REVIEW

Evidence-Based Opinions from Multidisciplinary Experts on Use of Naturopathic Herbal Remedies in Pain Management

Ali Mobasheri 10 1-4, Aliyyaa Spring-Charles⁵, Francesco Carlo Gamaleri 10 6, Joyce McSwan⁷, Manohar Garg⁸, Vidhu Sood Sethi⁹

¹Research Unit of Health Sciences and Technology, Physics and Technology, University of Oulu, Oulu, Finland; ²Department of Regenerative Medicine, State Research Institute Centre for Innovative Medicine, Vilnius, Lithuania; ³Department of Joint Surgery, Sun Yat-sen University, Guangzhou, People's Republic of China; ⁴World Health Organization Collaborating Center for Public Health Aspects of Musculoskeletal Health and Aging, Université de Liège, Liège, Belgium; ⁵Hope Spring Clinic, London, UK; ⁶Pharmacy San Rocco, Cornaredo (Milan), Italy; ⁷PainWISE Pty Ltd, Gold Coast, Queensland, Australia; ⁸Nutraceuticals Research Program, University of Newcastle, Callaghan, Newcastle, NSW, Australia; ⁹Medical Affairs, Haleon (Formerly GlaxoSmithKline Consumer Healthcare) Pte. Ltd., Singapore

Correspondence: Vidhu Sood Sethi, Haleon (Formerly GlaxoSmithKline Consumer Healthcare) Pte. Ltd., Singapore, Tel +65 96447203, Email vidhu.x.sethi@haleon.com

Background: Pharmacological approaches to acute and chronic pain management, including non-steroidal anti-inflammatory drugs (NSAIDs) and opioids, are respectively associated with adverse reactions (such as gastrointestinal, cardiovascular, and renal effects) that might limit their use in patients with comorbidities and controversy related to inappropriate use. Naturopathic remedies might offer patients alternative and integrative treatments with minimal side effects.

Objective: To explore the regional variation in the acceptance and use of naturopathic remedies in pain management.

Methods: Two expert panel discussions were held by GlaxoSmithKline Consumer Healthcare (now Haleon Pte. Ltd.) over 9 and 12 hours in 2020 and 2021, respectively, and attended by multidisciplinary experts in naturopathy, Ayurvedic medicine, community pharmacy, physiotherapy, clinical pharmacy, Western medicine, academics, and naturopathic pain relief. Experts shared and discussed their experiences of naturopathic treatments and relevant clinical evidence related to different types of pain (including joint and muscle pain, migraine, sleeplessness due to pain, and general pain) and examined barriers to providing support to patients.

Results: Experts agreed on the potential for curcumin (2020, 71.4% [5/7]; 2021, 91.7% [11/12]) and fish oil (2020, 100% [7/7]) for management of osteoarthritic joint pain although these are not uniformly recommended in osteoarthritis treatment guidelines. In treatment of migraines, coenzyme Q10 and magnesium were favored by experts (2021, 90.9% [10/11] and 63.6% [7/11], respectively). **Conclusion:** The need was emphasized for more and higher quality clinical studies to support naturopathic remedies, which might not be reflected in the latest treatment guidelines. The expert panel also highlighted missed opportunities for physicians and pharmacists to recommend effective naturopathic treatments.

Keywords: herbal remedies, naturopathy, osteoarthritis, migraine, expert panel discussion

Introduction

Pain has been recognized as important in the primary care of patients with chronic conditions. Similar to management of acute pain, treatments including acetaminophen and non-steroidal anti-inflammatory drugs (NSAIDs), and their combinations, are amongst the most popular medicines currently used for reducing various types of pain. Acetaminophen is believed to operate by activation of descending serotonergic pathways and NSAIDs act via inhibition of the pro-inflammatory enzyme cyclooxygenase (COX), which is involved in the generation of inflammation and biochemical recognition of pain. However, use of NSAIDs is associated with complications of the upper gastrointestinal, cardiovascular (CV), and renal systems. Opioids have become increasingly used for management of chronic pain; however, these medications are associated with controversies, owing to concerns regarding

inappropriate use and related mortality and morbidity. Opioids account for more than 70% of 0.5 global million deaths attributable to drug use, with deaths by overdose accounting for >30% of those cases. 10 Increased misuse of pain medications has prompted the World Health Organization (WHO) to revise its guidance on management of pain, 11 motivating the need for alternative pain management solutions that avoid problems associated with NSAIDs and opioids.

Natural products and dietary supplements (hereafter referred to as naturopathic remedies/products) have relatively low side effects, which may favor their use in patients with comorbidities and represent potential alternative pain management solutions for inflammatory pain and hyperalgesia. 12 Many patients with both acute and chronic pain are turning to non-pharmacological therapies for additional relief and research activity, and interest in these alternative remedies have increased. 13 Among existing recommendations on naturopathic remedies, some treatment guidelines now incorporate recommendations on use of naturopathic remedies in conditions, including osteoarthritis, 14 back pain, 15 neuropathic pain, 16 and dysmenorrhea. 17 For example, the European League Against Rheumatism (EULAR) 18 and European Society for Clinical and Economic Aspects of Osteoporosis and Osteoarthritis (ESCEO)¹⁹ have recognized the roles of naturopathic remedies. The Australian guidelines for rheumatoid arthritis also reflect evidence of the anti-inflammatory effects of curcumin.²⁰ Magnesium has been recommended for treating migraines in some countries.^{21,22} However, there is a lack of a global consensus on the use of naturopathic remedies in clinical guidelines that address pain management.

