



## Magnetic Stimulation Therapy in Patients with COPD: A Systematic Review

Massimiliano Polastri <sup>a</sup>, Vittoria Comellini<sup>b</sup>, Angela Maria Grazia Pacilli <sup>c</sup>, and Stefano Nava <sup>b,c</sup>

<sup>a</sup>Medical Department of Continuity of Care and Disability, Physical Medicine and Rehabilitation, University Hospital St. Orsola-Malpighi, Bologna, Italy;

<sup>b</sup>Respiratory and Critical Care Unit, University Hospital St. Orsola-Malpighi, Bologna, Italy; <sup>c</sup>Department of Specialistic, Diagnostic and Experimental Medicine (DIMES), Alma Mater Studiorum University of Bologna, Bologna, Italy

### ABSTRACT

Magnetotherapy (MT) is a therapeutic treatment based on the use of magnetic fields (MF) that can have an anti-inflammatory and analgesic effect. MT represents a possible treatment or an ancillary therapeutic intervention for a wide range of diseases and it is often used in the field of physiotherapeutic practices. A crucial point in the treatment of chronic obstructive pulmonary disease (COPD) patients, to counteract muscular depletion and respiratory symptoms, is represented by physiotherapy. Nevertheless, the knowledge about the application of MF as a therapeutic option in COPD patients is very limited. The purpose of the present study was to define what is currently known about the use of MF in patients with COPD. A systematic review of the literature was conducted during the month of October 2017, searching three main databases. Only those citations providing detailed informations about the use of MF to treat COPD symptoms either during an acute or a chronic phase of the disease, were selected. Following the selection process three articles were included in the final analysis. The present review focused on a total of thirty-six patients with COPD, and on the effects of the application of MF. In the majority of cases, the treatment sessions with MF were carried-out in an outpatient setting, and they differed with regard to the duration; frequency of application; dosage; intensity of the applied MF. Basing on the available informations, it seems that MF is a feasible, well tolerated, safe therapeutic option, for the treatment of motor-related COPD symptoms.

### ARTICLE HISTORY

Received 5 December 2017

Revised 6 February 2018

Accepted 7 February 2018

### KEYWORDS

Magnetic fields; COPD; human body; patient care team; recurrence of symptoms; rehabilitation; respiratory care units

### Introduction

Magnetotherapy (MT) is a therapeutic treatment based on the use of magnetic fields (MF) to maintain health and treat illness. Starting from 1960s, the interest on this technique has increased, and its use as well as its diffusion have reached the maximum mostly in the Eastern European countries (most of the publications about this topic are in Russian language) where the MT was firstly applied for the treatment of different pathological conditions, including chronic pain, and acquired bone nonunion (1–12).

Researchers' interest on the use of MT still continue nowadays, as confirmed by more recent publications indexed in the US National Library of Medicine (13–16). MF can have an anti-inflammatory (17) and analgesic effect thanks to their ability to induce vasodilatation, myorelaxation, and modulation of ion exchange across cell membrane (18,19). Therefore, MT represents a possible treatment or an ancillary therapeutic intervention for a wide range of diseases, but its use in respiratory patients is still very limited, given the lack of consistent evidence. The most common respiratory disease is chronic obstructive pulmonary disease (COPD) (20–23). Typically, experience shortness of breath, reduced and worsening of quality of life (24). The indication for the treatment of this class of patients (25–28), and an increasing interest is in the feasibility of a home-based modality (29–31); although MF are used in physiotherapeutic practices,

the knowledge about the application of MF as a therapeutic option in COPD patients is very limited. To date there are not published reviews examining the application of MF in COPD patients, and very few studies are available in the literature regarding magnetic stimulation for patients with respiratory diseases. The purpose of the present study was to define what is currently known about the use of MF in patients with COPD.

### Methods

A systematic review of the literature was conducted during the month of October 2017, searching three main databases: US National Library of Medicine (PubMed), Scopus, and Web of Science.

### Search strategy

Due to the very narrowed nature of the topic, it has been chosen a search strategy that allowed to retrieve the highest number of results, using two key words: 'magnetic fields' and 'COPD' linked by the Boolean operator AND. Search was conducted in the title and abstract fields for each database. No limits were used for the date of publication.

