Balneotherapy in Medicine: A Review

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Abstract

Bathing in water (balneotherapy or spa therapy) has been frequently and widely used in classical medicine as a cure for diseases. This paper reviews the present literature on the use of balneotherapy in dermatologic, chronic musculoskeletal (inflammatory and non-inflammatory), metabolic and psychological conditions.

We performed a systematic review on related papers appearing in the Medline and Cochrane Library database from 1966 to 2003 that included randomized controlled and non-randomized clinical trials using balneotherapy. We also determined to reflect where possible the chemical compositions of spas.

The major dermatologic and musculoskeletal diseases that are frequently treated by balneotherapy with a remarkable rate of success are atopic dermatitis, psoriasis, rheumatoid arthritis (RA), ankylosing spondylitis, osteoarthritis and low back pain. Moreover, the effects of spa therapy on several metabolic conditions are discussed. The mechanisms by which broad spectrums of diseases respond to spa therapy probably incorporate chemical, thermal and mechanical effects.

The importance of balneotherapy either alone or as complement to other therapies should be considered after, or accompanying, orthodox medical treatments.

Key words: spa therapy, balneotherapy, review

Introduction

Over the past decades, a re-assessment of the use of mineral water for the treatment of several diseases has taken place around the world (1). Many rheumatologists and dermatologists now acknowledge the medical significance of bathing. The term balneotherapy comes from the latin balneum (bath). The term is classically used for bathing in thermal or mineral waters. Bathing is usually combined with other treatments, such as physical exercise, hydrotherapy, and mud packs.

Thousands of health resort areas have developed around these hot springs. Spa resorts are differentiated according to their location (sea side, mountain area) and the chemical composition of their mineral water. They are also classified as being low mineralized (0.6-2 g/l), mildly mineralized (>2-10 g/l) or highly mineralized (>10 g/l). Water temperature is described as

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being cold (<20°C), hypothermal (20–30°C), thermal (>30–40°C); or hyperthermal (>40°C) (2).

Absorption of minerals through the skin seems to be limited. The dermatological therapeutic effect would therefore appear to lie in a local interaction between the mineral water and the structure of the skin surface.

The effects of spa therapy can be divided into three categories: mechanical, thermal and chemical.

Mechanical effects

Immersion allows the patient to mobilize joints and strengthen muscles with minimal discomfort. This hydrostatic effect is increased when the water is more concentrated. Hydrostatic pressure also causes displacement of fluids from the extremities to the trunk, thus causing hemodilution and increased diuresis. It has been shown that immersion for 1 hour increases water excretion by about 50% (3).

Thermal effects

The hot water causes superficial vasodilation and it has been shown to reduce vascular spasm and stasis in the nail bed and conjunctiva (4). The in vivo proliferative response of human peripheral blood lymphocytes to phytohemagglutinin and concanavalin A was enhanced markedly when cultured at

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Table 1	Balneotherapy in	patients with	dermatologic	disorders

Disease	Authors	Mode of therapy	n	Treatment duration	Efficacy	*Evidence (Design and journal type)
Atopic dermatitis	Kubota et al. (8)	Acidic hot-spring bath pH: 2.0 plus Mn 1.4 mg/l, I 0.3 mg/l	70	Median of 2 months	Significant clearing of skin lesions and reduction in itch	B (B)
	Shani et al. (9)	Balneotherapy in the Dead Sea	1408	4 to 6 weeks	Complete clearance of lesions in 90% of patients	A (B)
Psoriasis	Halvey et al. (11)	Dead Sea balneotherapy	25	3 weeks	Mild improvement comparing to those treated with common salts bath	B (A)
	Leaute-labreze et al. (12)	Saline water balneotherapy (sodium concentration, 250 g/l; magnesium, 980 mg/l)	71	3 weeks	Minor therapeutic effects with saline spa water alone, and no beneficial effect of bathing to enhance phototherapy	C (A)
	Pinton et al. (18)	Immersion and drinking selenium rich spa (70 µg/l)	92	3 weeks	Improvement in psoriatic plaques	A (B)
Cutaneous microcirculation	Hartmann et al. (15)	Balneotherapy with CO ₂ (1200 mg/kg water) water	18	20 minutes	Cutaneous vasodilation and increased oxygen utilization	A (A)

* Evidence. A: effective, B: probably effective, C: may not be effective.