Issues preventing wider recommendations of naturopathic remedies include limited and low-quality evidence of their effectiveness and/or slower onset of action compared with pharmacological treatments, as well as a lack of education and insufficient collaboration among healthcare professionals.²³ Standardization of naturopathic medicines also poses challenges, based on differences in preparation and plant species, which may alter the chemical profile dependent upon geographical area, seasonal variations, method of harvest, extraction procedures, and/or storage conditions.²⁴ Falling under the umbrella of complementary and alternative remedies, naturopathic remedies are not uniformly regulated across different regions, and in some cases not regulated at all.^{25–27}

Although generally recognized as being associated with relatively few adverse drug reactions, there are risks and contraindications associated with naturopathic remedies that warrant caution.²⁸ Lower risks associated with naturopathic products depend on appropriate levels of education, and safe standards of practice and mechanisms should be enacted to ensure these standards are met. Naturopathic products are often provided as over-the-counter (OTC) medicines and are accessible for self-selection, and there is the potential for incorrect dosage and interactions with other drugs²⁹ due to a lack of clinical guidance. These features place a particular burden on pharmacists, who may be more easily accessed by the public than appointment-based naturopaths or general practitioners (GPs).³⁰ Furthermore, pharmacists are now under greater pressure to deliver frontline guidance to patients, owing to the global COVID-19 pandemic, which has led to restricted access to GPs by patients.³¹ Patients are likely to use pharmacists as their as first point of contact, which represents an opportunity for re-education of these healthcare providers to play a more active role in advising on naturopathic remedies.

The expanding use of complementary remedies necessitates healthcare education that addresses misconceptions of the public and healthcare providers, potentially involving appropriate government oversight.³² Any such measures should be supported by further clinical and basic research into complementary and alternative medicine³³ and studies of their costeffectiveness and how to best integrate these options within existing public health frameworks.

In this review, we examine variation in ways of treating pain and the use of naturopathic products across different geographic regions and cultures. The panel evaluated current clinical evidence to identify evidence gaps in the use of naturopathic products for pain management. The panel also examined barriers to the uptake of naturopathic remedies, the path of patients using naturopathic treatments and potential pitfalls, as well as the scope for multidisciplinary integration of holistic approaches with existing healthcare systems. The format of these meetings provided a unique forum, bringing together experts from a range of healthcare ecosystems to discuss and share opinions on topics related to herbal remedies for consideration by industry stakeholders.

Journal of Pain Research 2024:17 600

Dovepress Mobasheri et al

Methods

Two online events were held over 2 days each in 2020 (9 hours total) and 2021 (12 hours total) and attended by a multidisciplinary panel of expert practitioners in naturopathy, Ayurvedic medicine, community pharmacy, pain medicine, clinical pharmacy, joint pain and osteoarthritis (OA), and naturopathic pain relief. In 2020, 7 experts joined from India, Malaysia, Singapore, and Australia. In 2021, 11 experts were in attendance from Saudi Arabia, United Arab Emirates, Finland, Italy, United Kingdom, India, Malaysia, Singapore, and Australia. The panel was selected based on a desk search of the literature, experts' affiliations in national and international organizations relevant to naturopathy, and institutional/personal websites for individuals active in the above areas of expertise and regions, who expressed an interest in participating in discussions on these topics. The duration and timing of the meetings were determined through consultation with the experts. Prior to the expert panel discussions, practicing attendees were surveyed on their current naturopathic treatment recommendations (Supplementary Tables 1 and 2). During the discussions, experts delivered presentations on naturopathic treatments for specific and relevant clinical evidence related to different types of pain, including joint and muscle pain, migraine, sleeplessness due to pain, and general pain (meeting agendas are shown in Supplementary Tables 3-6). The topics were discussed further by the panel, and consensus polling was conducted.

Results and Discussion

Joint and Muscle Pain

Osteoarthritis is the leading cause of disability and joint pain worldwide.³⁴ As a complex and multifactorial degenerative joint disease, osteoarthritis is primarily a disease of the articular cartilage that causes local inflammation and mechanical changes. Nutrition and dietary supplementation have been shown to have meaningful effects on the disease 35,36

Curcumin is a popular treatment option for joint pain/muscle pain due to its strong antioxidant and anti-inflammatory effects.³⁷ These effects are well supported by clinical evidence including a 2016 systematic review and meta-analysis, which, although based on a small number of RCTs, concluded that turmeric extract is effective in managing pain from osteoarthritis. 38 Boswellia has been investigated in a few recent clinical trials that have reported conflicting results. 39,40 A recent meta-analysis 41 found that boswellia and its extracts may be an effective and safe treatment option for patients with osteoarthritis. Use of fish oils in osteoarthritis has been widely studied; however, a meta-analysis has noted the lowto-moderate quality of current supporting evidence.⁴²

Experts who attended the 2021 panel discussion reported that 5–30% of their patients were seeking non-medicated pain relief from osteoarthritic joint pain. A higher percentage of patients reportedly sought non-medicated solutions for joint and muscle pain in the Middle East (50-90%) compared with Singapore/Malaysia (10-20%) and Australia (5-10%).

In 2020 experts responded to a pre-meeting survey question on their top 3 naturopathic solutions used to treat joint pain. Curcumin and fish oil were both strongly recommended, with each receiving 7 votes, followed by magnesium and boswellia (2 votes each). Glucosamine, palmitoylethanolamide with nattokinase, and ginger root also received 1 vote each. During the meeting, the experts voted for curcumin (5 of 7 experts) and ginger root (1 of 7 experts) as their preferred naturopathic remedies for joint pain.

In a consensus discussion on the use of curcumin complex, the 2021 panel agreed (11 of 12 experts with 1 abstention) that the complex offered benefits in treating joint pain. Common factors that motivate the selection of treatment options were safety and efficacy of the natural compounds, as supported by naturopaths, pharmacists, and researchers (9 votes); personal experience/observation (5 votes); and access to high-quality products (2 votes).

