



Mechatronic Wearable Exoskeletons for Bionic Bipedal Standing and Walking: A New Synthetic Approach

Gelu Onose^{1,2*}, Vladimir Cârdei³, Ștefan T. Crăciunoiu³, Valeriu Avramescu³, Ioan Opreș⁴, Mikhail A. Lebedev⁵ and Marian Vladimir Constantinescu⁶

¹ Department of Physical and Rehabilitation Medicine, University of Medicine and Pharmacy “Carol Davila”, Bucharest, Romania, ² Teaching Emergency Hospital “Bagdasar-Arseni”, Bucharest, Romania, ³ Research and Technological Design Institute for Machines Construction, Bucharest, Romania, ⁴ Miller School of Medicine, University of Miami, Miami, FL, USA, ⁵ Department of Neurobiology, Duke University, Durham, NC, USA, ⁶ Holistic Dental Medicine Institute – ROPOSTURO, Bucharest, Romania

OPEN ACCESS

Edited by:

Mircea Dragoman,
National Institute for Research and
Development in Microtechnologies,
Romania

Reviewed by:

Giuseppe D’Avenio,
Istituto Superiore di Sanità, Italy
Giovanni Stellan,
Telerobot Labs Srl, Italy

*Correspondence:

Gelu Onose
geluonose@gmail.com

Specialty section:

This article was submitted to
Neural Technology,
a section of the journal
Frontiers in Neuroscience

Received: 09 April 2016

Accepted: 08 July 2016

Published: 29 September 2016

Citation:

Onose G, Cârdei V, Crăciunoiu ȘT, Avramescu V, Opreș I, Lebedev MA and Constantinescu MV (2016) Mechatronic Wearable Exoskeletons for Bionic Bipedal Standing and Walking: A New Synthetic Approach. *Front. Neurosci.* 10:343. doi: 10.3389/fnins.2016.00343

During the last few years, interest has been growing to mechatronic and robotic technologies utilized in wearable powered exoskeletons that assist standing and walking. The available literature includes single-case reports, clinical studies conducted in small groups of subjects, and several recent systematic reviews. These publications have fulfilled promotional and marketing objectives but have not yet resulted in a fully optimized, practical wearable exoskeleton. Here we evaluate the progress and future directions in this field from a joint perspective of health professionals, manufacturers, and consumers. We describe the taxonomy of existing technologies and highlight the main improvements needed for the development and functional optimization of the practical exoskeletons.

Keywords: mechatronic, portable/wearable exoskeletons, bipedal standing, bionic walk assistance, models

HUMAN POSTURE AND WALKING, AND THEIR RESTORATION TO THE DISABLED

Vertical posture and bipedal gait are two hallmarks of human biomechanics and motor physiology, which emerged over the millions of years of evolution. While our body size and overall physical performance are not particularly impressive compared to other animal species, having our hands relieved from body support against gravity gives us a number of advantages, dexterity being one of the most important ones. In addition to motor physiology, virtually any human physiological function is adapted to the bipedal posture and walking, for example breathing, digestion, and excretion (Uebelhart et al., 2000)¹. Moreover, human bipedalism has had a great influence on our daily social interactions and the manmade habitat.

While the surrounding objects and structure of the buildings all fit the needs of healthy humans, the conditions are quite different for disabled, wheelchair bound people. No matter how many facilities would be made in order to overcome their numberless structural barriers, new hurdles are always to be expected. Wheelchair users will never be able to completely adapt to the habitat built for the people who walk bipedally and stand upright. By contrast, the cartoon of a person in the wheelchair is the worldwide used symbol for handicap.

¹http://www.sci-info-pages.com/other_issues.html#Osteoporosis

Although there is no cure for many lesions of the spinal cord and the brain (Talley Watts et al., 2015)—the major causes of paralysis—essential improvements can be made to the quality of life of handicapped people using devices that assist vertical stance and bipedal walking (Louie et al., 2015). These assistive technologies are based on mechatronic, robotic, and bionic exoskeletons (Chen et al., 2013). Such devices clearly represent one of the most important developments in rehabilitation of paralysis, even though they are currently not expected to deliver a truly spectacular, translational progress. Rather, they are considered as viable intermediary solutions, which bring us closer to the long-awaited recovery of posture and gait to the paralyzed patients. While much work is still needed to improve these external aids, they already have shown effective daily use capabilities.

