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REVIEW

# Cupping (Hijama) in Rheumatic Diseases: The Evidence

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## **ABSTRACT**

Although practiced initially for the alleviation of pain, cupping therapy (Hijama in Arabic) has been used for a variety of medical conditions, including autoimmune and autoinflammatory diseases, with variable outcomes. In recent years, scientific research on the effectiveness of cupping in the treatment of various diseases has accelerated. Relevant literature to identify the types of cupping along with its association with certain rheumatic conditions was screened through a search of the online databases (MEDLINE, PubMed, and Google Scholar) for an indefinite period. Many reports have drawn serious rheumatic side effects which led medical providers to raise the voice against its practice. Moreover, the rare induction of rheumatic conditions has been partly referred to the immunomodulatory effect that cupping exerts in the body. Cupping therapy still needs more evidence to be labelled as therapeutic procedure for rheumatic conditions. Many studies agree that cupping works best when used in combination to pharmacotherapy. Other studies find no clinically significant beneficial role.

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#### INTRODUCTION

Cupping therapy, Hijama in Arabic, is a traditional alternative medicine that has been practiced in different cultures. The earliest recorded references to cupping practice are found in the *Ebers Papyrus*, written by Ancient Egyptians in Hieroglyphics about 1550 B.C. It is noted that cupping use in Egypt dates to 3500 B.C., as documented in hieroglyphic writing. In addition to ancient Egyptians, the Chinese culture, Arabic medicine, Greek

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medicine, Prophetic medicine, and the recent European and American medicine, have all practiced the different types of cupping.<sup>3</sup> Although the therapy has been tradi-

tionally performed for the alleviation of pain, numerous medical conditions have been targeted. The American College of Physicians has listed cupping therapy as one of the several methods of complementary medicine for the treatment of chronic lumbar spine pain.<sup>4</sup> However, cupping therapy is not yet approved for the treatment of any other medical condition.4 In recent years, scientific research on the effectiveness of cupping in the treatment of various diseases has accelerated. Autoimmune and connective tissue diseases are among the common diseases that have been treated with cupping, with variable outcomes. Many reports have drawn serious rheumatologic side effects of cupping which led medical providers to raise the voice against its practice. This review briefly describes the types of cupping and its proposed biological mechanism of action before drawing the role of cupping in certain rheumatological conditions. The aim of this review is to shed light on the use of cupping in rheumatology as a beneficial or inadequate method of treatment based on evidence from the literature.

## **CUPPING TECHNIQUE**

Although several types of cupping exist, such as dry cupping, wet cupping, moving cupping, and fire cupping, the wet and dry types are most commonly used.<sup>5</sup>

Dry cupping involves coating a thick glass cup with alcohol.<sup>6</sup> A flame is applied and, just before extinguishing it, the coated cup is applied to the patient's skin. The flame should heat the air in the cup, and not the cup itself.<sup>7</sup> With the aid of negative pressure, the skin is suctioned into the cup.<sup>8</sup> The cup is kept in place for around 10 to 15 minutes.<sup>9</sup>

Wet cupping involves bloodletting, and has two main forms. One form is referred to as the cupping, puncturing, and cupping method (CPC), which essentially begins with dry cupping. CPC consists of six steps in total: demarcating the skin, sterilising the area, cupping, puncturing, cupping, and sterilizing once again. The other form is the puncturing and cupping (PC) method, which begins with puncturing. Five steps are instilled here: demarcating the skin, sterilising the area, puncturing, cupping, and sterilising one again. CPC is used more commonly in the Middle East, and is traditionally known as "Al-hijamah". The PC method has been commonly used in the Far East and Europe. The puncturing of the step of

# **MECHANISM OF ACTION**

Cupping has been practiced for four main aims: pain reduction, decrease of inflammation, immunomodulation, and haematological adjustment. For these different aims, many mechanisms of action theories have been proposed.

The three main possible theories that can explain mechanisms of pain reduction are: pain-Gate Theory (PGT), Diffuse Noxious Inhibitory Controls (DNICs), and Reflex Zone Theory (RZT). PGT is one of the most influential theories of pain reduction.11 The stimulation of pain receptors increases the frequency of impulses, ultimately leading to closure of the pain gates, and hence, pain reduction. 12 Subadi and colleagues injected rats taken as a control group with complete Freund's adjuvant (CFA) at the footpad. Wet cupping therapy was performed at the paralumbar regions 48 hours after the CFA injection in the rats belonging to the treatment group. Pain threshold was assessed 24 hours thereafter. Heat shock protein 70 (HSP70) and B-endorphin were assessed in the treatment group. The expression of HSP70 and \( \beta \- endorphin \) was significantly higher in the keratinocytes of the treatment group than the control group. Also, the pain threshold after wet cupping therapy was significantly higher in the treatment group than the control group. The authors concluded that wet cupping might be beneficial in pain