Design and journal type: A: Randomized clinical trial (RCT) in a major journal, a: RCT in a non-major journal, B: non-randomized clinical study in a major journal, b: non-randomized clinical study in a non-major journal.

40°C compared with the conventional temperature of 37°C (5).

Chemical effects

The solutes or additives in spa water act primarily on the skin, but there is no doubt that under certain conditions resorption of minerals is possible. The composition and physical properties of various spa waters vary. They are salty, sulfurous, bicarbonated, carbonic, radon-rich, selenium-rich, arsenical and ferruginous, etc. It is still not clear which elements are essential and what is the ideal concentration of each element in order to attain an optimal response to treatment.

In this report, we screened published papers from 1966 to 2003 for randomized controlled trials (RCT) and nonrandomized clinical studies of balneotherapy on dermatological, chronic musculoskeletal (inflammatory and non-inflammatory), metabolic and psychological conditions. Key words in the studies were: balneotherapy, spa therapy, RCT and clinical trials. To perform an adequate assessment the language of the publications had to be mainly English. Of the studies not written in English, those with an English abstract that had sufficient information about the study design were selected for this review. We also provided where possible the chemical compositions of the spas. Based on the results and design of each study, and the journal in which the paper published, we classified each study under the following criteria: Evidence; A: effective, B: probably effective, C: may not be effective. Journal type and design; A: RCT in a major journal, a: RCT in a non-major journal, B: non-randomized clinical study in a major journal, b: non-randomized clinical study in a non-major journal. The summarized effects of balneotherapy are shown in Tables 1 to 5.

Dermatologic effects

Inoue et al. (6) reported that balneotherapy using Kusatsu hot spring water (Japan) is useful for controlling the skin

symptoms of acute flares/exacerbations of refractory cases of atopic dermatitis. It is now widely accepted that patients with atopic dermatitis are prone to cutaneous Staphylococcus aureus (S. aureus) infection during phases of acute exacerbation and also an increased density of S. aureus is found to correlate well with the severity of skin manifestations (7, 8). It has been demonstrated that S. aureus on the skin surface decreased in number or disappeared after Kusatsu balneotherapy (6). The bactericidal activity of hot spring water was expressed by manganese and iodide ions in acidic conditions. Balneotherapy using acidic hot-spring water (Kusatsu, Japan) was shown to be useful for controlling the skin symptoms of acute flares of refractory cases of atopic dermatitis in comparison to a hot plain-water shower (8).

The Dead Sea is a famous place for its balneologic properties and its effects, especially on ailments of dermatologic and rheumatologic origin. The Dead Sea has a salt content of about 320 g/l, of which potassium chloride, magnesium chloride, calcium chloride and sodium chloride are the major components. The average mineral salt contents (g/l) are as follows: sodium, 5.44; potassium, 4.16; calcium, 65.28; magnesium, 15.69; chloride, 24.96; sulfate, 24.96 and carbonate, 74.24. Total concentration of salt and minerals are 32%, compared to a total concentration of 3% in the ocean (2). Two studies provided evidence for the therapeutic potential of Dead Sea spa therapy for atopic dermatitis (9, 10). Complete clearance of lesions was recorded in 90% of 1408 patients after 4 to 6 weeks therapy at the Dead Sea area. A reduction in itching was recorded during the first week of stay at the Dead Sea area. The percentage of patients who improved during the spring and summer (91%) was higher than in the autumn (86%) and winter (74%) (9). Giryes et al. (10) reported the efficacy of Dead Sea climatotherapy for atopic dermatitis in a non-published study. The climatotherapy regimen consisted of daily sun exposure (maximum, 3-4 hours a day), bathing in Dead Sea water (20 minutes twice a day) and free application of emollients. Acute exacerbation of atopic dermatitis must first be given specific pharmacologic treatment, however, bathing can prepare the skin for the application of moisturizers. Rest and the healthy environment provided by spas can also be positive factor in healing atopic dermatitis.

