for 40 days.

The patients in the present study had severe OA of the hip, were refractory to clinical treatment, 224 225 and complained of severe pain and many functional limitations. The results showed pain improvement by patients, who complained about intense pain, especially during the first 2 226 227 months after the blockade, with a difference of 2 points from the initial VAS and a subsequent 228 stabilization. But in the third assessment there was a slight regression of the score, concurrent 229 with the patients complaining of pain once again, a fact that may be related to the duration of 230 the phenol effect, since its effects are not permanent and a functional re-innervation occurs over a period of months-usually between 2 and 6 months. The time of action of this procedure varies 231 232 with the concentration, injected volume, duration of exposure, and injection technique.

If the phenol is injected near nerves that have a predominance of sensory fibers, it may cause dysesthesia or anesthesia that can last up to 4 months.<sup>12</sup> In the study there was only one complain about side effects, 1 patient said that felt lost of motor strength but no lost in function and no dysesthesia. The improvement in the HHS happened in terms of mobility, with larger angles achieved in goniometry, as well as in the perception of ease in activities like tying shoes, taking a bus, and going up and down stairs.

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Regarding the dolorimetry parameter, although no statistical difference was found, the patients presented a mild worsening of pain perception which, according to Imamura et al.<sup>13</sup> can be associated with the myofascial pain syndrome, a condition associated with weakness and muscle shortening, which is common in patients with OA of the hip, especially in the areas being evaluated such as the piriformis, gluteus medius and minimus, and hip adductor muscles. The patients in the study had struggle with pain for long periods of time, already with a chronic pain, affecting other joints, altering the dynamic of walk and resulting in more myofascial syndrome.

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The present study had some limitations, such as the lack of a control group and the sample size. Another bias was the large number of comorbidities in the patients, with many already having osteoarthritis elsewhere, such as gonarthrosis and widespread pain, which affects the pain perception of the patient and would require a more comprehensive treatment; this would include providing gait supports, which in the Brazilian Health System can take up to 6 months to obtain, or the need to continue with physical therapy, where there is also a lack of time slots and professionals in the public network.

#### 256 257 **CONCLUSION**

The application of phenol to the anterior branch of the obturator nerve can alleviate the pain and improve the quality of life in patients with hip OA, being an alternative treatment for patients who have not undergone THR, either because they lack the clinical conditions or because of long waiting lists for the procedure.

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