In a roundtable discussion, the experts considered the importance of communicating the effects of different ingredients for different types of pain, the role of oral versus topical use, and the holistic effects of therapies in terms of the gut microbiome. Integration of naturopaths and pharmacists, who often advise patients, was also discussed as a means of promoting the role of naturopathic remedies in the healthcare system.

https://doi.org/10.2147/JPR.S432090 Journal of Pain Research 2024:17 60 I

Sleeplessness Due to Pain

Population-based studies have found that up to a third of people may report sleep disorders; ⁴³ however, sleep complaints are more prevalent in people with chronic pain disorders (50–88%).⁴⁴ Furthermore, a bidirectional relationship between sleep and pain exists, whereby pain reduces sleep quality⁴⁵ and poor sleep quality amplifies pain perception.⁴⁶ Sleeplessness due to pain can increase a patient's risk for depression and anxiety, reduce functional ability. 45 and increase systemic/widespread pain.⁴⁷

Among naturopathic treatments for sleeplessness, the use of Valeriana officinalis is widely supported in the literature. 48-50 In 2021, experts from most regions, except Europe, concurred that V. officinalis has a role in managing sleeplessness due to pain, although there were some differing practices. Australia experts also recommended magnesium and St John's Wort/Ziziphus (2 votes each). Experts from Europe (ie, UK, Finland, and Italy) preferred magnesium, boswellia, and melatonin (1 vote each). Singapore/Malaysia experts preferred ashwagandha (Withania somnifera) (2 votes) and passiflora (1 vote). Saudi Arabia/UAE experts preferred passiflora and melatonin (1 vote each).

A consensus discussion focused on the use of ashwagandha to promote sleep in patients with associated chronic pain. While experts reported positive feedback from patients using ashwagandha for this purpose, the experts were divided on the use of ashwagandha for this specific novel combination of indications (6 of 12 in favor) but concluded that ashwagandha might be useful in managing sleeplessness with pain.

Use of melatonin and its combination with other nutrients are supported for use in sleeplessness; however, long-term use of OTC products can have adverse effects on regulation of endogenous melatonin production and release by the pineal glands.⁵¹ Experts across different specialties concurred that the most convincing evidence for treating sleeplessness due to pain was in melatonin. In Saudi Arabia and Italy, fewer patients sought naturopathic solutions (0% and 3%, respectively) for sleeplessness compared with 50% of patients in Australia, which might reflect the status of melatonin as a prescription drug in the former regions. Consumers from these countries may be more likely to seek medicated solutions for sleeplessness (eg., prescription melatonin and potentially other pharmacological treatments) rather than natural options.

Headache/Migraine

Acute and chronic pain from migraine and headache may derive from a variety of diseases⁵² and can negatively affect many important aspects of life.⁵³ Naturopathic therapies are likely to have an important role in managing migraine, owing to their fewer associated adverse effects.⁵⁴

For feverfew (Tanacetum parthenium), there is clinical evidence to support its use in treating headache; however, a need for further and higher quality studies is noted.⁵⁵ Feverfew is indicated for migraines in some treatment guidelines. 56-58

Magnesium has a more established role in treating migraine. Although a 2019 systematic review found insufficient evidence for magnesium in combination with coenzyme Q10 (vitamin B) for migraine prophylaxis, ⁵⁹ a later 2020 systematic review of reviews⁶⁰ found support for its effects in reducing the intensity/frequency of migraines. Magnesium is currently recommended in several guidelines for treatment of migraines. 21,22,56-58,61-63

There is also evidence to support the effectiveness of coenzyme Q10 in reducing the frequency and duration of migraines, 62 and a number of guidelines include this recommendation. 21,58,61,62,64,65

Experts were asked for their top 3 naturopathic therapies for migraine. Coenzyme Q10 (vitamin B) received the greatest number of recommendations (10 votes) followed by magnesium (3 votes). Feverfew also received 1 mention, in addition to Ginkgo biloba, calcium, curcumin/turmeric, and probiotics. During the meeting, 7 experts voted magnesium as the top treatment option for migraine. Pre-meeting survey results from the 2021 meeting indicated a similar preference for magnesium, with some regional differences: Australian and European experts preferred feverfew (4 votes), magnesium (3 votes) and willow bark (2 votes). Singapore/Malaysia experts preferred magnesium (1 vote), boswellia (1 vote), and palmitoylethanolamide (1 vote). Saudi Arabia/UAE experts preferred feverfew (1 vote) and menthol (1 vote).

Following a consensus discussion, experts were divided on the use of feverfew, with 6 of 12 agreeing that it might be effective for migraine prophylaxis (preventive care) or for episodic relief of migraine.

Journal of Pain Research 2024:17 602

Doverress Mobasheri et al

General Pain

General pain includes both chronic and acute pain resulting from conditions such as back pain, 66 menstrual pain, 67 abdominal pain, 68 and musculoskeletal pain. 69

Meta-analyses have indicated the effectiveness of magnesium for various types of pain, 70 and specifically in dysmenorrhea. ⁷¹ Although magnesium is recommended for different types of pain (including aforementioned migraines) few treatment guidelines currently recommend use of magnesium for dysmenorrhea.⁷²

Although widely recommended for general pain, boswellia has not been extensively studied outside of the indication for osteoarthritis. Potential anti-oxidative, anti-inflammatory, anti-amyloidogenic, and anti-apoptotic properties have been recognized; however, there is a need for further clinical evidence to support use of boswellia for different pain types.⁷³

Curcumin/turmeric received the most recommendations for treatment of general pain from experts who indicated their top 3 preferred solutions prior to the 2020 expert panel discussion (3 votes). Omega-3 fatty acids and resveratrol also received 2 mentions each, and magnesium, willow bark, Jamaican dogwood received 1 vote each. During the meeting, experts voted for curcumin/turmeric (4 of 7 experts) and omega-3 fatty acids (3 of 7 experts) as their preferred naturopathic solutions for general pain.

