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The history of light therapy in hospital physiotherapy and medicine with emphasis on Australia: Evolution into novel areas of practice

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ABSTRACT

Objective: The objective of this narrative review was to investigate the history of light therapy in hospital settings, with reference to physiotherapy and particularly in an Australian context.

Types of articles and search method: a review of available literature was conducted on PubMed, Medline and Google Scholar using keywords light therapy, photobiomodulation, physiotherapy, low-level laser, heliotherapy. Physiotherapy textbooks from Sydney University Library were searched. Historical records were accessed from the San Hospital library. Interviews were conducted with the San Hospital Chief Librarian and a retired former Head Physiotherapist from Royal Prince Alfred Hospital.

Summary: Historically, light treatment has been used in both medical and physiotherapy practice. From its roots in ancient Egypt, India, and Greece, through to medieval times, the modern renaissance in 'light as therapy ' was begun by Florence Nightingale who, in the 1850s, advocated the use of clean air and an abundance of sunlight to restore health. Modern light therapy (phototherapy) had a marked uptake in use in medicine in Scandinavia, America, and Australia from 1903, following the pioneering work of Niels Finsen in the late 19th century, which culminated in Dr Finsen receiving the Nobel Prize for Medicine for the treatment of tuberculosis scarring with ultraviolet (UV) light, and treatment of smallpox scarring with red light. Treatment with light, especially UVB light, has been widely applied by physiotherapists in hospitals for dermatological conditions since the 1950s, particularly in Australia, Scandinavia, USA, England and Canada. In parallel, light treatment in hospitals for hyperbilirubinemia was used for neonatal jaundice. Since the 1980s light was also used in the medical specialties of ophthalmology, dermatology, and cardiology. In more recent years in physiotherapy, light was mostly used as an adjunct to the management of orthopedic/rheumatological conditions. Since the 1990s, there has been global use of light, in the form of photobiomodulation for the management of lymphedema, including in supportive cancer care. Photobiomodulation in the form of low-level laser has been used by physiotherapists and pain doctors since the 1990s in the management of chronic pain. The use of light as therapy is exemplified by its use in the San Hospital in Sydney, where light therapy was introduced in 1903 (after Dr. John Harvey Kellogg visited Niels Finsen in Denmark) and is practiced by nurses, physiotherapists and doctors until the present day. The use of light has expanded into new and exciting practices including supportive cancer care, and treatment of depression, oral mucositis, retinopathy of prematurity, and cardiac surgery complications. Light is also being used in the treatment of neurological diseases, such as Parkinson's disease, traumatic brain injury, and multiple sclerosis. The innovative uses of light in physiotherapy treatment would not be possible without the previous experience of successful application of light treatment.

Conclusion: Light therapy has had a long tradition in medicine and physiotherapy. Although it has fallen somewhat out of favour over the past decades, there has been a renewed interest using modern techniques in recent times. There has been continuous use of light as a therapy in hospitals in Australia, most particularly the San Hospital in Sydney where it has been in use for almost 120 years.

Introduction

This paper will explore the relationship between the history of the use of light in physiotherapy and its potential as a novel treatment for a number of intractable diseases. A number of examples of traditional light therapy are presented along with the evolution of light therapy into photobiomodulation and experimental treatments for cardiovascular disease, neurological diseases such as multiple sclerosis and Parkinson's disease,

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and supportive cancer care, particularly the treatment of lymphedema and oral mucositis.

The history of light treatment in physiotherapy is closely paralleled by the history of light treatment in medicine. Light has been used in medicine in many guises, importantly as a diagnostic tool such as in microscopy, spectrometry, quantitative PCR and ELISA techniques. It is increasingly used for sophisticated optogenetic research in diagnosis and treatment. Light is used as a trigger for photodynamic therapy (PDT), where photosensitive dyes are activated to release reactive oxygen species in order to kill infective microorganisms or cancer cells (Hamblin, 2016). Light can also be used as a direct therapy, including thermal and non-thermal laser applications, light emitting diodes (LED), bright light and intravenous and transdermal UVB applications.