management through increased HSP70 and ß-endorphin expression.<sup>13</sup> DNIC describes the inhibitory activity of a wide dynamic range-type of nociceptive spinal neurons triggered by a second, spatially remote, noxious stimulus. Local damage of the skin and capillary vessels induced by cupping may cause a nociceptive stimulus that activates the distraction effects of DNICs.<sup>14</sup> RZT depends on the hypothesis that signs and symptoms of illness related to one dermatome may be reflected in changes in neighbouring dermatomes.<sup>15</sup> When the diseased organ sends a signal to the skin through the autonomic nerves. the skin responds by becoming tender and painful with swelling. Skin receptors are activated when cups are applied to the skin. The entire process will result in the increment of the blood circulation and blood supply to the skin and the internal organs through the neural connections.16

The anti-inflammatory effect of cupping is mainly mediated by nitric oxide (NO). NO mediates vasodilatation, and regulates blood flow and volume. Tagil and colleagues implemented wet cupping to 31 healthy volunteers. Venous blood samples and wet cupping blood samples were taken concurrently. Serum NO, malondialdehyde levels, and activity of superoxide dismutase and myeloperoxidase were measured by spectrophotometer. Wet cupping blood had higher activity of myeloperoxidase, lower activity of superoxide dismutase, higher levels of malondialdehyde and NO compared to the venous blood. The authors concluded that wet cupping helps to remove oxidants and decreases oxidative stress. 18

The immunomodulatory effect of cupping comes through regulating the activity of immunoglobulins and haemoglobin. Numerous studies have assessed the effects of cupping on the immune system. Ahmed et al. showed that cupping significantly reduces the pain and laboratory markers of rheumatoid arthritis activity. Also, it tends to modulate the activity of the innate immune response especially the natural killer cells as well as the adaptive cellular immune through the activity of Soluble Interleukin 2 Receptor SIL-2R.19 Mohammad Reza and colleagues evaluated the Interferon Gamma (IFNy) and Interleukin 4 (IL-4) concentrations in supernatant of vein and cupping blood cultures with or without the presence of phytohemagglutinin (PHA) mitogen. The results showed IFN-y and IL-4 concentrations in cupping blood samples were higher compared to venous blood samples without presentation of PHA mitogen. In the presence of PHA mitogen, the levels of IFN-γ and IL-4 in cupping blood samples were equally low as in venous blood samples which suggested that lymphocytes in cupping blood samples may not have their natural function, so they cannot properly respond to stimulation of mitogen.<sup>20</sup> Li et al. reported that cupping can upregulate the oxyhaemoglobin and deoxyhaemoglobin. The red blood cell works to recognize antigens, and eliminate

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immune complex, tumour cells, and effector cells, as well as bind germs and viruses, and regulate immune function.<sup>21</sup> Guo and colleagues proposed that physical and biological signals induced by cupping activate the neuroendocrine-immune system, which produces the therapeutic effect.<sup>22</sup>

The haematological adjustment of cupping takes place by detoxification. Several studies reported significant differences in many of the biochemical, haematological, and immunological parameters between the venous blood and the cupping blood.<sup>9</sup>

## **METHODS**

A PubMed search was conducted with keywords "cupping", "hijama", "rheumatology", "arthritis", "osteoarthritis", "fibromyalgia", "rheumatoid arthritis", "ankylosing spondylitis", "gout", "psoriasis", "vasculitis", "scleroderma", "lupus", and "myositis" for all available English literature on PubMed Central. All the case reports, case series, cohort studies, and randomised clinical trials reporting the outcomes after the use of any type of cupping for any autoimmune or connective tissue disease were included. Additionally, reference lists of selected manuscripts were checked manually for eligible articles. All articles not indexed on PubMed/MEDLINE were not included. Articles were selected to fit the scope of our topic, reporting the evidence of benefits or inadequacy of cupping therapy in the background of rheumatology.

# RHEUMATOLOGY AND CUPPING

Fibromyalgia

The fibromyalgia syndrome is characterised by chronic widespread pain in combination with fatigue, cognitive disturbances, sleep disorders, and pronounced somatic and/or psychological distress.<sup>23</sup> practical criteria for clinical diagnosis of fibromyalgia that are suitable for use in primary and specialty care and that do not require a tender point examination, and to provide a severity scale for characteristic fibromyalgia symptoms.\ nMETHODS: We performed a multicenter study of 829 previously diagnosed fibromyalgia patients and controls using physician physical and interview examinations, including a widespread pain index (WPI In total, four early randomised controlled trials (RCTs) showed the efficacy of cupping in fibromyalgia. Two systematic reviews concluded that cupping can be more helpful in treating fibromyalgia when combined with acupuncture and pharmacotherapy than pharmacotherapy alone. One prospective case series found that cupping is beneficial, whereas one recent RCT denied any effect of cupping. Figure 1 lists all the papers that discussed the effect of cupping in fibromyalgia.