### Inclusion and exclusion criteria

Only papers written in English, French, Italian, or Spanish were included; moreover, only those citations providing detailed

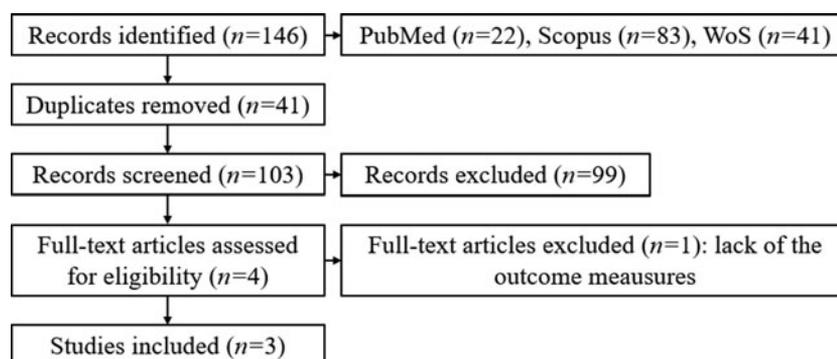


Figure 1. Flow chart.

information about the use of MF to treat COPD symptoms either during an acute or a chronic phase of the disease, were selected. Abstracts, opinion pieces, letters to the editors, and conference proceedings were not considered suitable for the inclusion in the present review. Those citations discussing the use of MF in COPD patients as a diagnostic instrument, were also considered not suitable for inclusion.

### Studies selection

Two reviewers independently revised the literature according to the abovementioned criteria; they operated a selection of the citations, and then a consensus for the final inclusion has been reached. Studies describing the use of MF in COPD patients were included. A case report, describing the use of MF in a COPD patient, was initially considered suitable for inclusion (32), but it was excluded after the full-text analysis due to the lack of outcome measures, and in accordance with the exclusion criteria (Figure 1). A full-text analysis of the included articles was performed and the results are shown below.

### Results

A total of one hundred forty-six citations published in the selected databases up to October 2017 has been found; after the selection process three articles were included in the final analysis, as shown in Figure 1. All those three papers have

been published in the last fifteen years; two out of three were randomized controlled trials (33,34).

### MF application

In one case (18), the study design was not defined by the authors: we considered it as an interventional study (35) with subjects allocated in four arms. In that study patients with COPD were allocated in two arms treated with a placebo or with sessions of pulsatile electromagnetic fields, respectively. Magnetic stimulation was performed once daily for 20 minutes for 10 days using the apparatus MTU 500H (Therapy System, Brno, Czech Republic); however, it was not described how MF were applied to the body.

### MF applied to lower limbs

The remaining studies by Bustamante and Colleagues (33,34) included two groups: in the control group the COPD patients followed a clinical monitoring, while in the treatment group they underwent to stimulation of quadriceps of both lower limbs, using the apparatus Medtronic Magpro electromagnet (Medtronic Denmark A/S, Copenhagen, Denmark; provided with a 60-mm refrigerated MCF 125 circular stimulating coil). In both studies the treatment was applied in 15-minute sessions, with the intensity and frequency of stimulation adjusted according to patient's tolerance (Table 1).

Table 1. Demographics characteristics of patients with COPD ( $n = 36$ ) treated with MF, and description of the intervention.

Variables	Studies		
	Bustamante et al. (2010)	Bustamante et al. (2008)	Sadlonova et al. (2002)
Female (n. of)	—	—	4
Male (n. of)	10	10	12
Age (years)	$61 \pm 6^*$	$59 (8)^{\S}$	$54 \pm 4^{\ddagger}$
BMI ( $\text{kg}/\text{m}^2$ )	$25.3 \pm 3.8^*$	$26.4 (6.3)^{\S}$	—
GOLD classification (stage)	III–IV	IV	—
Tiffeneau Index (%)	$32 \pm 8^*$	$31 (10)^{\S}$	—
6MWD (meters)	$397 \pm 138^*$	$440 (188)^{\S}$	—
Design	RCT Out-patient	RCT Out-patient	Interventional In-patient
Frequency	24 Quadriceps MS with coils 15–7 Hz; 40–70% of the max. output (0.8–1.4 T) 3 days/week for 8 weeks	24 Quadriceps MS with coils 15–8 Hz; 40–70% of the max. output 3 days/week for 8 weeks	10 PEMF 4.5 Hz; 3 mT daily

COPD = chronic obstructive pulmonary disease; MF = magnetic fields; BMI = body mass index; GOLD = global initiative for chronic obstructive lung disease; 6MWD = six-minute walking distance; RCT = randomized controlled trial; MS = magnetic stimulation; PEMF = pulsatile electromagnetic fields; Hz = Hertz; T = Tesla; mT = milliTesla.