In 2021, the top 3 naturopathic ingredients recommended to treat general pain included magnesium (7 votes), curcumin (6 votes), and boswellia (6 votes). In Australia, curcumin (2 votes), magnesium (2 votes) and boswellia (2 votes) were preferred. European (UK/Finland/Italy) experts preferred curcumin (2 votes), willow bark extract (2 votes), and boswellia (1 vote). In Singapore/Malaysia, magnesium (1 vote) and boswellia (1 vote) were preferred. Usage in Saudi Arabia/UAE included Arnica montana extract (1 vote), Vitis vinifera leaf extract (1 vote) and menthol (1 vote).

The panel noted that, particularly for acute pain, naturopathic products may be administered at higher doses or used as an adjunct to pharmaceutical pain killers to reduce doses required by the patient. Naturopathic products may also allow for gradual tapering from strong painkillers once a patient's pain has improved.

Barriers to the Use of Naturopathic Products

Many consumers seek rapid solutions for their pain, particularly in the case of acute pain, which might result in the underlying causes for their symptoms going undiagnosed. This treatment-seeking behavior may be at odds with naturopathic approaches to pain management, which consider patient symptoms from a holistic context. For acute pain, naturopathic approaches may be effective in use alongside conventional pharmaceutical therapies; however, naturopathic remedies together with holistic lifestyle changes could offer preventative and greater longer-term benefits to patients and reduce reliance on pharmaceutical pain killers. There remains a lack of high-quality evidence to demonstrate the broader efficacy of naturopathic consultations. However, the 2021 panel unanimously agreed (11/11) that natural pain medications have a role in both symptomatic and preventative care.

The panels identified gaps in the training of healthcare professionals. Those trained in Western medicine may have misconceptions about naturopathic treatment and prevent or discourage patients from using naturopathic treatments. Hence, referrals to naturopaths from healthcare practitioners (HCPs) were noted to be rare across all regions. HCPs and researchers often consult treatment guidelines, many of which exclude naturopathic solutions because of poor quality evidence. Hence, there is also a need to both improve the quality and the volume of studies supporting naturopathic remedies, including non-inferiority trials.

As an additional barrier, many naturopathic products are more expensive than regular pharmaceutical products. Hence, there is a need to improve affordability and accessibility of naturopathic remedies to realize their full value in treatment as primary medicines.

Patients often seek naturopathic options owing to mistrust of medical doctors and concern about side effects of standard medications. Pre-conceptions about naturopathic therapies, unrealistic expectations for immediate effects and the differences between effectiveness towards acute and chronic pain may lead to low trust in the products. In selfadministering naturopathic remedies, patients might not be aware of the time required for response and fail to complete their full treatment course. Use of naturopathic products is not without risk, and there is a need to provide more information to consumers about contraindications and drug interactions. There may also be perceptions among general

https://doi.org/10.2147/JPR.S432090 Journal of Pain Research 2024:17 603 consumers that naturopathic remedies are poor quality, high in cost, and/or ineffective compared with pharmaceuticals. Therefore, to establish patient trust, production methods and quality assurance approaches to naturopathic remedies should mirror requirements of the pharmaceutical industry. Standardization of active ingredients should help to build greater trust in naturopathic products among both healthcare professionals and patients.

Importance of Pharmacy Education and Labelling

The patient journey to consult with a naturopathic practitioner typically starts with a desire to avoid starting a course of pharmaceutical treatment or to substitute or decrease reliance on their current medication. Amid growing interest in alternative therapies among patients and the absence of guidance from GPs and pharmacists, patients are likely to seek alternative sources of information.

Social influencers in this space have raised awareness of alternative therapies, but provision of unreliable information could potentially lower trust in naturopathic remedies. When regional agencies seek to revise their guidelines through the consensus approach, it may be useful to engage with a trusted spokesperson from the naturopathic community to raise awareness of naturopathic treatments among experts. There may also be benefits from developing national and international networks of naturopathic practitioners to align on delivery of treatment and consistent patient messaging.³²

A large proportion of patients who experience pain are likely to first approach a pharmacist for an OTC solution. Pharmacy visits represent an important opportunity for providing patients with information about effectiveness, safety, dosage, and duration of naturopathic therapies. Product labelling is also crucial, and packaging information on indications treated and specific doses for symptomatic and preventive care might benefit patients. The use of online resources accessed through QR codes on packages could provide more information on the product and about naturopathy. Regardless of the source and delivery method, patient information about naturopathic products should avoid overpromising treatment benefits and emphasize the holistic context in which these products best serve patients. 74,75

There is a clear long-term need for inclusion of content on naturopathic medicine in the curriculum for medical students and pharmacists. This might be achieved by further research into course design in collaboration with instructors, collecting evaluations and reflections from participating students. Incorporating naturopathic medicine within existing healthcare systems may improve the rate of appropriate referrals from GPs to naturopaths. This may in turn help to address misconceptions among the public and build trust in naturopathic remedies and an understanding of their role alongside pharmaceuticals.

Conversations with pharmacists and consumers should also be promoted. Surveys to evaluate pharmacist knowledge and engagement with naturopathic remedies might offer useful insights. This might enable the development of factsheets to be shared by pharmacists and could be a useful tool for delivering dependable and accessible product information. Pharmacy assistants could be educated through bite-sized engaging materials to increase their confidence in recommending naturopathic medicine in the OTC setting and ensure that doses are tailored to the patient.