Light has been used as a treatment for medical conditions since ancient times, with its origins as "heliotherapy" (natural sunlight) for the treatment of skin diseases. Some 3,500 years ago, the ancient Egyptians used a weed grown on the banks of the Nile (*Amul majus*) followed by sun exposure, to treat vitiligo (Hönigsmann, 2013). This treatment is analogous to the modern photochemotherapy (PUVA) first applied in 1974, with extension of its use to various other conditions (Hönigsmann, 2013). The field of photochemotherapy has recently expanded into the treatment of multiple sclerosis (MS) using UVB (Hart, Finlay-Jones, and Gorman, 2009; Hart et al., 2018) after the work on the immunosuppressive effects of UVB (McGonigle et al., 2017).

Modern phototherapy has its origins in the latter part of the 19th century (Figure 1), with the observation, in 1877, that sunlight was beneficial in the treatment of anthrax infections (Downes and Blunt, 1877) and then in 1890, when it was employed for the treatment of rickets (Palm, 1890).

The combination of this early research in medicine, as well as the influence of Florence Nightingale with her view that ... "It is the unqualified result of all my experience with the sick that, second only to their need of fresh air, is their need of light; that, after a close room, what hurts them most is a dark room and that it is not only light but direct sunlight they want." (Nightingale, 1859) was the beginning of light therapy, both as heliotherapy (natural sunlight) and phototherapy (artificial light). Arnold Rikli also advocated the healing potential of light and heliotherapy, with the establishment of heliohydrotherapy centers in the 1850s in Bled (Slovenia). The awarding of the 1903 Nobel Prize in Medicine to Niels Finsen, for his work on the use of light in treatment of disease, paved the way for the continued and expanded use of light therapy in the 20th century and beyond, including the use of light in physiotherapy.

History of light treatment in modern physiotherapy

Nonthermal light treatment in the disciplines of medicine and physiotherapy began with the treatment of dermatological conditions (Figure 1). In the late 1890s, Niels Finsen began treating lupus vulgaris with blue and UV light and smallpox scarring with red light, using lamps and filters. His results were published in the British Medical Journal in 1895 (Finsen, 1895) and in The Lancet in 1903 (Finsen, 1903). The Editorial in the BMJ (Valdemar, 1899) included "the uptake shows the treatment spread quickly around the world". One example of this uptake was by Dr. John Harvey Kellogg, who traveled to Denmark to work with Finsen and introduced light boxes to his clinics in Battle Creek, Michigan in 1891 (Schwarz, 2006). His brother, Dr. Merritt Kellogg, introduced light therapy into the newly formed Sydney Adventist (San) Hospital, also in 1903 (Michael Rigby, San Librarian, pers. com., Sydney Adventist Hospital archive marg\wpold\history). A full account of the treatment rationale and protocol was published by Dr. John Harvey Kellogg in 1910 (Kellogg, 1910) and as a result of his early medical practice, light therapy has been in use in the Seventh Day Adventist health institutions in various forms for over a hundred years.

Light treatment was originally performed by nurses and massage therapists. In 1906 these therapists joined the Australian Massage Association, which, in Australia, evolved into "physiotherapy" during World War I and the term was formalized in 1939 (McMeeken, 2008; Sellentin, 2015). Nurses at the San Hospital were taught massage and hydrotherapy from 1906 and in the 1940s were offered membership of the Society of Massage and Electrotherapy and given automatic registration into the society in 1946 (Michael Rigby, San Librarian, pers. com., Sydney Adventist Hospital archive marg\wpold \history) (Figures 1 and 2).

There was a period after the extensive use of light in the 1930s, when phototherapy, which includes UV and farinfrared (thermal) wavelengths, resulted in injury with home-use applications. This came about due to untrained persons, who without adequate considerations of dose and timing, caused burns. This controversy surrounding light therapy was well documented (Woloshyn, 2017). After Medical Board consultation in Britain, UVB light therapy was restricted to hospital use to be administered by health professionals including physiotherapists and nurses.