A prospective, double blind, controlled study (11) evaluated the therapeutic effect of Dead Sea salts in patients with psoriasis. Twenty-five patients with psoriasis vulgaris were randomly allocated to 2 groups treated with either Dead Sea salt baths or common salt baths. After 3 weeks of treatment, mild improvement was observed in patients treated with Dead Sea salts compared with those treated with common salts. However, saline spa water alone at Salies de Bearn in France (sodium concentration, 250 g/l; magnesium, 980 mg/l) was reported to have a minor therapeutic effect on psoriasis compared with UV-B exposure in an RCT on 90 patients (12).

Acne vulgaris is another dermatologic disease that benefits from balneotherapy. A non-randomized clinical study of 86 patients treated for acne vulgaris in the Dead Sea area showed a significant improvement manifested by a reduced number of comedones and pustules (9).

Sulfur-rich spas attract special interest for their dermatologic effects. The sulfur that penetrates the skin is oxidized and evokes various physiologic responses in the skin, such as vasodilation in the microcirculation, an analgesic influence on the pain receptors, and inhibition of the immune response. Sulfur also interacts with oxygen radicals in the deeper layers of the epidermis, producing sulfur and disulfur hydrogen, which may be transformed into pentathionic acid, and this may be the source of the antibactericidal and antifungal activity of sulfur water (2). The therapeutic action of sulfur water is related mainly to sulfur's keratolytic effect, resulting in peeling (13).

Schempp et al. (14) demonstrated in both in vivo (a 5% concentration of $MgCl_2$) and in vitro (a 1% concentration of $MgCl_2$) studies that magnesium ions specifically inhibit the antigen-presenting capacity of Langerhans cells and may thus

contribute to the efficacy of magnesium-rich spa water in the treatment of inflammatory skin diseases.

Therapeutic activities of CO_2 water baths (700–1300 mg CO_2 per kg water) are explained by a synergism between hydrostatic pressure and the chemical properties of carbon dioxide that acts directly on the blood vessels of the skin, causing vasodilation and increased oxygen utilization (15).

Balneotherapy with arsenical-ferruginose (no data on concentrations) water from the spa at Terme di Levico (Italy) showed effectiveness in the treatment of cervico-vaginitis (n=20) in comparison to placebo suppositories (n=10). Both clinical signs and symptoms were reduced in the intervention group with good tolerability (16).

Apart from immersion, drinking spa water has also been investigated for treatment of dermatologic conditions. Drinking of low-salt Avene (France) spring water (sodium, 4.9 mg/l; magnesium, 22.5 mg/l; calcium, 44.3 mg/l; bicarbonate, 234.8 mg/l) for a period of 18 days normalized the intestinal permeability in patients with atopic dermatitis (17). Drinking and immersion in a selenium-rich spa water (selenate 70 µg/l) for three weeks at the care center of La Roche-Posay (France) demonstrated an improvement in patients with psoriasis (18). Patients who responded to treatment had a significant increase in their plasma selenium level. In patients with psoriasis, inflammatory reactions in the skin may lead to an increased loss of selenium (18).

Chronic musculoskeletal effects

Balneotherapy has been practiced for centuries in the management of chronic musculoskeletal diseases. It is based on the indigenous natural remedies of the spas, such as thermal and mineral water and gases (CO_2 , radon). The net beneficial effect of balneotherapy on musculoskeletal diseases is probably a result of mechanical, thermal and chemical effects.

Table 2 Balneotherapy in patients with chronic inflammatory musculoskeletal diseases

Disease	Authors	Mode of therapy	n	Treatment duration	Efficacy	*Evidence (Design and journal type)
Rheumatoid arthritis	Sukenik et al. (19)	Sulfur baths, mud packs and combination	40	2 weeks	Improvement in clinical indices up to 3 months	A (A)
	Elkayam et al. (20)	Mineral water baths and mud packs (rich in sodium chloride and sulfate)	41	2 weeks	Temporary improvement in clinical indices	A (A)
	Sukenik et al. (21)	Dead Sea, sulfur bath or combination	36	3 months	Improvement in clinical indices in all three intervention groups	A (A)
	Franke et al. (23)	Combined radon (1.3 KBq/l) carbon dioxide (1.6 g/l) spa	60	4 weeks	Lower pain intensity after 6 months comparing to taking only a carbon dioxide bath	B (a)
Ankylosing spondylitis	Tubergen et al. (24)	Combination of spa-exercise therapy	120	3 weeks	Improvement in clinical indices up to 40 weeks	A (A)
	Tishler et al. (25)	Combination of hot mineral baths (38°C) and mud packs (45°C)	14	2 weeks	Improvement in morning stiffness and overall well-being	B (B)
Fibromyalgia	Buskila et al. (27)	Sulfur baths (2000 mg/l)	48	10 days	Improvement in clinical indices up to 3 months	A (A)