The first Chinese clinical study on cupping as a fibromyalgia treatment was published in 2003. Cao and Li compared the efficacy of acupuncture, cupping, and pharmacotherapy versus pharmacotherapy with regards to pain and depression of fibromyalgia. The authors concluded that the combination therapy alleviated pain and depression better than a single therapy.5 Li et al. concluded that a combination of acupuncture with cupping therapy and amitriptyline is significantly more effective than amitriptyline an effective therapy for fibromyalgia syndrome when it came to both pain and depression.<sup>24</sup> Jang and colleagues used a multi-central randomised controlled method to evaluate the clinical effect of combination of acupuncture, cupping, and pharmacotherapy for treatment of fibromyalgia syndrome. Similar to the previous study, the authors suggested that the combination of acupuncture, cupping, and amitriptyline alleviates pain and depression better than pharmacotherapy alone or complimentary medicine alone.<sup>25</sup>

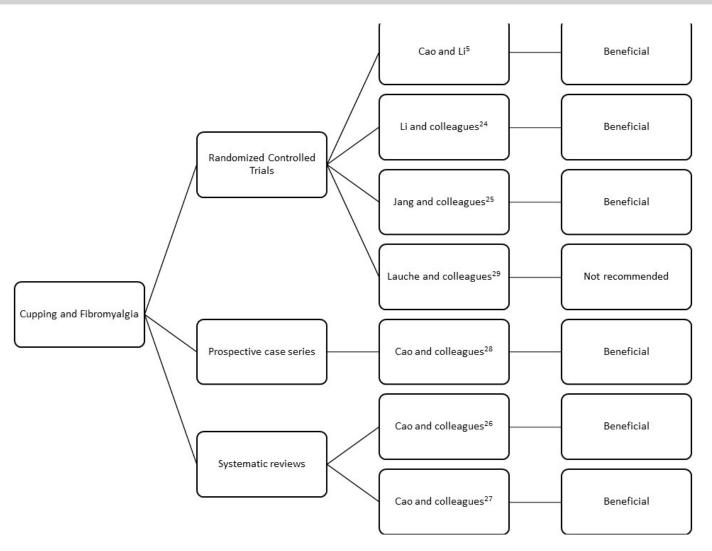
A meta-analysis by Cao and colleagues showed also that a combination of acupuncture and cupping therapy was better than conventional medications for reducing pain and for improving depression scores in fibromyalgia patients. <sup>26</sup> Also, another systematic review by Cao and colleagues concluded that Acupoint stimulation therapy, which includes acupuncture and cupping, appears to be effective in treating fibromyalgia compared with medications. However, the authors strongly suggested that further large trials should be performed with better design than the previous work. <sup>27</sup>

A prospective case series by Cao and colleagues was the first to assess the relationship between fibromyalgia and cupping exclusively. Thirty patients diagnosed with fibromyalgia according to the 1990 criteria of the American College of Rheumatology (ACR) were investigated. Pain, assessed via a 10-point visual analogue scale (VAS), and the number of tender points were noted throughout a 2-week treatment with bamboo cup boiled in herbal decoction. A total of 29 patients completed the whole therapy which suggested a reduction in pain and tender points by 48%.<sup>28</sup>

Working according to the 2010 ACR guidelines for fibromyalgia, Lauche et al. aimed to investigate the efficacy of cupping therapy compared to usual care by assessing the pain intensity as well as functional disability, quality of life, fatigue and sleep quality post cupping compared to usual care and sham treatment. Cupping therapy was more effective than usual care to improve pain intensity and quality of life. However, its effects were small and comparable to those of a sham treatment. The authors concluded that cupping could not be recommended for the treatment of fibromyalgia.<sup>29</sup>

## Osteoarthritis

Osteoarthritis (OA) is a clinical syndrome of joint pain which involves localised loss of articular cartilage and new bone formation in places of destructive bone loss at joint margin. 30" source": "PubMed", "event-place": "Lon-



**Figure 1.** A summary of the randomised controlled trials, prospective case series, and systematic reviews which analysed the effect of cupping in fibromyalgia along with the conclusion of each study.

don", "abstract": "This guideline applies to people with a working diagnosis of osteoarthritis who present for treatment or whose activities of daily living are significantly affected by their osteoarthritis. The management of neck or back pain related to degenerative changes in spine are not part of this guideline. People presenting to health professionals with osteoarthritis complain of joint pain. they do not complain of radiological change. Thus, these guidelines are primarily about the management of older patients presenting for treatment of peripheral joint pain. treatment of the pain itself and of the consequences of such pain for patients who have a working diagnosis of osteoarthritis. The Guideline Development Group (GDG Five RCTs and a single meta-analysis supported the hypothesis that cupping is beneficial for the alleviation of OA pain, whereas three systematic reviews found weak evidence in the support of cupping as a treatment of OA. Figure 2 lists all the studies which analysed the effect of cupping in osteoarthritis.