Mrs Pam Wilster, the former Head Physiotherapist of the Royal Prince Alfred (RPA) Hospital (Sydney) for over three decades, reported administering phototherapy to children in The London Hospital, for such diverse



New horizons for treatment of intractable diseases

Figure 1. Timeline of the history of the use of light as therapy in physiotherapy and medicine. (1) Nightingale (1859); (2)https://flashbak.com/light-therapy-for-naked-children-delicate-adults-sick-pigs-and-quacks-photos-1900-1950-41389/(3 Finsen (1895); (4) Finsen (1903); (4) Hönigsmann (2013); (5) pers. com Pam Wilster, former Head Physiotherapist, Royal Prince Alfred Hospital, Sydney; (6) Mester, Juhász, Varga, and Karika (1968); (7) Piller and Thelander (1996); (8) Ralf Uwe et al. (1994); (9) Horvath et al. (2001); (10) https://spinoff.nasa.gov/database/spinoffDetail.php?this=/spinoff/msfc/MSFC-SO-161; (11) Schiffer et al. (2009); (12) Chow, Johnson, Lopes-Martins, and Bjordal (2009); (13) Nussbaum, Gabel, Bjordal, and Houghton (2011); (14) Stone, Johnstone, and Mitrofanis (2013); (15) Hallett (2015); (16) Liebert, Bicknell, and Adams (2014); (17) Natoli et al. (2013); (18) Hamblin (2016); (19) https://www.nobelprize.org/prizes/lists/all-nobel-laureates-in-physiology-or-medicine/(2018); (20) Hamilton et al. (2018); (21) Kiat (2017); (22) Hart et al. (2018); (23) Liebert et al. (2019); (24) Liebert and Bicknell (2019).

diseases as rickets, dermatitis, and autoimmune disorders such as asthma (Pam Wilster, pers. com.). These free treatments were available in London hospitals, including the Royal London Hospital, from 1899 (https://flashbak. com/light-therapy-for-naked-children-delicate-adults-

sick-pigs-and-quacks-photos-1900-1950-41389/), and were given by nursing staff and masseurs, who went on to become registered physiotherapists (Woloshyn, 2017). Pam Wilster instigated this treatment at RPA in 1958 for the prevention of rickets and asthma. Physiotherapists were applying UVB treatment at RPA hospital in Sydney continuously until 1995 when the procedures were taken over by nurses and other health professionals. Light treatment was also used in Canada from the 19th century until the 1940s for the treatment of rickets (UVB), infantile paralysis and nervous diseases (electric light cabinets) (Connor and Pope, 1999). The treatments, which included light baths that were understood to have analgesic as well as anti-bacterial properties, was ongoing in many hospitals throughout the world at this time but is less widely used in current physiotherapy practice (Nuhu, Mohammed, and Muhammad, 2014). Newly graduated physiotherapists have therefore been denied the opportunity to experience the use of phototherapy and to be a part of an integrated approach to treatment for the medically compromised patient.

While UVB treatment continued to be used in the 1960s, there was also an increase in light treatment using red and infrared wavelengths. Light therapy increased in clinical applications in the 1970s for its application in the treatment of both dermatitis and hyperbilirubinemia (Brown, Kim, Wu, and Bryla, 1985). Intravenous therapy using UVB has also been used for many decades (Wu, Hu, and Hamblin, 2016).

Light treatment expanded after an incidental discovery in 1967 by Dr. Endre Mester who began to perform the first experiments using laser light to accelerate wound healing (as well as re-growing hair) in animals and patients (Mester, Juhász, Varga, and Karika, 1968). This treatment effect was biphasic in nature, such that increased dosage beyond an energy threshold resulted in no further increase in healing effect. Mester also documented a systemic effect of laser, where the stimulatory effect of the treatment could be seen in areas adjacent and remote from the point of application of the laser. Laser technology was championed by Dr. Tiina Karu, an Estonian scientist, who published more than 190 papers in the field of low-level laser therapy, particularly practiced by doctors and physiotherapists in Russia; later to expand to Israel and Iran (Moskvin, 2017; Oron et al., 2001). The effectiveness of the therapy was demonstrated after the Chernobyl nuclear accident in 1986, when radiation wounds were successfully treated with