ASSISTIVE TECHNOLOGIES: MANUFACTURER AND CONSUMER PERSPECTIVES

In the last decade and mostly in its second half, a series of exoskeletons have been designed, built, clinically tested in small groups of subjects, and reviewed in the literature^{2–9}. These previous publications have fulfilled the promotional and marketing objectives of the manufacturers, and several exoskeletons are currently commercially available. Yet, an optimal, fully functional assistive device has not yet emerged from this research and development^{10,11} (Chen et al., 2013; Arazpour et al., 2015b; Lajeunesse et al., 2015; Louie et al., 2015).

An optimal, practical exoskeleton should fulfill the following requirements:

- The system should be safe.
- It should be effectively wearable¹¹ in the common sense of the term, mainly regarding don and doff issues. Additionally, it should be psychologically acceptable in terms of self-esteem, miniaturization, and esthetics. Ideally, the exo-suit should be thin and wearable like clothes or underwear.
- The exoskeleton should be appropriate for long time performance, including the performance in community (Louie et al., 2015).

²http://www.cyberdyne.jp/company/download/20160212_financialresultsummary.pdf

³<http://www.digitaltrends.com/cool-tech/rewalk-6-0-exoskeleton/>

⁴<http://www.rexbionics.com/investors/>

⁵<http://www.engadget.com/2011/05/02/new-zealand-paralympian-buys-first-rex-bionics-exoskeleton-take/>

⁶<http://www.fastcompany.com/1822791/eksos-exoskeletons-let-paraplegics-walk-will-anyone-actually-wear-one>

⁷<http://world.honda.com/news/2015/p150721eng.html>

⁸<http://www.indego.com/indego/en/Product>

⁹<http://www.fastcodesign.com/3056049/a-budget-exoskeleton-allows-paraplegics-to-walk-for-the-price-of-a-car7>

¹⁰http://www.b-temia.com/category/press_releases/

¹¹<https://www.cadth.ca/sites/default/files/pdf/htis/july-2015/RA0784>

%20Powered%20Wearable%20Walking%20Assistive%20Devices%20Final.pdf

- It should produce very low, practically imperceptible noise when functioning.
- The assistive device should be truly affordable and cost effective.

We believe that achieving these requirements should be the main goal of the field, so we focus on these practical, consumer-oriented objectives in this article. We start with proposing a taxonomy of exoskeletons. We then discuss the improvements of these assistive devices that need to be achieved in near-term and long-term. Our views reflect both the professional, multi-disciplinary expertise of the authors and consumer perspective. The first author (**Figure 1**) sees both sides of the coin: he is an academic physician in Physical and Rehabilitation Medicine (with special focus on Neurorehabilitation) and in Gerontology and Geriatrics, and also a chronic, complete paraplegic, dependent on the wheelchair.

Our multidisciplinary approach intends—without underestimating the legitimate competition between different technical solutions and producers—to foster the optimization of assistive and rehabilitation devices, so that they could eventually become suitable and available to as many people in need as possible. Being one of the consumers too, the first author is looking forward to welcome this long awaited accomplishment, no matter which exoskeleton model meets the expectations first.

MAIN INDICATIONS

The currently available wearable powered exoskeletons serve two main functions. The first function is medical: providing assistive and rehabilitation technology for disabled people¹². The second function is non-medical: augmenting normal human physical capabilities.

As to the non-medical domain, we are covering it here only briefly. Noteworthy, non-medical exoskeletons have indirect linkages to medical issues, such as contributions to technology development and prevention of occupational injuries. The other uses encompass military devices for enhancement of soldier speed, power, and interactions with the equipment^{13–15}; exoskeletons for people professionally involved in critical rescue situations, such transportation of wounded individuals to safety from the war or civil disaster areas^{13–16}; civil applications for lifting and carrying heavy objects and patients¹⁷; and exoskeletons that augment human gait by assisting leg muscles and supporting a comfortable orthostatic posture. The target population for these applications includes the elderly with weak legs, personnel engaged in long-distance walking and/or performing flexion and extension movements with their lower limbs, and people utilizing exoskeleton appliances for entertainment¹⁸.