In 2008, Qin and colleagues compared the results of treating OA with an integrated Chinese that involves cupping and Western medicine group compared to the Western medicine alone as well as traditional Chinese medicine alone. The randomised controlled study concluded that combined Chinese medicine and Western medicine treatment has rapid and definite therapeutic effect in reducing pain and improving mobility of knee joints and daily living ability in Caucasian patients with knee OA.31 safety and tolerability of different therapies in Caucasian patients with osteoarthritis (QA Teut and colleagues assessed the effectiveness of dry cupping exclusively in the treatment of knee OA. The pain and quality of life were improved in the treatment group compared to the control group, although the total number of consumed pain killers was not significantly different between both groups.32 Gao and colleagues performed a randomised controlled trial to assess the difference in pain between a group of knee OA patients treated

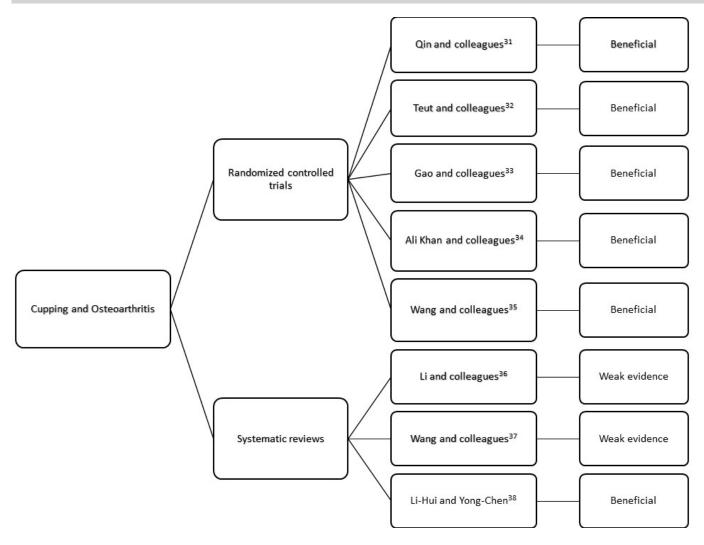


Figure 2. A summary of the randomised controlled trials and systematic reviews which analysed the effect of cupping in osteoarthritis along with the conclusion of each study.

only with acupuncture and another group treated with acupuncture plus cupping. The pain score of joint after treatment, as well as the clinical cure rate, was superior in the combination group.3350 cases in each one. The comprehensive treatment of fire needles at bones combined with cupping and Tuina on local area of affected knee was applied in the comprehensive group. The Ashi points were mainly selected in the fire needles at bones therapy, once every other day. The cupping and Tuina therapy was adopted once a day. The conventional acupuncture was applied in the acupuncture group, in which Dubi (ST 35 As per the RCT by Ali Khan and colleagues, the efficacy of treatment with cupping therapy in relieving signs and symptoms of knee OA was comparable to that of acetaminophen 650 mg three times per day, in terms of analgesia, anti-inflammatory, and resolution of oedema with minimal and temporary side-effects.<sup>34</sup> To evaluate the clinical efficacy of the pricking-cupping therapy for knee OA, a multicentre randomised parallel controlled

trial was performed by Wang and colleagues. This study concluded that the pain score, stiffness score, physical function score and total score of the Western Ontario and McMaster Universities Osteoarthritis Index (WOMAC) were all reduced after 4 weeks of treatment and during the follow-up visit in the pricking-cupping group and the conventional acupuncture group. However, each score and the total score of WOMAC in the pricking-cupping group were lower than those in the conventional acupuncture group after 4 weeks of treatment and during the follow-up visit.<sup>35</sup>

In contrast to all the positive results, a systematic review performed by Li et al. included 7 RCTs. It concluded that only weak evidence can support the hypothesis that cupping therapy can effectively improve the treatment efficacy and physical function in patients with knee OA.<sup>36</sup> Also, Wang and colleagues included 5 studies in their systematic review to conclude that there is weak evidence to support the hypothesis that cupping therapy has ben-

eficial effects on reducing the pain intensity and improving the physical function in patients with knee OA.<sup>37</sup> Recently, Li-Hui and Yong-Chen included 19 eligible studies in their meta-analysis on acupuncture and collaterals cupping therapy for knee OA. The authors found that acupuncture and cupping therapy of traditional Chinese medicine is effective and safe in treating knee OA.<sup>38</sup>

## Rheumatoid Arthritis

Rheumatoid arthritis is an autoimmune disease that has a symmetric, inflammatory peripheral polyarthritis as well as many extra-articular manifestations.