laser by the American military (Ralf Uwe, 2002; Ralf Uwe et al., 1994). This contrasted with the non-treated, non-healing radiation ulcers that were the rule after the nuclear blasts in Japan during World War II, that did not resolve in the lifetime of the victims. The evidence that red and near-infrared light was effective in healing wounds continued to grow after this time and set the stage for their contemporary use in radiation injuries and radiodermatitis resulting from cancer therapies (Robijns, Lodewijckx, and Mebis, 2019). The early work with laser light therapy has led to the use of what is now known as photobiomodulation therapy (Khan and Arany, 2015), which is the use of non-thermal light to stimulate beneficial cellular processes for therapeutic effect. This is a rapidly expanding field with physiotherapy applications reaching into treatment of musculoskeletal pain (Chow, Johnson, Lopes-Martins, and Bjordal, 2009), osteoarthritis pain (Stausholm et al., 2019), neurological conditions (Hamblin, 2016) and cancer-related lymphedema and oral mucositis.

In parallel with the early use of light as a therapy and its development over the next century (Figure 1), advances in the uses of light was also occurring in other areas of medicine. This included the use of blue light to treat hyperbilirubinemia in new-borns in the 1970s, and the wider use of light in the 1970s for diseases such as skin lesions and vitiligo, due in part to the development of lasers in the early 1960s, and the later development of cheaper LEDs.

Paradoxically, the expanded use of light in medicine (i.e. high-powered surgical lasers in dermatology, orthopedics, oncology, and ophthalmology) has been accompanied by decreased use in physiotherapy and a consequent diminishing of the knowledge and practice of light therapy for wounds and dermatological conditions by newly qualified physiotherapists. For example, in Nigeria, the lack of knowledge was reported to be accompanied by a lack of access to UV equipment, both being a barrier to effective treatment (Nuhu, Mohammed, and Muhammad, 2014). It is somewhat ironic that the continued use of UVB by physiotherapists since the 1950s for the treatment of dermatological conditions in the hospital setting (Fitzpatrick, 1988) (Sydney Adventist Hospital archive marg\wpold\history), has allowed the expedited use of UVB for the treatment of acute immune diseases, such as multiple sclerosis (MS) in medical research (Prue Hart, Research Fellow, University of Westerrn Australia, pers. com.).

The use of UVB in the treatment of MS is thought to regulate the vitamin D axis in a manner analogous to rickets (Hart et al., 2018). Other forms of light are also being investigated for the effect on cases of MS (Lyons, clinical trial https://clinicaltrials.gov/ct2/show/NCT03691766). Knowledge of the historical use of

light and its effects is important in this context and its inclusion in teaching curricula would be beneficial.

Light therapy applications expanded in the eighties into the field of dentistry, for the treatment of oral wounds (including "dry sockets"); and for treating the cancer treatment side-effect of oral mucositis; a debilitating condition of the mucous membranes, where patients may need to be tube fed due to the pain and inflammation. PBM is used by many hospitals around the world now use photobiomodulation to successfully treat oral mucositis. It is now considered standard treatment for oral mucositis caused by chemotherapy or radiotherapy, under the Multinational Association for Supportive Care in Cancer (MASCC) guidelines (Zadik et al., 2019) and the National Institute for Health and Care Excellence (NICE) guidelines in Britain. https://www.nice.org.uk/guidance/ ipg615. In Australia, this treatment has been introduced into The Royal Children's Hospital in Melbourne by Dr. Kerrod Hallett, after a successful pilot study (Hallett, 2015) and has been extended to the San Hospital in 2021. The use of photobiomodulation therapy and previously low-level laser therapy for lymphedema was first proposed following an incidental finding in 1989, where lowlevel laser treatment of arthritis by physiotherapists coincidentally showed an improvement in long-standing lymphedema. This led to extensive research at Flinders University in both animal and clinical models (Figure 1) to show the effective dosage and applications for this treatment (Piller and Thelander, 1996). A laser device, specifically designed to treat lymphedema has approval by both the Therapeutic Goods Association (Australia) and the Food and Drug Administration (USA).