\* Mean  $\pm$  standard deviation;  $\S$  median (interquartile range);  $\ddagger$  mean  $\pm$  standard error of mean (S.E.M.).

**Table 2.** Main results using MF in patients with COPD ( $n = 36$ ).

Variables	Studies		
	Bustamante et al. (2010)	Bustamante et al. (2008)	Sadlonova et al. (2002)
<b>Motor function-related parameters</b>			
6MWD (meters)	Increased by 23.4*	Increased by 25 <sup>§</sup>	—
Vastus lateralis type I fibers size ( $\mu\text{m}^2$ )	—	Increased by 734.2 <sup>§</sup>	—
Quadriceps TwQ (%)	Increased by 15.6*	—	—
Quadriceps MVC (%)	Increased by 17.5*	—	—
Quadriceps endurance (seconds)	Increased by 158*	—	—
<b>Quality of life</b>			
SF-36 (score)	Significant improvements in the following fields: physical functioning, role limitations due to physical problems, bodily pain, social functioning, mental health, role limitations due to emotional problems, vitality, general health perception, changes over time.	—	—
<b>SGRQ (score)</b>	Significant improvements in all four areas.	—	—
<b>Lung function-related parameters</b>			
FVC (L)	—	—	Decreased by 0.03*
FEV1 (L)	No significant differences	—	Increased by 0.27*
Tiffeneau Index (%)	No significant differences	—	Increased by 7.50*
PEF (L/min)	—	—	Increased by 0.66*

MF = magnetic fields; COPD = chronic obstructive pulmonary disease; 6MWD = six-minute walking distance;  $\mu\text{m}^2$  = micrometer; TwQ = twitch tension (Kg); MVC = maximal voluntary manoeuvres (Kg); SF-36 = short form; SGRQ = St. George's respiratory questionnaire; FVC = forced vital capacity; FEV1 = forced expiratory volume in 1 second; PEF = peak expiratory flow.

\* Mean; § median.

The present review focused on a total of thirty-six patients with COPD, and on the effects of the application of MF. The main outcomes investigated in the included studies were quadriceps muscle function (*i.e.*, structure, strength, and endurance), exercise capacity (6MWD), health-related quality of life, and pulmonary function tests, as summarized in Table 2. For twenty subjects, the diagnosis of COPD was sustained by the GOLD guidelines and Tiffeneau Index measured at baseline. Demographics data and other characteristics of the patients are illustrated in Table 1. In the majority of cases, the treatment sessions with MF were carried-out in an outpatient setting, and they differed with regard to the duration; frequency of application; dosage; intensity of the MF applied; and type of device used, as shown in Table 1. MF were primarily used for the muscle stimulation of the quadriceps resulting in increased walked distance, muscular strength, and endurance after treatment; additional improvements were also detected in the quality of life domains (Table 2).

## Discussion

The use of MF as a therapy for dysfunctions and pain of the musculoskeletal system has a long-term history, it is a solid concept, and its efficacy is sustained by several studies (36–38). To the best of our knowledge, there are not previously published reviews on the use of MF in respiratory patients, and the trials available are very limited. COPD is an invalidating disease that affects both respiratory and motor functions during the acute and the chronic phase. Data from the literature demonstrate that pulmonary rehabilitation is the most used non-pharmacological intervention provided in order to treat the underlying symptoms of COPD, namely dyspnea and muscle depletion (39–41). An appropriate planning of the rehabilitation activities for COPD patients is a cornerstone in order to reduce hospital readmission rate, which is very high in this population (42,43).

Data gathered from the retrieved citations showed that MF are used mainly to enhance muscle function as highlighted in the studies by Bustamante and Colleagues (33,34). Muscle weakness of the quadriceps, is one of the most invalidating condition occurring in COPD patients, resulting in exercise limitation, and consequent reduced motor activity. One of the most interesting aspects emerging from this review is represented by the fact that MF seems to have a positive effect on increasing size of the quadricep fibers in COPD patients (34). Lower limbs muscular action, together with all the others motor abilities related to daily life activities, is a fundamental biomechanical component of the walking. Thus, a dedicated treatment addressed to improve muscle strength is perfectly in line with the COPD characteristics, and patients' needs. On the other hand, electrical muscle stimulation has been proven as a suitable tool for the prevention of muscle atrophy in different class of subjects including patients with COPD (44–49).