In 2005, Ahmed and colleagues showed that wet cupping significantly reduces the pain and laboratory markers of rheumatoid arthritis (RA) activity. Also, the therapy tends to modulate the activity of the innate immune response mainly the natural killer cells as well as the adaptive cellular immune through the Soluble Interleukin 2 Receptor (SIL-2R).<sup>19</sup> El Sayed and colleagues showed that Al-hijamah combined with conventional can decrease the rheumatoid factor (RF) levels in RA better than what the conventional therapy can do.<sup>39</sup> In some patients who suffered from methotrexate-induced leukopenia, Hijama induced leucocytosis, which counteracted the effect of methotrexate.<sup>40</sup> Cupping has been shown to decrease the levels of serum ferritin which is an inflammatory marker that is elevated in many autoimmune disease including RA.<sup>41</sup>

## Gout

Gout is a broad term for a spectrum of clinical conditions related to an excess of serum uric acid. Gouty arthritis is commonly the first clinical manifestation of gout.<sup>42</sup> Only one RCT concluded that cupping is beneficial in treating acute gouty arthritis, and one prospective cohort study was able to conclude that cupping is helpful.

Zhao et al. assessed the effectiveness of electro-acupuncture with wet cupping in treating gouty arthritis compared to Probenecid. The wet cupping therapy group had a lower serum uric acid than the Probenecid group.<sup>43</sup> Zhang and colleagues treated 34 cases of acute gouty arthritis with wet cupping plus herbal medicine. 21 cases were cured and 13 cases improved.<sup>44</sup>

## Ankylosing spondylitis

Ankylosing spondylitis (AS) is a common chronic inflammatory disorder of unknown aetiology, characterised by enthesitis, sacroiliitis, bone hypertrophy, and new bone formation.<sup>45</sup> One RCT found out that cupping is helpful for the alleviation of AS symptoms, whereas one systematic review found a weak evidence to support that cupping is beneficial for AS.

Wan compared the results of treating a group of patients with AS by acupuncture combined with cupping therapy and treating a control group by simple acupuncture. The author found out that the combination of acupuncture

and cupping therapy in the therapeutic effect on AS is better than simple acupuncture, with shorter therapeutic course and lower recurrence rate. A systematic review by Ma and colleagues which included 5 RCTs concluded that when compared with western medicine alone, cupping therapy plus western medicine has a favourable statistically significant effects on the Bath Ankylosing Spondylitis Functional Index (BASFI) and Bath Ankylosing Disease Activity Index (BASDAI) with low heterogeneity. Furthermore, the effect of the combination of cupping therapy plus western medicine on inflammatory markers is more significant than western medicine alone. However, the authors concluded that when taken together, only weak evidence supported the hypothesis that cupping therapy had potential benefits for patients with AS.

#### Psoriasis

Psoriasis is a T-cell mediated autoimmune inflammatory skin disease characterised by skin surface inflammation, epidermal proliferation, hyperkeratosis, angiogenesis, and abnormal keratinisation.<sup>48</sup> Three case reports in total discussed the topic of cupping therapy in psoriasis. Only one case report found cupping to be beneficial.

Malik and colleagues reported a case of psoriasis with a psoriasis area severity index of 2 who had a decrease in the size of the plaques few days after starting wet cupping. However, some lesions appeared at his elbow after 6 months of treatment.<sup>49</sup> Yu and colleagues reported a case of Köebner phenomenon induction at the cupping sites in a patient with psoriasis.<sup>50</sup> Similarly, Vender and Vender reported a case of a biopsy-proven cupping-induced localized psoriasis in a 45-year-old Asian male.<sup>51</sup>

## Vasculitis

The term vasculitis refers to inflammation directed at blood vessels of any size, which leads to destruction of the vessel wall.<sup>52</sup> One case report supported a beneficence of cupping in treating a form of vasculitis. Another case report illustrated a dermatological side effect of cupping in the use for vasculitis.