Light, including sunlight, is important in all aspects of critical illness and has been shown to be an important consideration and to have direct modulating effects on illness (Castro, Angus, and Rosengart, 2011). For example, lighting in hospitals is incorporated into the architectural design of newly built hospitals to assist in medical care and optimal oncological management. Oncologists also recommend exposure to red light in the morning as being therapeutic (Holick, 2008). Differences in prenatal light exposure has also been linked with the risk of: Parkinson's disease (Gardener et al., 2010); MS (Hart et al., 2018); autism (Cieslinska et al., 2017); and schizophrenia (Kendell and Adams, 2002). These observations suggest potentially rich research areas of preventative medicine for otherwise intractable diseases. Bright light therapy is also used for seasonal affective disorder (Wirz-Justice et al., 1996) and Parkinson's disease (Willis, Boda, and Freelance, 2018). In summary Florence Nightingale's principle is still being applied in health practice and appears to be assuming an increased clinical importance.

Recent advances in the application of photobiomodulation have been pioneered in Australia, America, Canada, Norway, and Brazil. The advances follow from the understanding of light modulation on human tissues, physiology, and disease obtained by NASA (Whelan, 2002) in the 1970s when wounds that would not heal in microgravity, were shown to heal under the influence of light in the experimental hydroponic units of the space shuttle. These findings led to NASA spinoff technology and the development of the WARP 10 LED device (https://ntrs.nasa.gov/archive/nasa/casi.ntrs.nasa. gov/20090002502.pdf). The understanding of the mechanisms of action and the effect of various wavelengths, energy, modulation frequency, and other parameters of light treatment has improved the outcomes of light therapy and has led to increasingly successful clinical therapy trials. These trials have included the use of photobiomodulation to treat: chronic neck pain (Chow, Johnson, Lopes-Martins, and Bjordal, 2009); lymphedema (Smoot et al., 2015); arthritis (Alfredo et al., 2018); and muscle performance (Ferraresi, Huang, and Hamblin, 2016).

In the 1980s, an Australian manipulative physiotherapist Louis Gifford (1987) pioneered the study of circadian rhythms. He examined the importance of circadian rhythms on joint stiffness, by awakening volunteers every hour over a 24-hour period to measure changes in joint flexibility. He noted changes in tissue elasticity and collagen. This work was a precursor to the principles of chronobiology and has been an important background principle to photobiomodulation and how light more generally impacts health. The importance of chronobiology in health and disease was recognized with a Nobel Prize in 2017 to the researchers Jeffery Hall, Michael Rosbash and Michael Young "for their discoveries of molecular mechanisms controlling the circadian rhythm" (https://www.nobelprize.org/prizes/lists/allnobel-laureates-in-physiology-or-medicine/).

Chronobiology appears to have an important clinical role for the timing of surgery, hospital lighting, prediction of the risk of such diverse diseases as Parkinson's disease, MS and some cancers, and the progression of diseases such as Alzheimer's disease. These recent advances in chronobiology and circadian rhythm research and their effects on seasonal affective disorder, miscarriage in shift workers, jetlag and so on, have led to treatment of these disorders using light (Lushington et al., 2002).

Research in chronobiology and photobiomodulation suggests the possibility of incorporating phototherapy in the treatment of a range of diseases and the possibility for physiotherapists, as well as physicians, dentists and other health professionals, to further embrace phototherapy, including photobiomodulation therapy (red and near-infrared laser and LED) into their existing treatment options.

Since the late 1990s, photobiomodulation therapy has been taught in the Physiotherapy curricula in the University of Queensland and Griffith University in Australia. This follows from Professor Liisa Laakso and her long association with these universities subsequent to her PhD studies into mechanisms of low-level laser therapy (Laakso, Cramond, Richardson, and Galligan, 1994). Photobiomodulation is also in the scope of practice of physiotherapy, as well as the curricula at many universities worldwide, including in Brazil, Canada, Russia, Iran, and Europe. The breadth of photobiomodulation research has expanded from treatment of muscle, bone, and nerve disorders, to include applications in muscle strength and athletic performance, supportive cancer care, and neurological disorders.