From the analyzed studies (Table 2) and in particular from the article by Sadlonova et al. (18) MF seems to produce a favorable response in patients with COPD. However, this response was mainly subjective and related to improvement of ventilation and mucociliary clearance as reported by the patients treated with the pulsatile electromagnetic fields in comparison to those in the placebo group. Nevertheless, the amount and density of mucus were not measured; mucus clearance was evaluated only on the basis of patients reported informations, limiting the level of evidence of the observations made. On the other side, the informations regarding quantitative outcomes, such as the spirometry parameters, could demonstrate an improvement of some indices (Tiffeneau, forced expiratory volume in 1 sec., maximal expiratory flow at 25%, maximal expiratory flow at 50%, peak expiratory flow), but it wasn't identified any statistically significant effect of the technique on lung function.

A relevant aspect of this technique is its safe profile: MF are well tolerated and painless. Unexpected or significant side

effects have not been reported even in the case of repeated stimulations; the only absolute contraindication is the presence of metal objects where magnetic stimulation is applied (i.e., pacemaker) (50). From data analyzed in the present review, it was not clear whether MF could also have a direct effect on the airways (i.e., prevent the remodeling of the airways), or on the lung parenchyma (i.e., avoid the alveolar damage): further studies are needed to assess these hypotheses.

Due to the paucity of data on the topic, further experimental research should be encouraged to deeply investigate the effectiveness of MF on COPD respiratory symptoms, especially because it is well established that peripheral muscle function correlates with the severity of the disease: indeed, it is a relevant parameter influencing dyspnea, exercise capacity and it is taken into account in the evaluation of the BODE index (51,52). Depletion of the muscle function is related to utilization of health care resources (53). Moreover, from a clinical perspective, it's noteworthy that many patients with COPD may have an exercise capacity limited by muscle fatigability rather than dyspnea, even when the respiratory function is impaired (54). Basing on the data currently available, it was not possible to get detailed informations about the operating costs of magnetic stimulation for patients with COPD. However, in one study (33) the authors highlighted that the stimulating device used in their research had a high cost.

### Limitations

The present review bears a major limitation: since the topic is very narrowed, we found only a very small number of citations. On the other hand, the additional searching in the grey literature (materials outside of the traditional academic publishing) did not return supplementary data; this confirmed the underestimation of the use of MF in COPD, and of the related lack of knowledge. Moreover, although our findings are based on the analysis of experimental studies, the very small number of patients included in these analyses make it impossible to extend the results reported here to the whole COPD population. Hence, the information discussed here should be considered with caution. However, in spite of the above listed limitations, we believe that the paucity of data could represent rather a positive encouragement for further and additional investigations to improve our knowledge on the application of MF in COPD, particularly in the regard of the respiratory issues.

### Conclusions

We found that the use of MF is described in the literature for the treatment of muscular impairment (quadriceps), and respiratory symptoms in patients with COPD. To date very few investigations have been conducted to explore the effectiveness of MF in COPD; it seems that there is an increasing interest in MF as a therapeutic option in patients with COPD to counteract muscular depletion. In fact, most of the studies found that most relevant studies were published in the field of muscular rehabilitation rather than that of respiratory system. This aspect remains to be clearly defined, and further studies are necessary to clarify the potential direct effect of magnetic stimulation on the airways and lungs. At the

same time, basing on the available informations, it seems that MF represent a feasible, well tolerated, safe therapeutic option, for the treatment of motor-related COPD symptoms. Furthermore, it should be better investigated which groups of patients with COPD (and at which stage) can benefit most from magnetic stimulation treatment, and indicated more precisely the fields and modalities of applications of this therapy.

### Acknowledgment

We thank Dr Paolo Mondardini (Istituto di Medicina dello Sport, Bologna, Italy) for his suggestions.

### Declaration of interest

The authors have no conflicts of interest to declare.

### Funding

The authors have not received any financial support.