Strengths and Limitations

There are some limitations to insights obtained from these expert panel discussions. Notably, no formal consensus or qualitative research methodology was applied to either meeting; furthermore, expert views are not supported by a comprehensive review and quality assessment of the clinical literature. Nevertheless, the independent insights from this large panel of international experts serves to highlight regional differences and similarities in provision of naturopathic remedies for pain management. There was a potential risk of selection bias in deciding the panel. Panel members may also have felt compelled to vote in line with the majority during the meetings. However, both pre-meeting voting and voting following discussions in the meetings demonstrated strong alignment among experts on the effectiveness of specific ingredients and on qualitative steps for integrating naturopathic practices with conventional healthcare systems. Although not all the same experts were able to attend both meetings, there was overall agreement of several key experts between the meetings. Furthermore, although the panels agreed that naturopathic remedies may be safer than conventional pharmaceuticals, there remains a risk of adverse reactions by individuals to certain ingredients and a need for caution in combining natural remedies with other therapies. Further exploration of appropriate dosages and contraindications of the ingredients are warranted.

https://doi.org/10.2147/JPR.S432090 Journal of Pain Research 2024:17 **Dove**press Mobasheri et al

Conclusions

Notable regional variation was observed in the use of naturopathic products by patients with pain across this multidisciplinary and geographically diverse panel. For example, views differed on the benefits of ashwagandha for sleeplessness due to pain. However, the panels agreed on the benefits of certain therapies, including curcumin and fish oil for osteoarthritic joint pain and coenzyme Q10 and magnesium for migraines. Here, we try to integrate differing views from a range of practitioners representing these regions and develop integration of naturopathic products with conventional pain management solutions. However, further studies should seek to identify appropriate dosages and consider potential safety issues associated with specific naturopathic products.

Consumers and patients may not be aware of the effects of the ingredients in naturopathic medicine. Hence, there is a need to ensure that naturopathic products for pain are marketed based on the indication they treat rather than the ingredients. Labelling on products should also clearly indicate dosage and the treatment period to ensure that patients have realistic expectations and maintain compliance with naturopathic products. Any important safety considerations in using the medicine should also be clearly highlighted on labels. Labelling could be augmented with digital media, through QR codes, to provide more information on products.

A key barrier to recommending naturopathic remedies in treatment guidelines is the lack of quality evidence. Thus, new clinical trials with larger populations are required, to generate high-quality evidence related to effectiveness and safety of naturopathic products. There are inconsistencies across regional recommending bodies regarding naturopathic treatments, which are based on existing evidence of effectiveness. Involvement of the naturopathic community in medical expert panel discussions and consensus meetings could help to ensure that naturopathic products are fairly evaluated and recommended where appropriate.

Pharmacy assistants in most countries may be the first point of contact within a pharmacy and there is a need to add content on naturopathic products to the curriculum of pharmacists to improve awareness among the profession and to provide consumers with trustworthy information on the effectiveness of the products and their safety. Provision of information brochures and factsheets with products that may also be shared with consumers could help to improve acceptance of the products among experienced pharmacists and pharmacy assistants and support staff.

Ethics Approval

Our study is a narrative review/commentary based on a consensus discussion of professionals who gave written consent to participate in the meetings and signed standard agreements, which assigned Haleon full rights to the content generated, including consent for scientific publication. There was no intervention, and no human subjects were involved in the study. On this basis, the study is not subject to the Declaration of Helsinki and exempt from institutional review board ethical review.

Acknowledgments

The authors thank Andrew Jackson of MIMS, Singapore, for providing medical writing support, which was funded by Haleon (formerly GlaxoSmithKline Consumer Healthcare) in accordance with Good Publication Practice (GPP3) guidelines (http://www.ismpp.org/gpp3).

Disclosure

VS is a current employee of Haleon (formerly GSK Consumer Healthcare); AM has consulted for and received honoraria from Haleon. AS-C, FCG, JM, and MG have all received honoraria from Haleon. The authors report no other conflicts of interest in this work.

References

- 1. Smith BH, Fors EA, Korwisi B, et al. IASP taskforce for the classification of chronic pain, the IASP classification of chronic pain for ICD-11: applicability in primary care. Pain. 2019;160:83-87. doi:10.1097/j.pain.0000000000001360
- 2. Allegri M, Clark MR, De Andres J, Jensen TS. Acute and chronic pain: where we are and where we have to go. Minerva Anestesiol. 2012;78:222-235.

Journal of Pain Research 2024:17 https://doi.org/10.2147/JPR.S432090 605 Mobasheri et al Dovepress

3. Bindu S, Mazumder S, Bandyopadhyay U. Non-steroidal anti-inflammatory drugs (NSAIDs) and organ damage: a current perspective. *Biochem Pharmacol*. 2020;180:114147. doi:10.1016/j.bcp.2020.114147