¹²<http://www.keego.com/biomechatronics/>

¹³<http://bleex.me.berkeley.edu/research/exoskeleton/bleex/>

¹⁴<http://bleex.me.berkeley.edu/research/exoskeleton/exohiker/>

¹⁵<http://bleex.me.berkeley.edu/research/exoskeleton/hulc/>

¹⁶<http://www.b-temia.com/defense-security-soldier-protection/>

¹⁷http://www.cyberdyne.jp/english/products/Lumbar_LaborSupport.html

¹⁸<http://asimo.honda.com/innovations/feature/body-weight-support-assist/>



FIGURE 1 | The picture of the first author Prof. Dr. Gelu Onose in wheelchair visiting patients in the NeuroRehabilitation Clinic Division of the 'Bagdasar-Arseni' Teaching Emergency Hospital, Bucharest, Romania. Reproduced with permission of the TV producer, Romanian Television.

The medical domain includes wearable devices utilized for prophylactic, rehabilitation, and assistance to disabled people. Taxonomy of these devices reflects the needs of different types of patients. A complete paraplegic would need a device that performs a set of sophisticated tasks, and there is currently no device with these capabilities. Such a device would require no effort from the paralyzed patient, and would do almost everything for the patient.

The device for a complete paraplegic would have to ensure:

- Complete assistance to orthostatic posture and walking, including the very difficult task of maintaining balance in order to avoid falls with severe consequences (Kannape and Lenggenhager, 2016).
- Prevention of spasticity and contractures (Arazpour et al., 2013) and related articular stiffness and pain.
- Prevention of venous and lymphatic stasis in the lower limbs, which could result in lower extremity edema and related complications due to prolonged standing (Louie et al., 2015).
- Partial compensation of the body weight to avoid injuries, such as fracture of the talus (Louie et al., 2015).

The complete paraplegia is classified by the scales of American Spinal Injury Association (ASIA) and International Spinal Cord Society (ISCoS) ("Standard Neurological Classification of Spinal Cord Injury"¹⁹, Frankel et al., 1969). Mechatronic wearable exoskeletons can be very useful as assistive devices in these cases because in complete chronic lesions of the spinal cord, as a rule, there are not enough remaining neural-muscular units to be trained. The possibility that exoskeletons could be used for rehabilitation in such cases, for example for rehabilitation of muscle activity, is still unclear (Arazpour et al., 2015b). The use

of exoskeletons still can induce neuroplasticity (Muresanu et al., 2012; Bryce et al., 2015; Louie et al., 2015) that improved patient interaction with the device²⁰. Additionally, a supplementary input from the patient's collected and decoded cortical activity might improve the man-machine interaction and the overall functional outcome.

The indications are different for incomplete paraplegics classified as AIS/ Frankel B-D, tetraplegics with low cervical spinal cord lesions classified as AIS/ Frankel C, and for hemiplegics and hemiparetics with fully preserved functionality of the non-affected hemi-body and limbs, and no sensorial, cognitive and balance serious impairments. The other potential beneficiaries of such exoskeletons are patients with neuromuscular and somatic impairments, including patients with polyneuropathia in lower limbs (Zeilig et al., 2012), multiple sclerosis, Parkinson's disease, hip or knee osteoarthritis²¹, limb fractures, and amputated limbs (Chen et al., 2013).

In hemiplegia cases, an H2 (Technaid S.L., Spain) robotic exoskeleton is more appropriate for gait rehabilitation (Bortole et al., 2015)—an exoskeleton based on two control strategies: (i) adaptive trajectory control for guiding the patient's limb within a desired path, the strategy that facilitates patient interaction with the device, and (ii) admittance control strategy that captures the user's movements during assistive training phase and reproduces it during active training phase (Bortole et al., 2013). The main feature of the wearable exoskeletons indicated to approach these kinds of patients is usually not an assistive control of the trunk. An essential feature of using such devices is training induced rehabilitation, where the exercise outcomes are compared to the time spent exercising (Louie et al., 2015).

²⁰http://who.int/disabilities/policies/actionplan/disability_action_plan_en.docx

²¹<http://www.keego.com/keego-in-action/>

The choice of the exoskeleton system and training regime critically depend on the extent of neurological impairment, the affected neural site (central, peripheral, or both), and the state of muscles (which can change with training). Additionally, it is important to take into account and correct by training such conditions as pain, contracture, osteoporosis, bed sores, and edema (Arazpour et al., 2013).