### ORCID

Massimiliano Polastri  <http://orcid.org/0000-0001-8548-0813>

Angela Maria Grazia Pacilli  <http://orcid.org/0000-0003-1976-8936>

Stefano Nava  <http://orcid.org/0000-0002-8404-2085>

### References

- Kordiuikov EV. Magnetotherapy of patients suffering from obliterating diseases of the peripheral vessels. *Vopr Kurortol Fizioter Lech Fiz Kult.* 1969;34(3):227–9.
- Degen IL. Magnetotherapy of humeroscapular peri-arthritis. *Ortop Travmatol Protez.* 1974;35(3):66–68.
- Solov'eva GR. Apparatus and areas of application for low-frequency magnetotherapy. *Med Tekh.* 1974;3(0):41–46.
- Skrinnik AV. Current status of the problem of magnetotherapy and the use of magnetic fields in ophthalmology. *Oftalmol Zh.* 1979;34(8):500–505.
- Claisse RH, Baron JB, Fauchier P, Miaux Y. Magnetotherapy in the treatment of ankylosing spondylarthritis. *Acta Belg Med Phys.* 1986;9(1):15–36.
- Ivanov SG, Smirnov VV, Solov'eva FV, Liashevskaja SP, Selezneva L. The magnetotherapy of hypertension patients. *Ter Arkh.* 1990;62(9):71–74.
- Binder A, Parr G, Hazleman B, Fitton-Jackson S. Pulsed electromagnetic field therapy of persistent rotator cuff tendinitis. A double-blind controlled-assessment. *Lancet* 1984;1(8379):695–698.
- Brighton CT, Black J, Friedenber ZB, Esterhai JL, Day LJ, Connolly JF. A multicenter study of the treatment of non-union with constant direct current. *J Bone Joint Surg Am.* 1981;63(1):2–13.
- He Z, Selvamurugan N, Wharshaw J, Partridge NC. Pulsed electromagnetic fields inhibit human osteoclast formation and gene expression via osteoblasts. *Bone* 2018;106:194–203.
- Ross CL, Teli T, Harrison BS. Electromagnetic field devices and their effects on nociception and peripheral inflammatory pain mechanisms. *Altern Ther Health Med.* 2016;22(3):52–64.
- Servodio Iammarrone C, Cadossi M, Sambri A, Grosso E, Corrado B, Servodio Iammarrone F. Is there a role of pulsed electromagnetic fields in management of patellofemoral pain syndrome? *Bioelectromagnetics* 2016;37(2):81–88.
- Griffin XL, Costa ML, Parsons N, Smith N. Electromagnetic field stimulation for treating delayed union or non-union of long bone fractures in adults. *Cochrane Database Syst Rev.* 2011;(4):CD008471.