- 4. Anderson BJ. Paracetamol (Acetaminophen): mechanisms of action. Paediatr Anaesth. 2008;18:915-921. doi:10.1111/j.1460-9592.2008.02764.x
- Ong CK, Lirk P, Tan CH, Seymour RA. An evidence-based update on nonsteroidal anti-inflammatory drugs. Clin Med Res. 2007;5:19–34. doi:10.3121/cmr.2007.698
- Ofman JJ, MacLean CH, Straus WL, et al. A meta analysis of severe upper gastrointestinal complications of nonsteroidal antiinflammatory drugs. J Rheumatol. 2002;29:804–812.
- Grosser T, Ricciotti E, FitzGerald GA. The cardiovascular pharmacology of nonsteroidal anti-inflammatory drugs. Trends Pharmacol Sci. 2017;38:733–748. doi:10.1016/j.tips.2017.05.008
- 8. Kalso E, Edwards JE, Moore AR, McQuay HJ. Opioids in chronic non-cancer pain: systematic review of efficacy and safety. *Pain*. 2004;112:372–380. doi:10.1016/j.pain.2004.09.019
- Els C, Jackson TD, Kunyk D, et al. Adverse events associated with medium- and long-term use of opioids for chronic non-cancer pain: an overview of Cochrane reviews. Cochrane Database Syst Rev. 2017;10:CD012509.
- 10. World Health Organization. Opioid overdose; 2021. Available from: https://www.who.int/news-room/fact-sheets/detail/opioid-overdose. Accessed March 17, 2023.
- 11. World Health Organization. Model list of essential medicines; 2021. Available from: https://www.who.int/publications/i/item/WHO-MHP-HPS-EML-2021.02. Accessed March 17, 2023.
- 12. Chen L, Michalsen A. Management of chronic pain using complementary and integrative medicine. BMJ. 2017;357:j1284.
- 13. Wang C, Meng Q. Global research trends of herbal medicine for pain in three decades (1990–2019): a bibliometric analysis. *J Pain Res.* 2021;14:1611–1626. doi:10.2147/JPR.S311311
- 14. Arden NK, Perry TA, Bannuru RR, et al. Non-surgical management of knee osteoarthritis: comparison of ESCEO and OARSI 2019 guidelines. *Nat Rev Rheumatol.* 2021;17:59–66. doi:10.1038/s41584-020-00523-9
- 15. Oliveira CB, Maher CG, Pinto RZ, et al. Clinical practice guidelines for the management of non-specific low back pain in primary care: an updated overview. Eur Spine J. 2018;27:2791–2803. doi:10.1007/s00586-018-5673-2
- Boyd A, Bleakley C, Hurley DA, et al. Herbal medicinal products or preparations for neuropathic pain. Cochrane Database Syst Rev. 2019;4: CD010528. doi:10.1002/14651858.CD010528.pub4
- Wattier JM. [Conventional analgesics and non-pharmacological multidisciplinary therapeutic treatment in endometriosis: CNGOF-HAS endometriosis guidelines]. Gynecol Obstet Fertil Senol. 2018;46:248–255. French. doi:10.1016/j.gofs.2018.02.002
- 18. Kloppenburg M, Kroon FP, Blanco FJ, et al. 2018 update of the EULAR recommendations for the management of hand osteoarthritis. *Ann Rheum Dis.* 2019;78:16–24. doi:10.1136/annrheumdis-2018-213826
- 19. Bruyere O, Honvo G, Veronese N, et al. An updated algorithm recommendation for the management of knee osteoarthritis from the European Society for Clinical and Economic Aspects of Osteoporosis, Osteoarthritis and Musculoskeletal Diseases (ESCEO). Semin Arthritis Rheum. 2019;49:337–350. doi:10.1016/j.semarthrit.2019.04.008
- 20. Therapeutic guidelines, managing symptoms in patients with rheumatoid arthritis; 2022. https://tgldcdp.tg.org.au/viewTopic?topicfile=rheumatoid-arthritis&guidelineName=Rheumatology#toc d1e438. Accessed March 17, 2023.
- 21. Pringsheim T, Davenport WJ, Mackie G, et al. Canadian Headache Society guideline for migraine prophylaxis. *Can J Neurol Sci.* 2012;39(2 Suppl 2):S1–S59.
- 22. Antonaci F, Dumitrache C, De Cillis I, Allena M. A review of current European treatment guidelines for migraine. *J Headache Pain*. 2010;11:13–19. doi:10.1007/s10194-009-0179-2
- 23. Sugito R, Son D. Obstacles to the use of complementary and alternative medicine by primary care physicians: preliminary study. *Tradit Kampo Med.* 2019;6:173–177. doi:10.1002/tkm2.1225
- Ruiz GG, Nelson EO, Kozin AF, Turner TC, Waters RF, Langland JO. A lack of bioactive predictability for marker compounds commonly used for herbal medicine standardization. *PLoS One*. 2016;11:e0159857. doi:10.1371/journal.pone.0159857
- 25. Wiesener S, Salamonsen A, Fønnebø V. Which risk understandings can be derived from the current disharmonized regulation of complementary and alternative medicine in Europe? *BMC Complement Altern Med.* 2018;18:11. doi:10.1186/s12906-017-2073-9
- 26. Boon H. Regulation of complementary/alternative medicine: a Canadian perspective. Complement Ther Med. 2002;10:14–19. doi:10.1054/ctim.2002.0499
- 27. Fischer F, Lewith G, Witt CM, et al. A research roadmap for complementary and alternative medicine what we need to know by 2020. Forsch Komplementmed. 2014;21A:e1–16. doi:10.1159/000360744
- 28. Xiong F, Guan YS. Cautiously using natural medicine to treat liver problems. World J Gastroenterol. 2017;23:3388–3395. doi:10.3748/wjg.v23. i19.3388
- 29. Eickhoff C, Hämmerlein A, Griese N, Schulz M. Nature and frequency of drug-related problems in self-medication (over-the-counter drugs) in daily community pharmacy practice in Germany. *Pharmacoepidemiol Drug Saf.* 2012;21:254–260. doi:10.1002/pds.2241
- 30. Jebara T, Cunningham S, MacLure K, Awaisu A, Pallivalapila A, Stewart D. Stakeholders' views and experiences of pharmacist prescribing: a systematic review. *Br J Clin Pharmacol*. 2018;84:1883–1905. doi:10.1111/bcp.13624
- 31. Basheti IA, Nassar R, Barakat M, et al. Pharmacists' readiness to deal with the coronavirus pandemic: assessing awareness and perception of roles. Res Social Adm Pharm. 2021;17:514–522. doi:10.1016/j.sapharm.2020.04.020
- 32. Gray AC, Steel A, Adams J. A critical integrative review of complementary medicine education research: key issues and empirical gaps. *BMC Complement Altern Med.* 2019;19:73. doi:10.1186/s12906-019-2466-z
- 33. Institute of Medicine (US) Committe on the Use of Complementary and Alternative Medicine by the American Public. Complementary and Alternative Medicine in the United States. US, Washington DC: National Academies Press; 2005. 5. State of Emerging Evidence on CAM
- 34. Safiri S, Kolahi AA, Smith E, et al. Global, regional and national burden of osteoarthritis 1990–2017: a systematic analysis of the Global Burden of Disease Study 2017. *Ann Rheum Dis.* 2020;79:819–828. doi:10.1136/annrheumdis-2019-216515
- 35. Liu X, Machado GC, Eyles JP, Ravi V, Hunter DJ. Dietary supplements for treating osteoarthritis: a systematic review and meta-analysis. *Br J Sports Med.* 2018;52:167–175. doi:10.1136/bjsports-2016-097333