Inflammatory diseases

Sukenik et al. (19) reported from an RCT that a two-week period of therapy with sulfur baths or mud packs (no data on mineral concentration), alone or in combination was effective in the treatment of patients with RA. Statistically significant improvements in most clinical indices were observed for a period of up to 3 months in treatment groups. In a similar study Elkayam et al. (20) allocated 41 RA patients at random to 2 groups. One group was treated with a combination of mineral baths (rich in sodium chloride and sulfate (no evidence on concentrations)) and mud packs for 20 minutes in Tiberias (Isreal), while the second group was treated with tap water only. Both groups showed a significant but only temporary improvement. The first group showed significant improvement in grip strength.

Sukenik et al. (21) again reported from a randomized trial that daily bath in the Dead Sea or a daily sulfur bath (no data on sulfur ions concentration), or a combination of both, resulted in improvements in clinical indices such as duration of morning stiffness, grip strength and number of active joints in 36 patients with RA during a 3 month follow-up period, compared to controls.

Radon spa therapy was found to be effective in treatment of RA (22). Sixty patients were grouped to take baths with either radon-carbon dioxide water (1.3 KBq/l and 1.6 g/l) or artificially enriched carbon dioxide water for four weeks. After six months of follow-up, pain intensity remained lower in patients receiving

radon-carbon dioxide bath than the other group (23).

Ankylosing spondylitis is another chronic inflammatory disease that benefits from balneotherapy (24–26). In a nonrandomized clinical (25) study 14 patients with ankylosing spondylitis were treated for 2 weeks at a Tiberias spa with a combination of hot mineral water baths (38°C) and mud packs (45°C) for 20 minutes (no data on mineral concentration). A significant improvement was observed in morning stiffness, finger to floor distance, and overall well-being as assessed by patients and physicians. Combined spa-exercise therapy (Bad Hofgastein, Austria), alongside standard treatment with drugs and weekly group physical therapy, has been shown to be more effective and show favorable cost-effectiveness and cost-utility ratios compared with orthodox treatment alone in patients with ankylosing spondylitis (26).

The beneficiary effect of balneotherapy was also observed in patients with fibromyalgia. Forty-eight patients with fibromyalgia were randomly assigned to a treatment group receiving sulfur baths (sulfur ions concentration of 2000 mg/l) for 20 minutes, and a control group with no treatment. All participants stayed for 10 days at a Dead Sea spa. Signs and symptoms of the disease improved in the patients receiving sulfur baths and improvement persisted for 3 months (27).

Although balneotherapy does not replace, but rather complements, conventional drug therapy, it is certainly beneficial in suitable cases.

Table 3	Balneotherapy in	patients with chronic non-inflamn	natory musculoskeletal diseases

Disease	Authors	Mode of therapy	n	Treatment duration	Efficacy	*Evidence (Design and journal type)
Osteoarthritis	Nguyen et al. (28)	Spa therapy at Vichey (France)	188	3 weeks	Improvements in pain and functional impairment up to 6 months	A (A)
	Kovacs et al. (29)	Spa therapy (Solute content 1675 mg/l) mainly sodium bicarbonate and fluoride)	58	15 days	Improvement in clinical indices compared to tap water	B (A)
	Green et al. (30)	Hydrotherapy in a deep pool, and home exercise	47	6 weeks	Improvement of symptoms in both groups	C (A)
	Wigler et al. (31)	Dead Sea balneotherapy	33	2 weeks	Improvement in clinical indices specially with the combination of mineral bath and mud packs	A (A)
Low back pain Guillemin et a (32) Constant et al. (33)		Spa therapy (total minerals <500 mg/l, mainly sulfate and sodium) with high pressure under water shower (36°C)	102	3 weeks	Positive short term effectiveness on chronic pain compared to controls	A (A)
	Spa therapy (total minerals 8073 mg/l, mixed bicarbon- ate, chlorine and sodium) and drug therapy compare with drug therapy alone	121	3 weeks	Improvements in signs and symptoms in the combination group	A (A)	
	Konard et al. (34)	Balneotherapy (bicarbonate (445.3 mg/l), sulfate (109.9 mg/l) and calcium (117.2 mg/l)), underwater massage, underwater traction bath	158	4 weeks	Reduction of prescription of analgesics and pain score in all intervention groups	B (B)
Chronic pain	Strauss-Blasche et al. (35)	Balneotherapy with mud and CO_2 applications	387	3 weeks	Seasonal variation in effectiveness, best response was between April and June	B (b)