In the disabled elderly, the degree and characteristics of multimorbidity should be considered (Salive, 2013), particularly balance disorders, cardio-respiratory impairments, and para-physiological sarcopenia (Morley, 2012). The assistive and rehabilitation devices should be adjusted to these conditions. For instance, if an older adult has preserved, although altered, ability to stand-up and walk, extended trunk stabilization may not be needed. However, if the senior beneficiary has marked sensory and/or balance impairment, a maximally assistive devices might be more appropriate, in order to avoid frequent falls in the older population²². The frequency of falls is variable (6–34%) in different countries and regions: (Kalache et al., 2007). Falls represent some of the most serious negative events with considerable pathological consequences, including disabilities and life threatening conditions. Such negative events result in economic burden for the patients, their families, and the society²² (Kalache et al., 2007). These considerations will become progressively more important in the future, as the global process of demographic ageing intensifies²² (Kalache et al., 2007; Chen et al., 2013).

CURRENT TECHNOLOGIES

Currently popular wearable powered exoskeletons include Isocentric Reciprocating Gait Orthosis (IRGO), Hybrid Assistive Leg/Limb (HAL), ReWalk, Ekso, Mina, Wearable Power-Assist Locomotor (WPAL), Rex, Indego, Keeogo, Kickstart, Stride Management Assist, and ExoAtlet²³. These devices have been recently reviewed¹¹ (Arazpour et al., 2013, 2015a,b; Chen et al., 2013; Lajeunesse et al., 2015; Louie et al., 2015). These review articles documented the following characteristics of the existing systems:

- Benefits, including compared to the robotic exoskeletons fixed to treadmills (Chen et al., 2013; Louie et al., 2015).
- Safety (Lajeunesse et al., 2015).
- Efficacy in restoring walking¹¹ (Louie et al., 2015), including the performance in community settings (Lajeunesse et al., 2015).
- Clinical effectiveness outcomes, including such parameters as stair climbing performance, speed, mobility, quality of life, and independence¹¹ (Chen et al., 2013; Lajeunesse et al., 2015; Louie et al., 2015; Arazpour et al., 2015a,b; Lajeunesse et al., 2015).
- Walking distance (Benson et al., 2016).
- Vertical ground reaction force (vGRF) that quantifies the pattern and magnitude of mechanical loading

(Fineberg et al., 2013). This characteristic is particularly important for exoskeletons used by completely paralyzed persons.

- Training protocols (Lajeunesse et al., 2015; Louie et al., 2015).
- User satisfaction and secondary benefits and skills achieved with the exoskeleton (Lajeunesse et al., 2015).
- Level of scientific evidence (Lajeunesse et al., 2015).
- Cost-effectiveness outcomes¹¹.

Overall, the current literature reports both successes of exoskeleton technology and remaining problems. One important achievement is that wearable exoskeletons enable patients with complete paralysis of the lower part of the body to stand and walk. The speed of walking is modest; it varies with the level of injury and training duration (Louie et al., 2015). In addition to paraplegics, wearable exoskeletons can be used by patients with post-stroke hemiparesis (Bortole et al., 2015), as well as non-stroke neurological pathologies²⁴ (Esquenazi et al., 2012), such as cerebral palsy, myelome-ningocele, traumatic brain injury and Guillain Barré syndrome (Zeilig et al., 2012).

Wearable exoskeletons have advantages compared to Hip-Knee-Ankle-Foot-Orthoses (HKAFO). For example, powered gait orthosis (PGO) and the IRGO outperform HKAFO, as evident from the improvements in physiological cost index (PCI), distance walked, and walking speed (Arazpour et al., 2013). Additionally, wearable exoskeletons have certain advantages compared to the treadmill-based ones. With wearable exoskeletons, patients get much greater autonomy. The autonomy in turn facilitates rehabilitative training because the device can be used at home in addition to specialized facilities (Chen et al., 2013; Louie et al., 2015). Yet, autonomous use of wearable exoskeletons in community settings still has to be demonstrated (Lajeunesse et al., 2015). Overall, there is still a discrepancy between the consumer high expectations of autonomy and versatility and the actual effectiveness of using wearable exoskeletons (Benson et al., 2016).

A critical analysis of the assessments of exoskeletons as health technology¹¹ indicates the need for more research, particularly regarding long-term performance (Lajeunesse et al., 2015), assessment of training (Louie et al., 2015), randomized controlled trials, and economic analysis¹¹. Overall, the level of scientific evidence remains low in clinical trials of exoskeletons (Lajeunesse et al., 2015). Prevention and detection of medical complications in the exoskeleton users is yet another area that needs more research (Benson et al., 2016). A unified framework is needed for assessment of exoskeleton performance (Bryce et al., 2015). Such a framework is needed to make different clinical trials comparable. The framework should include at least six modules: functional applications, personal factors, device factors, external factors, activities, and health outcomes (Bryce et al., 2015).