13. Paolucci T, Piccinini G, Iosa M., Piermattei C, de Angelis S, Grasso MR, et al. Efficacy of extremely low-frequency magnetic field in fibromyalgia pain: a pilot study. *J Rehabil Res Dev.* 2016;53(6):1023–1034.
14. Cichoń N, Bijak M, Miller E, Saluk J. Extremely low frequency electromagnetic field (ELF-EMF) reduces oxidative stress and improves functional and psychological status in ischemic stroke patients. *Bioelectromagnetics* 2017;38(5):386–396.
15. Kanat E, Alp A, Yurtkuran M. Magnetotherapy in hand osteoarthritis: a pilot trial. *Complement Ther Med.* 2013;21(6):603–608.
16. Sieroń A, Cieślak G. Application of variable magnetic fields in medicine-15 years experience. *Wiad Lek* 2003;56(9-10):434–441.
17. Servodio Iammarrone C, Cadossi M, Sambri A, Grosso E, Corrado B, Servodio Iammarrone F. Is there a role of pulsed electromagnetic fields in management of patellofemoral pain syndrome? *Bioelectromagnetics* 2016;37(2):81–88.
18. Sadlonova J, Korpas J, Vrabec M, Salat D, Buchancova J, Kudlicka J. The effect of the pulsatile electromagnetic field in patients suffering from chronic obstructive pulmonary disease and bronchial asthma. *Bratisl Lek Listy.* 2002;103(7-8):260–265.
19. Tenforde TS, Kaune WT. Interaction of extremely low frequency electric and magnetic fields with humans. *Health Phys.* 1987;53(6):585–606.
20. Global Initiative for Chronic Obstructive Lung Disease. Global strategy for the diagnosis, management, and prevention of chronic obstructive pulmonary disease. Available at: <http://www.goldcopd.org> 2017. (Accessed 30 November 2017).
21. Ye X, Wang M, Xiao H. Echo intensity of the rectus femoris in stable COPD patients. *Int J Chron Obstruct Pulmon Dis.* 2017;53:3007–3015.
22. Charusisin N, Dacha S, Gosselink R, Decramer M, van Leupoldt A, Reijnders T, et al. Respiratory muscle function and exercise limitation in patients with chronic obstructive pulmonary disease: a review. *Expert Rev Respir Med.* 2018;12(1):67–79.
23. Abdulai RM, Jensen T, Patel NR, Polkey MI, Jansson P, Celli BR, et al. Deterioration of limb muscle function during acute exacerbation of chronic obstructive pulmonary disease. *Am J Respir Crit Care Med.* 2018;197(4):433–449.
24. Halphin DM, Miravittles M, Metzendorf N, Celli B. Impact and prevention of severe exacerbations of COPD: a review of the evidence. *Int J Chron Obstruct Pulmon Dis.* 2017;12:2891–2908.
25. Broderick J, Mc Grath C, Cullen K, Talbot D, Gilmor J, Baily-Scanian M, et al. Effects of pulmonary rehabilitation on exercise capacity and disease impact in patients with chronic obstructive pulmonary disease and obesity. *Physiotherapy* 2017. pii: S0031-9406(17)30081-0.
26. McCarthy B, Casey D, Devane D, Murphy K, Murphy E, Lacasse Y. Pulmonary rehabilitation for chronic obstructive pulmonary disease. *Cochrane Database Syst Rev.* 2015;(2):CD003793.
27. Carone M, Patessio A, Ambrosino N, Baiardi P, Balbi B, Balzano G, et al. Efficacy of pulmonary rehabilitation in chronic respiratory failure (CRF) due to chronic obstructive pulmonary disease (COPD): the maugeri study. *Respir Med.* 2007;101(12):2447–2453.
28. Nava S. Rehabilitation of patients admitted to a respiratory intensive care unit. *Arch Phys Med Rehabil.* 1998;79(7):849–854.
29. Benzo RP, Kramer KM, Hoult JP, Anderson PM, Begue IM, Seifert SJ. Development and feasibility of a home pulmonary rehabilitation programme with health coaching. *Respir Care* 2018;63(2):131–140.
30. Chen Y, Niu M, Zhang X, Quian H, Xie A, Wang X. Effects of home-based lower limb resistance training on muscle strength and functional status in stable COPD patients. *J Clin Nurs.* 2017. doi:10.1111/jocn.14131.
31. ...old CF, Mahal A, Lee AL, Hill CJ, Burge AT, et al. ...ary rehabilitation for people with COPD: a qualitative study exploring the patient perspective. *Chron Respir Dis.* 2017. doi:10.1183/17455017.2017.2317729050.
32. ... V, Di Fede G. Utilization of extremely low frequency electromagnetic fields in chronic disease; five years experience: three case reports. *Electromagn Biol Med.* 2007;26(4):311–313.
33. Bustamante V, López de Santa Maria E, Gorostiza A, Jiménez U, Gáldiz JB. Muscle training with repetitive magnetic stimulation of the quadriceps in severe COPD. *Respir Med.* 2010;104(2):237–245.
34. Bustamante V, Casanova J, López de Santamaria E, Mas S, Sellarés J, Gea J, et al. Redox balance following magnetic stimulation training in the quadriceps of patients with severe COPD. *Free Radic Res.* 2008;42(11-12):939–948.