At the San Hospital, light therapy has been used in patient care for over 100 years (Figure 2). Physiotherapists at this hospital were thus in a position to embrace laser technology for the treatment of lymphedema when this became available. The routine use of light therapy over many years also allowed for research into light therapy and the appointment of a Director of Photomolecular Research (a physiotherapist) at the Australasian Research Institute at the hospital in 2016, following on from a Phototherapy Biological Research Laboratory established in 2011. This appointment was facilitated by the fact that the CEO of the hospital at that time had been a nurse manager, who was aware of the tradition of the therapy. A Photobiomodulation Clinic has also been established with a physiotherapy as well as a medical focus for the treatment of chronic pain, arthritis, cardiac surgery pre-conditioning, and sports injuries. Clinical trials are being conducted in this hospital (and other centers including the Mater Hospital in Brisbane) to assess the effect of photobiomodulation treatment on the symptoms of Parkinson's disease (ANZCTR registered trial ACTRN12618000038291). Current research and clinical trials include assessing the effects of photobiomodulation on red blood cell stability, cardiac surgery outcomes, and pre-conditioning against postoperative cognitive decline (Liebert, Chow, Bicknell, and Varigos, 2016; Liebert et al., 2017).

Expanded use of light for cardiovascular medicine

In medicine, the importance of light on individual health is apparent. Patients in brightly lit cardiac casualty wards have improved survival compared to those in dark wards (Beauchemin and Hays, 1998). The timing of heart attack events is influenced by the onset of daylight saving time changes, in that there are increases in heart attacks when clocks are put forward (one less hour of sleep) and decreases when clocks are put back (Sandhu, Seth, and Gurm, 2014).

Laser light treatment had also been used clinically for inoperable intractable heart disease. The transmyocardial reperfusion technique, carried out in the 1990s and early 2000s, used a focussed high-powered laser beam to create microscopic holes in cardiac muscle to increase blood perfusion. The procedure reduced pain in patients for 5 or more years (Horvath et al., 2001). Academic surgeons who performed the procedures (Prof Clifford Hughes, Head of Infection Control, Macquarie University, pers. com.) were of the opinion that the observed improvement in symptoms could also be attributed to a (low-level) laser effect myocardial tissues adjacent to the focussed laser; the so-called "bystander effect" (Najafi, Fardid, Hadadi, and Fardid, 2014) that may have been part of the mechanism that reduced angina and improved the quality of life outcomes in a majority of patients. This therapeutic technique was discontinued after several years, due to the high inherent operative risks, which were deemed to outweigh the symptomatic benefits (Briones, Lacalle, Marin-Leon, and Rueda, 2015).

777Following their initial work on intravascular lowpower red laser light as an adjunct to coronary stent implantation in porcine models (De Scheerder et al., 2001b), De Scheerder et al. (2001a) tested the feasibility and safety of intravascular low-power laser irradiation in an unblinded, single-arm clinical study of 68 patients undergoing percutaneous coronary intervention by stenting. The study showed a 6-month overall restenosis rate of 14.7%, from zero to 41.6% depending on the stented vessel size. In a similar study (Derkacz et al., 2013) with 52 patients receiving low-level laser treatment (49 in the control group), intra-coronary laser therapy showed a 6 month restenosis rate of 15% for the laser treatment group compared with 32.4% in the control group. The laser group also showed reduced pro-inflammatory markers (interleukin-1β, interleukin-6), reduced tumor growth factor- β 1, reduced fibroblast growth factor-2 and increased anti-inflammatory markers (interleukin-10) and nitric oxide compared to controls. Additionally, the cardiac event rate defined as death, Q and non-Q wave myocardial infarction, coronary artery by-pass grafting, and repeat revascularization of the target vessel/lesion was 7.7% for the lasertreated group at both 6- and 12-month follow-ups, compared to 14.3% and 18.5% in the control group for these follow-ups (Derkacz et al., 2010). There were also marked differences in late coronary lumen loss, late lumen loss index, minimal lumen diameter and lower

History of Light Treatment: Future directions



Figure 2. History of light treatment and future directions. (1) pers. com. Michael Rigby, Museum curator, San Hospital, Sydney; (2) Brown, Kim, Wu, and Bryla (1985); (3) Piller and Thelander (1996); (4) Chow, Johnson, Lopes-Martins, and Bjordal (2009); (5) Hamblin, Nelson, and Strahan (2018). (6) Liebert et al. (2017); (7) Zadik et al. (2019). (8) Vassão et al. (2020). (9) Liebert and Bicknell (2019).