Current exoskeletons (**Figure 2**) provide several secondary skills and benefits (Lajeunesse et al., 2015). Exoskeleton with compact design and small backpacks are particularly useful

²²<http://www.cdc.gov/Features/OlderAmericans/>

²³<http://www.exoatlet.com/>

²⁴<http://www.parker.com/literature/Exoskeleton/Parker%20Indego%20Brochure.pdf>



FIGURE 2 | Images of several types of exoskeletons. (A) Mindwalker mind-controlled exoskeleton could help the disabled walk again. With courtesy from Professor van der Kooij at University of Twente, NL. **(B)** The Keeego™ exoskeleton from B- TEMIA reveals the latest in an increase of the human systems (human augmentation systems) designed to help people walk more and better. Keeego™ eliminates several problems in patients with Parkinson's disease. With courtesy from Danielle Beaudoin at b-termia.com. **(C,D)** The ExoAtlet is a powered exoskeleton designed to assist patients during their rehabilitation after stroke, injury, or unsuccessful operation. ExoAtlet automatically repeats the natural patterns of walking, has electrical stimulation system, and physiological sensors. The control system of the ExoAtlet is unique: it collects data from body angles, allows to set the height and length of the step, which provides: (i) standing still; (ii) classic walking; (iii) walking on angled surface; (iv) stepping over obstacles; and (v) comfortable walking up & down stairs. ExoAtlet can be used in rehabilitation centers and at home²⁴. ExoAtlet can be controlled with the app on tablet when used in clinics. Experienced user of ExoAtlet use "thinking" crutch for control. With courtesy from Ekaterina Bereziy at exoatlet.ru.

in this respect (Chen et al., 2013) because they allow don, doff and wearing an exoskeleton while sitting in a wheelchair. These operations are possible using electrically actuated ReWalk and ExoAtlet, hydraulically actuated Ekso, and Indego, which combines electrically actuated hip and knee joints for both legs (Chen et al., 2013) with functional electrical stimulation (FES) of paralyzed, yet responsive muscles²³ (Chen et al., 2013). This combination of the exoskeleton action with FES training improves rehabilitation results.

FUTURE DIRECTIONS

Notwithstanding significant technological advancements in the field, the current exoskeleton systems still need much improvement for them to become practically efficient. The areas that need improvement remain almost the same as about one decade ago. Accordingly, our assessment made 8 years ago

(Onose et al., 2008) remains of current interest: exoskeletons need further miniaturization, optimization of actuators and sensors. Ideally, exoskeletons should become robotic orthotic suits wearable beneath the clothes.

In our perspectives from 2007 to 2008 (Figures 3, 5, 6), we conceived a soft cable-driven exo-suit device that can apply forces to the body to assist walking. Unlike traditional exoskeletons which contain rigid framing elements, the soft exosuit is worn like clothing and uses geared motors to pull on Bowden cables connected to the suit near the ankle (Asbeck et al., 2013).

Some of the authors of this article have been involved in extensive scientific research on this topic for almost 15 years (Onose et al., 2008, 2012a), and were awarded with the Gold Medal at the Inventions Saloon, in Genève, Switzerland, in 2008 (Figure 4).

We have not finished all the necessary tests and improvements of our prototype (Figure 5). Because of this delay, several

mechatronic wearable powered exoskeletons are in overall more advanced stage of development, including the clinical trials.

We re-emphasize ichnographically two important requirements to exoskeletons: (i) prevention of venous-lymphatic stasis in the lower limbs, and (ii) skin protection at the interface with the rigid structure of the exoskeleton.

Both issues can cause serious hurdles, especially during the long-time wearing of the assistive devices. Regarding the former issue, because of an impaired venous-lymphatic outflow in paralyzed persons (as well as patients with cardiovascular problems) (Guyton and Hall, 2006), regaining vertical posture with the aid of a wearable exoskeleton provokes regional stasis. This may cause discomfort and edema, including the risk of venous thrombosis, associated trophic somatic