606 https://doi.org/10.2147/JPR.5432090 Journal of Pain Research 2024:17

Dovepress Mobasheri et al

36. Thomas S, Browne H, Mobasheri A, Rayman MP. What is the evidence for a role for diet and nutrition in osteoarthritis? *Rheumatology (Oxford)*. 2018;57(suppl–4):iv61–iv74. doi:10.1093/rheumatology/key011

- 37. Kotha RR, Luthria DL. Curcumin: biological, pharmaceutical, nutraceutical, and analytical aspects. Molecules. 2019;24:2930
- 38. Daily JW, Yang M, Park S. Efficacy of turmeric extracts and curcumin for alleviating the symptoms of joint arthritis: a systematic review and meta-analysis of randomized clinical trials. *J Med Food*. 2016;19:717–729. doi:10.1089/jmf.2016.3705
- Żęgota Z, Goździk J, Głogowska-Szeląg J. Prospective, multicenter evaluation of a polyherbal supplement alongside standard-of-care treatment for mild knee osteoarthritis. Adv Orthop. 2021;2021:5589597. doi:10.1155/2021/5589597
- 40. Liu X, Robbins S, Eyles J, et al. Efficacy and safety of a supplement combination on hand pain among people with symptomatic hand osteoarthritis an internet-based, randomised clinical trial the RADIANT study. *Osteoarthritis Cartilage*. 2021;29:667–677. doi:10.1016/j.joca.2021.01.011
- 41. Yu G, Xiang W, Zhang T, Zeng L, Yang K, Li J. Effectiveness of boswellia and boswellia extract for osteoarthritis patients: a systematic review and meta-analysis. *BMC Complement Med Ther.* 2020;20:225. doi:10.1186/s12906-020-02985-6
- 42. Senftleber NK, Nielsen SM, Andersen JR, et al. Marine oil supplements for arthritis pain: a systematic review and meta-analysis of randomized trials. *Nutrients*. 2017;9:42. doi:10.3390/nu9010042
- 43. Ancoli-Israel S, Roth T. Characteristics of insomnia in the United States: results of the 1991 National Sleep Foundation Survey. I. Sleep. 1999;22 (22 Suppl 2):S347–S353.
- 44. Finan PH, Goodin BR, Smith MT. The association of sleep and pain: an update and a path forward. *J Pain*. 2013;14:1539–1552. doi:10.1016/j. jpain.2013.08.007
- 45. Baker S, McBeth J, Chew-Graham CA, Wilkie R. Musculoskeletal pain and co-morbid insomnia in adults; a population study of the prevalence and impact on restricted social participation. *BMC Fam Pract*. 2017;18:17. doi:10.1186/s12875-017-0593-5
- 46. Staffe AT, Bech MW, Clemmensen SLK, Nielsen HT, Larsen DB, Petersen KK. Total sleep deprivation increases pain sensitivity, impairs conditioned pain modulation and facilitates temporal summation of pain in healthy participants. PLoS One. 2019;14:e0225849. doi:10.1371/journal.pone.0225849
- 47. Wiklund T, Gerdle B, Linton SJ, Dragioti E, Larson B. Insomnia is a risk factor for spreading of chronic pain: a Swedish longitudinal population study (SwePain). Eur J Pain. 2020;24:1348–1356. doi:10.1002/ejp.1582
- 48. Azizi H, Shojaii A, Hashem-Dabaghian F, et al. Effects of *Valeriana officinalis* (Valerian) on tension-type headache: a randomized, placebo-controlled, double-blind clinical trial. *Avicenna J Phytomed*. 2020;10:297–304.
- 49. Bent S, Padula A, Moore D, Patterson M, Mehling W. Valerian for sleep: a systematic review and meta-analysis. *Am J Med.* 2006;119:1005–1012. doi:10.1016/j.amjmed.2006.02.026
- 50. Shinjyo N, Waddell G, Green J. Valerian root in treating sleep problems and associated disorders-a systematic review and meta-analysis. *J Evid Based Integr Med*. 2020;25:2515690X20967323. doi:10.1177/2515690X20967323
- 51. Bravaccio C, Terrone G, Rizzo R, et al. Use of nutritional supplements based on melatonin, tryptophan and vitamin B6 (Melamil Tripto[®]) in children with primary chronic headache, with or without sleep disorders: a pilot study. *Minerva Pediatr.* 2020;72:30–36.
- 52. Leonardi M, Raggi A. A narrative review on the burden of migraine: when the burden is the impact on people's life. *J Headache Pain*. 2019;20:41. doi:10.1186/s10194-019-0993-0
- 53. Buse DC, Fanning KM, Reed ML, et al. Life with migraine: effects on relationships, career, and finances from the Chronic Migraine Epidemiology and Outcomes (CaMEO) study. *Headache*. 2019;59:1286–1299. doi:10.1111/head.13613
- 54. Kaur K, Hernandez V, Al Hajaj SW, et al. The efficacy of herbal supplements and nutraceuticals for prevention of migraine: can they help? *Cureus*. 2021;13:e14868.
- 55. Wider B, Pittler MH, Ernst E. Feverfew for preventing migraine. Cochrane Database Syst Rev. 2015;4:CD002286. doi:10.1002/14651858. CD002286.pub3
- 56. Araki N, Takeshima T, Igarashi H, Shimizu T. Clinical practice guideline for chronic headache 2013. In: The Chronic Headache Clinical Practice Guideline Development Committee of Japan. Vol. 7. Neurology and Clinical Neuroscience; 2019:231–259.
- 57. Sarchielli P, Granella F, Prudenzano MP, et al. Italian guidelines for primary headaches: 2012 revised version. *J Headache Pain*. 2021;13(Suppl 2): S31–S70.
- 58. Evers S, Afra J, Frese A, et al. EFNS guideline on the drug treatment of migraine-revised report of an EFNS task force. Eur J Neurol. 2009;16:968–981. doi:10.1111/j.1468-1331.2009.02748.x
- 59. Okoli GN, Rabbani R, Kashani HH, et al. Vitamins and minerals for migraine prophylaxis: a systematic review and meta-analysis. *Can J Neurol Sci.* 2019;46:224–233.
- 60. Veronese N, Demurtas J, Pesolillo G, et al. Magnesium and health outcomes: an umbrella review of systematic reviews and meta-analyses of observational and intervention studies. *Eur J Nutr.* 2020;59:263–272. doi:10.1007/s00394-019-01905-w
- Institute of Health Economics, Alberta, Canada. Primary care management of headache in adults; 2016. Available from: https://actt.albertadoctors.org/CPGs/Lists/CPGDocumentList/Primary-Care-Management-of-Headache-in-Adults.pdf. Accessed March 17, 2023.
- 62. Becker WJ, Findlay T, Moga C, Scott NA, Harstall C, Taenzer P. Guideline for primary care management of headache in adults. *Can Fam Physician*. 2015;61:670–679.
- 63. Diener HC, Holle-Lee D, Nägel S, Dresler T, Gaul C, Göbel H. Treatment of migraine attacks and prevention of migraine: guidelines by the German Migraine and Headache Society and the German Society of Neurology. Clin Transl Neurosci. 2019;3:1–40.
- 64. Sazali S, Badrin S, Norhayati MN, Idris NS. Coenzyme Q10 supplementation for prophylaxis in adult patients with migraine-a meta-analysis. *BMJ Open.* 2021;11:e039358.
- 65. British Association for the study of headache, national headache management system for adults; 2018. https://bash.org.uk/downloads/guide lines2019/02_BASHNationalHeadache_Management_SystemforAdults_2019_guideline_versi1.pdf. Accessed March 17, 2023.
- 66. Urits I, Burshtein A, Sharma M, et al. Low back pain, a comprehensive review: pathophysiology, diagnosis, and treatment. *Curr Pain Headache Rep.* 2019;23:23. doi:10.1007/s11916-019-0757-1
- 67. Ferries-Rowe E, Corey E, Archer JS. Primary dysmenorrhea: diagnosis and therapy. Obstet Gynecol. 2020;136:1047–1058. doi:10.1097/AOG.000000000000000066
- Natesan S, Lee J, Volkamer H, Thoureen T. Evidence-based medicine approach to abdominal pain. Emerg Med Clin North Am. 2016;34:165–190. doi:10.1016/j.emc.2015.12.008