Mataix and colleagues reported the case of a 65-yearold male with multiple circular, erythematous, bullous lesions, symmetrically distributed, which occurred after the application of suction cups in the context of polymyalgia rheumatica.<sup>53</sup> However, Wand reported a case of superficial thrombotic phlebitis treated by wet cupping.<sup>54</sup>

## Scleroderma

The subsets of scleroderma include localised scleroderma, limited cutaneous systemic sclerosis, diffuse cutaneous systemic sclerosis, and systemic sclerosis sine scleroderma. <sup>55</sup> One case report described an adverse effect of cupping in scleroderma. One case series study found out that cupping can be beneficial for limited scleroderma.

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Peter reported a case of a man suffering from arterial occlusion developing deep ulcerations after cupping therapy of circumscribed scleroderma on his right thigh. <sup>56</sup> Zhou enrolled 52 patients with limited scleroderma and treated them with needling and wet cupping. The study observed that 14 cases obtained clinical short-term recovery, 29 cases had marked effect, and 9 cases had improvement. <sup>57</sup>

No studies have assessed yet the role of cupping in systemic lupus erythematosus (SLE) or in myositis.

#### CONCLUSION

As many studies have suggested, cupping therapy (Hijama) still needs more evidence to be labelled as a therapeutic procedure for rheumatological conditions. Many studies agree that cupping works best when used in combination to pharmacotherapy. Still, other studies find no significant beneficial role for cupping. While still rare, the induction of rheumatological conditions by cupping could be explained in part by the immunomodulatory effect that cupping exert in the body. Overall, more studies are still needed to define the role of cupping in rheumatology.

#### **CONFLICT OF INTEREST**

The authors declare no conflict of interest.

### **REFERENCES**

- Abbasi M, Norouzadeh R, Gholizadeh M, Heidari S, Gharaboghlo Z. Determining the Clients' knowledge about the Rules of Hijama. Health Spiritual Med Ethics 2014;1(2):33-41.
- Nickel JC. Management of urinary tract infections: historical perspective and current strategies: Part 1--Before antibiotics. J Urol 2005;173(1):21-6.
- 3. Qureshi NA, Ali Gl, Abushanab TS, et al. History of cupping (Hijama): a narrative review of literature. J Integr Med 2017;15(3):172-81.
- Brinkhaus B, Dobos GJ. The Crisis of Medicine and the Benefits of Complex Pain Therapy Procedures Such As Cupping: More Research Needed. Complement Med Res 2019;26(3):145-7.
- Cao H, Han M, Li X, et al. Clinical research evidence of cupping therapy in China: a systematic literature review. BMC Complement Altern Med 2010;10:70.
- King DF, Davis MW. Cupping: An erstwhile common modality of therapy. J Am Acad Dermatol 1983;8(4):563.
- Soliman Y, Hamed N, Khachemoune A. Cupping in dermatology: a critical review and update. Acta Dermatovenerol Alp Pannonica Adriat 2018;27(2).
- 8. Yoo SS, Tausk F. Cupping: East meets West. Int J Dermatol 2004;43(9):664-5.
- Al-Bedah AMN, Elsubai IS, Qureshi NA, et al. The medical perspective of cupping therapy: Effects and mechanisms of action. J Tradit Complement Med 2018;9(2):90-7.
- Mahmoud HS ESS. Methods of Wet Cupping Therapy (Al-Hijamah): In Light of Modern Medicine and Prophetic Medicine. Altern Integr Med 2013;02(03).
- 11. Moayedi M, Davis KD. Theories of pain: from specificity to gate control. J Neurophysiol 2013;109(1):5-12.
- Ahmadi A, Schwebel DC, Rezaei M. The efficacy of wet-cupping in the treatment of tension and migraine headache. Am J Chin Med 2008;36(1):37-44.
- 13. Subadi I, Nugraha B, Laswati H, Josomuljono H. Pain Relief with Wet Cupping Therapy in Rats is Mediated by Heat Shock Protein