Non-inflammatory diseases

The aim of balneotherapy in musculoskeletal diseases is to improve the range of joint movements, cause muscle strengthening, relieve muscle spasm, maintain or improve functional mobility, and as a consequence to relieve patients' suffering.

Balneotherapy for osteoarthritis provided satisfactory results (28-31). An RCT study was conducted in patients with osteoarthritis of the hip, knee or lumbar spine (28). Treatment was either spa therapy (n=91) of three weeks duration at Vichey (France) (no data on mineral components) or the usual therapy (n=97) at home. Changes in the assessment criteria after a 6 months follow-up period showed improvement in terms of pain, functional impairment and quality of life in the spa group. The effects of thermal water (36°C for 30 minutes) from Cserkeszolo (Hungary) were appraised in a randomized, double blind study on 58 patients with osteoarthritis of the knee (29). Balneotherapy was performed with spa water (Solute content of 1675 mg/l comprised primarily of sodium bicarbonate along with silicic acid (48 mg/l) and fluoride (1.8 mg/l)) and tap water. Both groups improved in signs and symptoms, but the magnitude of improvement was significantly greater in patients treated with spa water.

Low back pain is another non-inflammatory musculoskeletal disease that benefits from short and long-term effects of spa therapy (32, 33). Constant et al. (33) assessed the overall effectiveness of spa therapy at Saint-Nectaire (France) compared with the usual routine drug therapy in chronic low back pain. In an RCT, 121 patients were allocated to treatment (n=59) and control (n=62) groups. In the treatment group, patients underwent routine drug therapy and spa therapy (total mineral content of 8073 mg/l of mixed bicarbonate, chlorine and sodium composition) 6 days per week for 3 consecutive weeks. In the control group, patients received routine drug therapy. After 3 weeks of intervention and 6 months of follow-up, patients in the treatment group had significant improvement in signs and symptoms of the disease and also drug consumption (analgesic and anti-inflammatory).

Three treatments for non-specific lumbar pain: balneotherapy (total mineral concentration of 901 mg/l, mainly bicarbonate, 445.3 mg/l; sulfate, 109.9 mg/l and calcium, 117.2 mg/l electrolytes), underwater traction bath and underwater massage were assessed in a randomized prospective controlled trial in 158 outpatients (34). Each group was treated for 4 weeks and patients were reviewed at the end of this period and at 12 months following entry to the trial. The prescription of analgesics and pain score were significantly reduced in all three treated groups but there was no difference between the three groups. After 1 year, only consumption of analgesics remained significantly lower in the intervention groups than in the control group who received no treatment.

Strauss-Blasche et al. (35) reported seasonal variation in the effects of spa therapy (including mud and CO_2 applications) on chronic pain. A total of 387 patients with non-inflammatory chronic pain stayed at a spa in Austria for 3 weeks and received 2–4 treatments per day, including mudpacks, massages, and exercise therapy. In different groups of patients for 2 years, pain was measured at the beginning, end, and 6 weeks after spa therapy. The effect of spa therapy on pain was best between April and June and a medium decrease of pain was more likely between October and November. The magnitude of the seasonal variation was greater for back pain than for joint pain.

The effectiveness of balneotherapy in the management of patients with arthritis is attributed to physiological changes like increased diuresis and hemodilution, as well as biomechanical changes like joint unloading, relaxation and increased muscle function (4, 36, 37).