6-7month restenosis rate (15.0% vs 32.4%) in favor of the laser-treated group. Infrared laser is currently being used in a cardiac stent trial (ANZCTR trial registration ACTRN12618001312235) in Tasmania by Dr.Michael Fox, a medical doctor and registered physiotherapist (pers. com.) to determine the effect on rates of restenosis, mortality, and morbidity. To date, 190 participants have been treated. The trial was formulated on the basis of case studies conducted at the San Hospital in Sydney (Kiat, 2017)

The original transcranial reperfusion using high-powered laser and the concomitant bystander effect on myocardial symptoms has been the inspiration for the use of red and infrared low-level laser as a preconditioning and/or postconditioning treatment in cardiac surgery (Liebert et al., 2017). The technique has been used by physiotherapists and doctors in Russia and Iran for heart patients for more than a decade (Kazemi Khoo et al., 2014; Moskvin, 2017). A singlearm clinical study of ischemic heart disease found that lowlevel laser therapy improved angina symptoms, systolic blood pressure, and exercise capacity, while reducing frequency of ischemia (Zyciński et al., 2007). A controlled trial in patients undergoing elective coronary bypass surgery conducted by Kazemi Khoo et al. (2014) showed improved cardiac and other biomarkers (i.e. creatinine kinase activity, creatine kinase myocardial band, lactate dehydrogenase, neutrophil and lymphocyte counts) post-operatively for those who received perioperative low-level laser therapy compared to controls, which may indicate cardiac cellular damage and inflammation. Photobiomodulation therapy also has the potential to accelerate the healing and reduce the trauma and nonunion of the sternotomy wound that is a feature of coronary artery bypass graft surgery (Helmy et al., 2019).

The best-suited areas of cardiovascular medicine for low-level laser therapy, optimized treatment algorithm including energy spectrum, dose, timing, anatomic application sites, delivery modes (e.g. percutaneous and intravascular) remain pressing questions to be addressed by robust, prospective, double-blind, randomized, placebo-controlled and multicenter clinical testing and validation studies. Nonetheless, based on the prevailing extensive experimental research output and several clinical studies pertinent to the field, the clinical future of low-level laser therapy in cardiovascular medicine seems likely to expand.

Expanded use of light for neurodegenerative disease

It has long been recognized that light is important in the management of neurological disorders, including Alzheimer's disease, for circadian rhythm stabilization, and improving sleep and behavior (Ancoli-Israel et al., 2003; Satlin et al., 1992; Van Someren, Kessler, Mirmiran, and Swaab, 1997). Experimental evidence from animal models has extended understanding of the systemic effects of photobiomodulation including the effects of photobiomodulation on areas of the body remote from the application of light, known as the bystander and abscopal effects. The helmet experiment in 2012 (Stone, Johnstone, and Mitrofanis, 2013), demonstrated that photobiomodulation had the potential to protect against neurological damage in the brain even when light was used on a site remote to the head. This finding has paved the way for recent innovations in the use of light as photobiomodulation and bright light therapy for both the transcranial and remote site treatment of neurological diseases such as: Parkinson's disease (Ganeshan et al., 2019; Johnstone, Mitrofanis, and Stone, 2015; Liebert and Bicknell, 2019; Mitrofanis, 2017); Alzheimer's disease (Blivet, Meunier, Roman, and Touchon, 2018); traumatic brain injury (Hamblin, 2016); and MS (Hart et al., 2018). It is now understood that many of these diseases have a strong gut/microbiome/brain connection and, at least in animal models, that changing the gut microbiome is a potential treatment option (Lubomski et al., 2020). The observation that photobiomodulation can change the gut microbiome (Bicknell, Liebert, Johnstone, and Kiat, 2018) suggests both a mechanism for the effect of remote light application and also a treatment option to expand current practice. These treatment protocols have followed on, not only from the understanding of the principles of light treatment, especially the effect of light on ion channels (Liebert, Bicknell, and Adams, 2014) and metabolomics more generally (Liebert, 2018), but also from observations during physiotherapy treatment of neurological disorders and observations that light is important in the management of Alzheimer's symptoms (Figueiro et al., 2014). The effect of circadian rhythms on the metabolome and the microbiome and other oscillatory processes in humans has paved the way for research into "photobiomics", the effect of light on the microbiome/metabolome interaction (Liebert et al., 2019), as well as the application of remote light application to the abdomen of mice to modulate a model of Parkinson's disease (Bicknell, Liebert, Johnstone, and Kiat, 2018; El Massri et al., 2018) and Alzheimer's disease (Blivet, Meunier, Roman, and Touchon, 2018). The effect of photobiomodulation on the microbiome of mice has now been replicated in clinical case studies (Liebert and Bicknell, 2019).