- 70 and β-Endorphin. Iran J Med Sci 2017;42(4):384-91.
- 14. Melzack R, Wall PD. On the nature of cutaneous sensory mechanisms. Brain J Neurol 1962;85:331-56.
- Tiran D, Chummun H. The physiological basis of reflexology and its use as a potential diagnostic tool. Complement Ther Clin Pract 2005;11(1):58-64.
- Aboushanab TS, AlSanad S. Cupping Therapy: An Overview from a Modern Medicine Perspective. J Acupunct Meridian Stud 2018;11(3):83-7.
- 17. Faraci FM. Role of nitric oxide in regulation of basilar artery tone in vivo. Am J Physiol 1990;259(4 Pt 2):H1216-21.
- Tagil SM, Celik HT, Ciftci S, et al. Wet-cupping removes oxidants and decreases oxidative stress. Complement Ther Med 2014;22(6):1032-6.
- Ahmed SM, Madbouly NH, Maklad SS, Abu-Shady EA. Immunomodulatory effects of blood letting cupping therapy in patients with rheumatoid arthritis. Egypt J Immunol 2005;12(2):39-51.
- 20. Reza M, vaez Mahdavi MR, Ghazanfari T, Aghajani M, Danyali F, Naseri M. Evaluation of the Effects of Traditional Cupping on the Biochemical, Hematological and Immunological Factors of Human Venous Blood. Compend Essays Altern Ther.
- Li T, Li Y, Lin Y, Li K. Significant and sustaining elevation of blood oxygen induced by Chinese cupping therapy as assessed by near-infrared spectroscopy. Biomed Opt Express 2016;8(1):223-9.
- 22. Guo Y, Chen B, Wang D, Li MY, Lim CH, Guo Y, et al. Cupping regulates local immunomodulation to activate neural-endocrine-immune worknet. Complement Ther Clin Pract 2017;28:1-3.
- 23. Wolfe F, Clauw DJ, Fitzcharles M-A, Goldenberg DL, Katz RS, Mease P, et al. The American College of Rheumatology preliminary diagnostic criteria for fibromyalgia and measurement of symptom severity. Arthritis Care Res 2010;62(5):600-10.
- 24. Li C, Fu X, Jiang Z, Yang XG, Huang SQ, Wang QF, et al. [Clinical study on combination of acupuncture, cupping and medicine for treatment of fibromyalgia syndrome]. Zhongguo Zhen Jiu Chin Acupunct Moxibustion 2006;26(1):8-10.
- 25. Jang Z-Y, Li C-D, Qiu L, Guo JH, He LN, Yue Y, et al. [Combination of acupuncture, cupping and medicine for treatment of fibromyalgia syndrome: a multi-central randomized controlled trial]. Zhongguo Zhen Jiu Chin Acupunct Moxibustion 2010;30(4):265-9.
- 26. Cao H, Liu J, Lewith GT. Traditional Chinese Medicine for treatment of fibromyalgia: a systematic review of randomized controlled trials. J Altern Complement Med N Y N 2010;16(4):397-409.
- Cao H, Li X, Han M, Liu J. Acupoint stimulation for fibromyalgia: a systematic review of randomized controlled trials. Evid-Based Complement Altern Med ECAM 2013;2013:362831.
- 28. Cao H, Hu H, Colagiuri B, Liu J. Medicinal cupping therapy in 30 patients with fibromyalgia: a case series observation. Forsch Komplementarmedizin 2006 2011;18(3):122-6.
- Lauche R, Spitzer J, Schwahn B, Ostermann T, Bernardy K, Cramer H, et al. Efficacy of cupping therapy in patients with the fibromyalgia syndrome-a randomised placebo controlled trial. Sci Rep 2016:6.
- National Collaborating Centre for Chronic Conditions (UK).
   Osteoarthritis: National Clinical Guideline for Care and Management in Adults. Royal College of Physicians (UK); 2008.
- 31. Qin X-Y, Li X-X, Berghea F, Suteanu S. [Comparative study on Chinese medicine and western medicine for treatment of osteoarthritis of the knee in Caucasian patients]. Zhongguo Zhen Jiu Chin Acupunct Moxibustion 2008;28(6):459-62.
- Teut M, Kaiser S, Ortiz M, Roll S, Binting S, Willich SN, et al. Pulsatile dry cupping in patients with osteoarthritis of the knee - a randomized controlled exploratory trial. BMC Complement Altern Med 2012:12:184
- 33. Gao Y-L, Yao J-H, Guo J-X. [Clinical observation on fire needles at bones combined with cupping and Tuina for knee osteoarthritistis]. Zhongguo Zhen Jiu Chin Acupunct Moxibustion 2013;33(8):697-699.
- 34. Khan AA, Jahangir U, Urooj S. Management of knee osteoarthritis with cupping therapy. J Adv Pharm Technol Res 2013;4(4):217-23.