Condition	Authors	Mode of therapy	n	Treatment duration	Efficacy	*Evidence (Design and journal type)
Blood viscosity	Shirakura et al. (39)	Whole body bathing in hyperthermal water (42°C or higher)	7	10–15 minutes	Marked increase of blood viscosity and enhancement in the blood coagulation system	A (b)
Platelet aggregability	Ohtsuka et al. (43)	Hydrotherapy with water temperature between 30 to 40°C	12	4 weeks	Partial improvement of platelet gluthatione metabolism	B (b)
Beta-thromboglobulin	Take et al. (45)	Hydrotherapy with water temperature of 47 and 42°C	5	10 minutes	Plasma beta-thromboglobulin increased with 47°C bathing	B (a)
Bone structure	Ay et al. (48)	Aerobic exercise in a spa resort pool with water temperature of 29–30°C	41	6 months	Anabolic effects on the heel bone	A (A)
Plasma lipids	Strauss-Blasche et al. (49)	Spa therapy (CO ₂), exercise therapy, and dietary measures	395	3 weeks	Mild decrease in total cholesterol, HDL cholesterol and LDL cholesterol	B (b)
Stress hormones	Kuczera et al. (51)	Spa therapy at Wysowa (Poland)	175	20 days	Increase of plasma concentration of ACTH, cortisol, growth hormone and prolactin	A (b)
Plasma homocysteine	Leibetseder et al. (52)	Spa therapy with Sulfur as the main mineral (7.3 mg/l)	40	3 weeks	Reduction of plasma homocysteine in sulfur bath group compare to controls	A (A)

Table 4 Balneotherapy in metabolic conditions

Metabolic effects

Water temperature has an important role for the metabolic effects of balneotherapy. In Japan, the most outstanding feature of balneotherapy is repeated whole body immersion in hot water (42°C or higher). Ohtsuka et al. (38) reported that immersion in water at 42°C for 10 minutes resulted in oxidative stress by increasing the levels of lipid peroxides and decreasing the activities of glutathione peroxides in erythrocytes. Moreover, whole body bathing in hot water (42°C or higher for 10–15 minutes) is reported to induce a marked increase of blood viscosity and an enhancement of the blood coagulation system (39). However, drinking of electrolyte water containing (meq/l) sodium, 21; potassium, 7; calcium, 1 and chloride, 18.5 at midnight was associated with a decrease in blood viscosity from midnight to 8 AM (40).

Platelet glutathione metabolism is impaired among patients with diabetes mellitus and coronary heart disease, in which platelet aggregability is elevated (41, 42). Ohtsuka et al. (43) reported from a clinical controlled trial on 12 type II diabetic patients that four weeks of twice or thrice daily balneotherapy at a spa in Hokkaido (Japan), with the water temperature between 39 to 40°C (no data on mineral compositions), resulted in partial improvement of platelet glutathione metabolism. They concluded that balneotherapy was beneficial for patients whose platelet antioxidative defense system is damaged, such as those with diabetes mellitus and coronary heart disease. However, immersion in hot water (42°C or higher) for 10 minutes was reported to cause an increase in the activity of erythrocyte aldose reductase, which has a positive correlation with hemoglobin A1c (44). The change in erythrocyte aldose reductase activity might aggravate diabetic complications, and therefore, body exposure to hot environmental conditions is better avoided for diabetic patients.

To determine the involvement of activated platelets in the frequent thrombosis after hot hot-spring bathing, Take et al. (45) found that plasma levels of beta-thromboglobulin began to rise at 5 minutes and elevated significantly 10 minutes after the start of 47°C hot spring bathing at Kusatsu (Japan). This effect was not observed with bathing at 42°C. Concentrations of beta-thromboglobulin were significantly correlated with intimamedia thickness of the common carotid artery (46). Acute hyperthermal stress due to hot hot-spring water may also decrease the fibrinolytic capacity, leading to the occurrence of thrombotic events (47).

Some spa resorts have aerobic exercise facilities. Aerobic

exercise in the Ataturk spa resort (Turkey) for 6 months with the water temperature of 29–30°C was determined to have positive anabolic effects on the bone of 41 postmenopausal, sedentary subjects, in comparison to control group who were instructed to maintain their sedentary lifestyle for the duration of the study (48).