Future directions

Novel and expanding practice of light treatment is in the area of neurological disease such as: Parkinson's disease; multiple sclerosis; Alzheimer's disease; and motor neurone disease (Figure 2). Registered trials have been conducted in Sydney, Adelaide, and Brisbane, with preliminary results published in 2019 (Liebert and Bicknell, 2019; Vassão et al., 2020).

Depression as an accompanying symptom of chronic pain (Doan, Manders, and Wang, 2015), including the pain of Parkinson's disease (Martinez-Martin et al., 2017) and other central pains such as fibromyalgia and unresponsive cervicogenic headache (Liebert and Bicknell, 2017; Liebert et al., 2013) may be amenable to photobiomodulation therapy by laser and transcranial LED. Physiotherapists and pain doctors have been using photobiomodulation for chronic neck pain and accompanying headache since the 1990s with good effect (Chow, Johnson, Lopes-Martins, and Bjordal, 2009), including quality of life and mood improvements. A current trial for the reduction in symptoms of Parkinson's disease has shown an improvement in the non-motor symptoms of mood, sleep, and cognition (Liebert and Bicknell, 2019). A small trial using transcranial infrared light showed promising results for the reduction of symptoms of depression (Schiffer et al., 2009). There is also a potential role for managing other depressive symptoms that accompany chronic pain conditions using photobiomodulation, as a therapy delivered by physiotherapists (Askalsky and Iosifescu, 2019; Cassano et al., 2016; Hamblin, 2016). More opportunities will open up in the future and a strong historical basis makes this possible.

Conclusion

The use of light therapy in physiotherapy and medicine spans 120 years and has been continuous in some locations, especially in hospitals such as the San Hospital in Sydney. This continuity and intergenerational practice has allowed for the emergence of new and innovative treatments for familiar conditions (e.g. wounds, lymphedema, and pain) and also intractable diseases and conditions. These conditions include persistent ulcers caused by metabolic diseases such as diabetes, cognitive and musculoskeletal decline of aging, metabolic-related cardiovascular disease, cancer therapy sequalae such as oral mucositis, post-surgical complications such as post-operative cognitive dysfunction, and neurological diseases such as MS, motor neurone disease, Parkinson's and Alzheimer's diseases.

Inter-generational practice unique to the physiotherapy profession, including the employment of light therapy, from the 19th century use of light boxes to the 21st century use of lasers, is important to maintain a diverse professional foundation. The ability to understand and incorporate beneficial aspects of the history of physiotherapy is important for many reasons. It assists in maintaining the scope of practice and prevents the attrition and diminishment of therapies to other health practitioners. It also gives a historical as well as an evidence-based rationale for physiotherapy practice, the importance of which should never be underestimated. It is incumbent on all of us to encourage newly graduated physiotherapists to be aware of the past and simultaneously, to innovate for the future. With knowledge of what has been done before, young physiotherapists can have a positive attitude toward the therapeutic potential of light as an adjunct therapy. The opportunity for the physiotherapy profession to embrace past therapeutic practices and apply them in novel areas of health care, such as intractable diseases is very attractive.

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Declaration of Interest

The authors declare no conflict of interest.

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