- 35. Wang B, Liu X, Hu Z, Sun A, Ma Y, Chen Yingying, et al. [YANG's pricking-cupping therapy for knee osteoarthritis: a multi-center randomized controlled trial]. Zhongguo Zhen Jiu Chin Acupunct Moxibustion 2016;36(2):113-18.
- 36. Li J-Q, Guo W, Sun Z-G, Huang QS, Lee EY, Wang Y, et al. Cupping therapy for treating knee osteoarthritis: The evidence from systematic review and meta-analysis. Complement Ther Clin Pract 2017;28:152-160.
- 37. Wang Y-L, An C-M, Song S, Lei F-L, Wang Y. Cupping Therapy for Knee Osteoarthritis: A Synthesis of Evidence. Complement Med Res 2018;25(4):249-55.
- 38. Zhang Li-Hui ZY-C, Zhang Li-Hui ZY-C. Systematic evaluation and Meta-analysis of acupuncture and collaterals cupping therapy for Knee Osteoarthritis. TMR Mod Herb Med 2019;2(1):36-47.
- 39. El Sayed SM, Baghdadi H, Abou-Taleb A, Mahmoud HS, Maria RA, Ahmed NS, et al. Al-hijamah and oral honey for treating thalassemia, conditions of iron overload, and hyperferremia: toward improving the therapeutic outcomes. J Blood Med 2014;5:219-37.
- Mahmoud HS ESS. Medical and Scientific Bases of Wet Cupping Therapy (Al-hijamah): in Light of Modern Medicine and Prophetic Medicine. Altern Integr Med 2013;02(05).
- 41. Baghdadi H, Abdel-Aziz N, Ahmed NS, Mahmoud HS, Barghash A, Nasrat A, Nabo MM, et al. Ameliorating Role Exerted by Al-Hijamah in Autoimmune Diseases: Effect on Serum Autoantibodies and Inflammatory Mediators. Int J Health Sci 2015;9(2):207-32.
- 42. Khanna D, Fitzgerald JD, Khanna PP, Bae S, Singh MK, Neogi T, Pillinger MH, et al. 2012 American College of Rheumatology guidelines for management of gout. Part 1: systematic nonpharmacologic and pharmacologic therapeutic approaches to hyperuricemia. Arthritis Care Res 2012;64(10):1431-46.
- 43. Zhao Q, Liu J, Qu X, Li W, Wang S, Gao Y, et al. [Observation on therapeutic effect of electroacupuncture plus blood-letting puncture and cupping combined with diet intervention for treatment of acute gouty arthritis]. Zhongguo Zhen Jiu Chin Acupunct Moxibustion 2009;29(9):711-3.
- 44. Zhang S-J, Liu J-P, He K-Q. Treatment of acute gouty arthritis by blood-letting cupping plus herbal medicine. J Tradit Chin Med Chung Tsa Chih Ying Wen Pan 2010;30(1):18-20.
- 45. Jones G. What's new in osteoarthritis pathogenesis? Intern Med J 2016:46(2):229-36.
- 46. Wan X. [Clinical observation on acupuncture combined with cupping therapy for treatment of ankylosing spondylitis]. Zhongguo Zhen Jiu Chin Acupunct Moxibustion. 2005;25(8):551-552.
- 47. Ma S-Y, Wang Y, Xu J-Q, Zheng L. Cupping therapy for treating ankylosing spondylitis: The evidence from systematic review and meta-analysis. Complement Ther Clin Pract 2018;32:187-94.
- 48. Rahman M, Alam K, Ahmad MZ, Gupta G, Afzal M, Akhter S, et al. Classical to current approach for treatment of psoriasis: a review. Endocr Metab Immune Disord Drug Targets 2012;12(3):287-302.
- 49. Malik IA, Akhter S, Kamal MA. Treatment of psoriasis by using Hijamah: A case report. Saudi J Biol Sci 2015;22(1):117-21.
- 50. Yu R-X, Hui Y, Li C-R. Köebner phenomenon induced by cupping therapy in a psoriasis patient. Dermatol Online J 2013;19(6):18575.
- Vender R, Vender R. Paradoxical, Cupping-Induced Localized Psoriasis: A Koebner Phenomenon. J Cutan Med Surg 2015;19(3):320-2.
- 52. Lakdawala N, Fedeles F. Vasculitis: Kids are not just little people. Clin Dermatol 2017;35(6):530-40.
- Mataix J, Belinchón I, Bañuls J, Pastor N, Betlloch I. [Skin lesions from the application of suction cups for therapeutic purposes]. Actas Dermosifiliogr 2006;97(3):212-4.
- 54. Wang T. [Superficial thrombotic phlebitis treated by blood-letting and cupping therapy]. Zhongguo Zhen Jiu Chin Acupunct Moxibustion 2011;31(1):22.
- Fett N. Scleroderma: Nomenclature, etiology, pathogenesis, prognosis, and treatments: Facts and controversies. Clin Dermatol 2013;31(4):432-7.
- Peter L. [Necrotizing circumscribed scleroderma after cupping therapy in arterial occlusion]. Z Hautkr 1986;61(13):953-7.

 Zhou Y. Observation on therapeutic effects of combined surrounding needling, bloodletting with cupping therapy for localized scleroderma. J Acupunct Tuina Sci 2011;9(3):154-5.