Spa therapy has also been studied for its possible lipidlowering effect. Strauss-Blasche et al. (49) reported that a 3week spa therapy (consisting of mud packs, CO_2 baths and massage and exercise therapy) at Bad Tatzmannsdorf (Austria) in 395 patients with musculoskeletal chronic pain over 2 years was associated with a mild decrease of total cholesterol, HDL cholesterol and LDL cholesterol.

Kuczera et al. (50) found that spa therapy (no data on mineral components) in Wysowa (Poland) for 20 days was associated with a significant increase of plasma erythropoietin, iron, ferritin and saturation of transferrin with iron plus an increase in the concentration of stress reaction hormones (51).

Leibetseder et al. (52) provided evidence that sulfur baths (S⁻² concentration of 7.3 mg/l) exert beneficial effects on plasma homocysteine. Plasma homocysteine is a risk factor for cardiovascular diseases (53). Forty subjects were randomized into two equal groups, a treatment group who received stationary spa therapy plus daily sulfur baths, and a control group who received spa therapy alone. Plasma homocysteine was significantly reduced in the sulfur bath group and rose insignificantly in the control group after three weeks stay at the spa resort. The researchers did not find any significant change in the urinary levels of 8-hydroxydeoxy Guanosine in both groups (52). Studies on rats provided evidence of an increase in heat-shock protein expression after bathing for 15 minutes in 41°C hot water for 4 weeks (54, 55). Heat-shock proteins are known as important endogenous cell-protective proteins induced in response to a wide variety of stresses (56, 57).

Psychological effects

A few studies addressed the psychological effects of balneotherapy. Steam baths, saunas and whirlpools are standard equipment in many spa resorts, with the main objective being to relax and strengthen the body and mind, and to prevent development of disease. A medium (3-month) to long term (6-month) intervention at a spa resort with bathing and health education resulted in less psychological tension and self-rated depression, and more vigorousness, among a group of middle-aged and elderly women (58, 59).

Condition	Authors	Mode of therapy	n	Treatment duration	Efficacy	*Evidence (Design and journal type)
Psychological tension	Kamioka et al. (58)	Once a week balneotherapy (41.5°C) with lifestyle edu- cation and physical exercise	56	3 months	Less psychological tension in the intervention group than controls	B (a)
Self-rated depression and vigorousness	Kamioka et al. (59)	Once a week balneotherapy (41.5°C) with lifestyle edu- cation and physical exercise	56	6 months	Less self-rated depression and increas- ing vigorousness in the intervention group than controls	B (a)

 Table 5
 Balneotherapy in psychological conditions

Rest is an important element in a spa resort treatment. The physical and emotional distance from the stress and concerns of daily life and worries over livelihood could be beneficial in reducing daily stress. Stress reduction has been shown to reduce disease activity by its effect on the soluble interleukin-2 receptor level in RA (60).

General contraindications for balneotherapy should also be considered. These contraindications include severe psychiatric conditions, acute alcoholic states, epilepsy, cardiac dysrhythmias, inadequate balance, severe varicose veins, open wounds and hypersensitivity to mineral baths (61). Furthermore, hyperthermic immersion might produce potentially dangerous cardiovascular effects, such as ectopic beats and excessive tachycardia (62).

Conclusion

Although there is a tendency to publish statistically significant findings and there is a lack of access to many nonsignificant studies, existing data support the beneficial effects of spa therapy on several diseases and conditions. Besides any

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specific effect of any natural waters, a proper regimen of treatment and rehabilitation with physiotherapy and hydrotherapy in an atmosphere free from stress will undoubtedly help many patients. In interpretation of the findings, the methodological aspects for each study should be taken into consideration. Can the tap water be adequately disguised? Can immersion be organized in a double-blind fashion? Can the wider aspects of climatotherapy be controlled? In addition, some other important questions should also be answered: Could the positive findings have resulted from patient expectations? How long do the positive effects persist? Were the benefits worth the costs? It is not clearly recognizable whether the improvements seen were due to the direct effect of spas or from the resort stay. Since the temperature and chemical compositions of each spa is a characteristic of that spa resort, extrapolating the results to other spa resorts should be done cautiously.

Most of the studies reported in this review were conducted on a small number of subjects. With such a smaller number, clinically relevant effects might be missed. Performing randomized studies using a larger study population is recommended so that statistical significance comes close to clinical relevance.

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