35. Thiese MS. Observational and interventional study design types: an overview. *Biochem Med (Zagreb)* 2014;24(2):199–210.
36. Vallbona C, Richards T. Evolution of magnetic therapy from alternative to traditional medicine. *Phys Med Rehabil Clin N Am.* 1999;10(3):729–754.
37. Borg MJ, Marcuccio F, Poerio AM, Vangone A. Magnetic fields in physical therapy. Experience in orthopedics and traumatology rehabilitation. *Minerva Med.* 1996;87(10):495–497.
38. Sadlonova J, Korpas J. Personal experience in the use of magnetotherapy in diseases of the musculoskeletal system. *Bratisl Lek Listy.* 1999;100(12):678–681.
39. Bisca GW, Camillo CA, Cavalheri V, Pitta F, Osadnik CR. Peripheral muscle training in patients with chronic obstructive pulmonary disease: novel approaches and recent advances. *Expert Rev Respir Med.* 2017;11(5):413–423.
40. Katajisto M, Laitinen T. Estimating the effectiveness of pulmonary rehabilitation for COPD exacerbations: reduction of hospital inpatient days during the following year. *Int J Chron Obstruct Pulmon Dis.* 2017;12:2763–2769.
41. Blánquez Moreno C, Colungo Francia C, Alvira Balada MC, Kostov B, González-de Paz L, Sisó-Almirall A. Effectiveness of an educational program for respiratory rehabilitation of chronic obstructive pulmonary disease patients in primary care in improving the quality of life, symptoms, and clinical risk. *Aten Primaria.* 2017. doi:10.1016/j.aprim.2017.03.019.
42. Lalmolda C, Coll-Fernández R, Martínez N, Baré M, Teixidó Colet M, Epelde F, et al. Effect of a rehabilitation-based chronic disease management program targeting severe COPD exacerbations on readmission patterns. *Int J Chron Obstruct Pulmon Dis.* 2017;12:2531–2538.
43. Polastri M, Pisani L, Dell'Amore A, Nava S. Revolving door respiratory patients: a rehabilitative perspective. *Monaldi Arch Chest Dis.* 2017;87(3):94–95.
44. Dirks ML, Hansen D, Van Assche A, Dendale P, Van Loon LJ. Neuromuscular electrical stimulation prevents muscle wasting in critically ill comatose patients. *Clin Sci (Lond)* 2015;128(6):357–365.
45. Wageck B, Nunes GS, Silva FL, Damasceno MC, de Noronha M. Application and effects of neuromuscular electrical stimulation in critically ill patients: systematic review. *Med Intensiva.* 2014;38(7):444–454.
46. Kucio C, Niesporek J, Kucio E, Narloch D, Węgrzyn B. Evaluation of the effects of neuromuscular electrical stimulation of the lower limbs combined with pulmonary rehabilitation on exercise tolerance in patients with chronic obstructive pulmonary disease. *J Hum Kinet.* 2016;54:75–82.
47. Chen RC, Li XY, Guan LL, Guo BP, Wu WL, Zhou ZQ, et al. Effectiveness of neuromuscular electrical stimulation for the rehabilitation of moderate-to-severe COPD: a meta-analysis. *Int J Chron Obstruct Pulmon Dis.* 2016;11:2965–2975.
48. Coquart JB, Grosbois JM, Olivier C, Bart F, Castres I, Wallaert B. Home-based neuromuscular electrical stimulation improves exercise tolerance and health-related quality of life in patients with COPD. *Int J Chron Obstruct Pulmon Dis.* 2016;11:1189–1197.
49. Maddocks M, Nolan CM, Man WD, Polkey MI, Hart N, Gao W, et al. Neuromuscular electrical stimulation to improve exercise capacity in patients with severe COPD: a randomised double-blind, placebo-controlled trial. *Lancet Respir Med.* 2016;4(1):27–36.
50. Man WD, Moxham J, Polkey MI. Magnetic stimulation for the measurement of respiratory and skeletal muscle function. *Eur Respir J.* 2004;24(5):846–860.

51. Celli BR, Cote CG, Marin JM, Casanova C, Montes de Oca M, Mendez RA, et al. The body-mass index, airflow obstruction, dyspnea and exercise capacity index in chronic obstructive pulmonary disease. *N Engl J Med.* 2004;350(10):1005–1012.
52. Gosselink R, Troosters T, Decramer M. Peripheral muscle weakness contributes to exercise limitation in COPD. *Am J Respir Crit Care Med.* 1996;153(3):976–80.
53. Decramer M, Gosselink R, Troosters T, Verschueren M, Evers G. Muscle weakness is related to utilization of health care resources in COPD patients. *Eur Respir J.* 1997;10(2):417–423.
54. Killian KJ, Leblanc P, Martin DH, Summers E, Jones NL, Campbell EJ. Exercise capacity and ventilatory, circulatory, and symptom limitation in patients with chronic limitation. *Am Rev Respir Dis.* 1993;146(4):935–940.