Journal of Pain Research 2024:17 https://doi.org/10.2147/JPR.5432090 **607**

Mobasheri et al **Dove**press

- 69. Hawker GA. The assessment of musculoskeletal pain. Clin Exp Rheumatol. 2017;35(Suppl 107):8-12.
- 70. Shin HJ, Na HS, Do SH. Magnesium and pain. Nutrients. 2020;12:2184. doi:10.3390/nu12082184
- 71. Pattanittum P, Kunyanone N, Brown J, et al. Dietary supplements for dysmenorrhoea. Cochrane Database Syst Rev. 2016;3:CD002124. doi:10.1002/14651858.CD002124.pub2
- 72. Colegio Mexicano de Especialistas en Ginecología y Obstetricia. Diagnóstico Y Tratamiento de la Dismenorrea Primaria en Adolescentes; 2008. Available from: http://comego.org.mx/GPC TextoCompleto/8-Diagnostico%20y%20tratamiento%20de%20la%20dismenorrea%20primaria%20en %20adolescentes.pdf. Accessed March 17, 2023.
- 73. Rajabian A, Sadeghnia H, Fanoudi S, Hosseini A. Genus Boswellia as a new candidate for neurodegenerative disorders. Iran J Basic Med Sci. 2020;23:277-286. doi:10.22038/IJBMS.2020.35288.8419
- 74. McSwan J, Gudin J, Song XJ, et al. Self-healing: a concept for musculoskeletal body pain management scientific evidence and mode of action. J Pain Res. 2021;14:2943-2958. doi:10.2147/JPR.S321037
- 75. Mobasheri A. "Self-healing": a novel and integrated multimodal concept for the management of musculoskeletal pain. J Pain Res. 2022;15:3479–3482. doi:10.2147/JPR.S386508

Journal of Pain Research

Dovepress

Publish your work in this journal

The Journal of Pain Research is an international, peer reviewed, open access, online journal that welcomes laboratory and clinical findings in the fields of pain research and the prevention and management of pain. Original research, reviews, symposium reports, hypothesis formation and commentaries are all considered for publication. The manuscript management system is completely online and includes a very quick and fair peer-review system, which is all easy to use. Visit http://www.dovepress.com/testimonials.php to read real quotes from published authors.

Submit your manuscript here: https://www.dovepress.com/journal-of-pain-research-journal