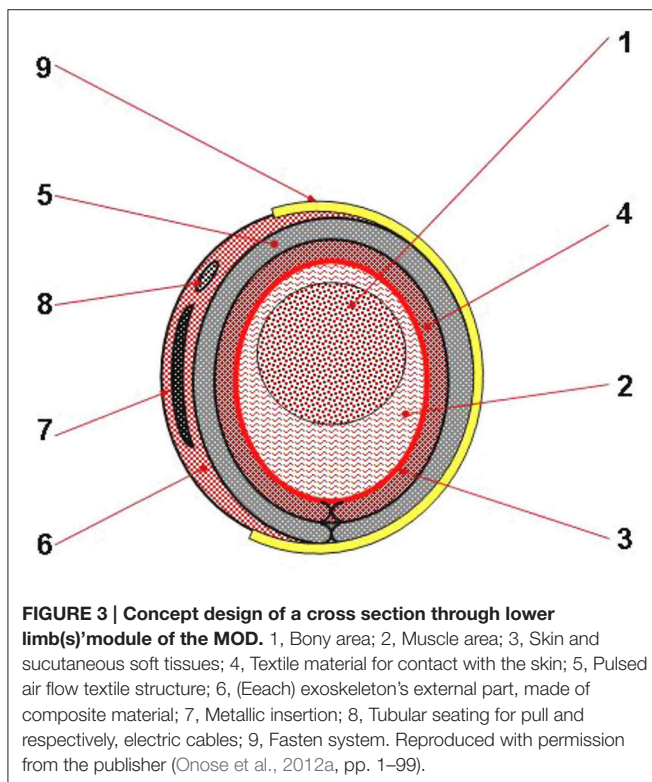
lesions, and secondary reactive heart conditions. As for the latter, an improper interface between the exoskeleton and the limb can result in mechanically generated lesions of the heel, respectively of the skin and subjacent soft tissue, which thus need for protection at the interface with the rigid structure of such devices (compressive, friction and tensile forces), as determined through performance metrics (van der Kooij et al., 2006). Therefore, we have conceived—for counteracting these problems—two components of our MOD.

Furthermore, regarding the underwear exoskeleton, we consider it ambitious and difficult, but worth trying developmental direction. Such an underwear exoskeleton would require a soft supporting structure actuated by strengthening/relaxing device. Specifically, we have foreseen, since 2006–2007 (Onose, 2007) the usage of electro-active polymers (EAPs). EAP materials seem to be the most applicable because of their large actuation forces and highest robustness (Bar-Cohen, 2005). These materials could be improved to act as artificial muscles (Bar-Cohen, 2005; Mirfakhrai et al., 2007).

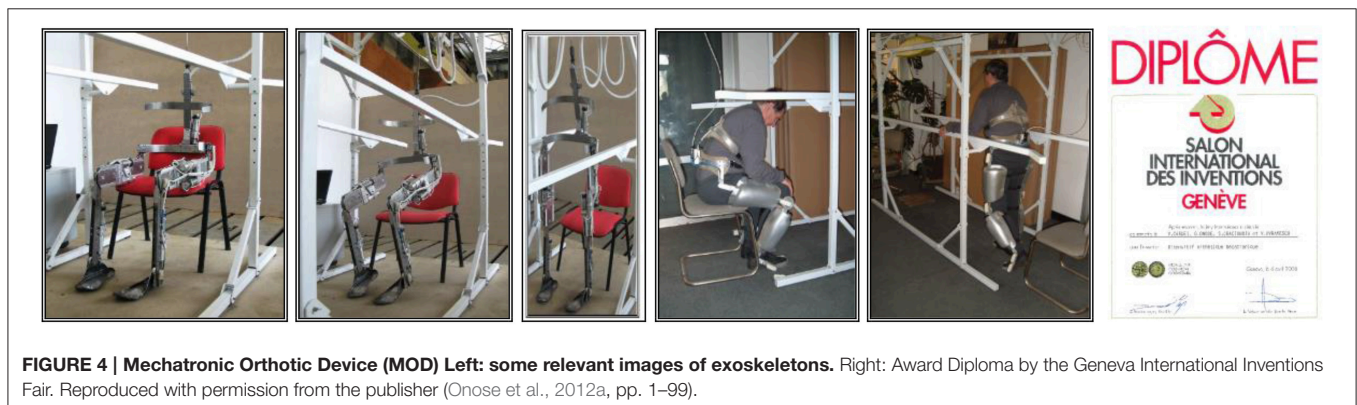
The development of such a device would be a revolutionary breakthrough, since the soft support structure in cadence with the artificial muscle contraction could not only move the lower limbs but also apply thrusts to the user's calf and thighs, thus mimicking a “muscle pump” and compensating for the returning, venous-lymphatic circulation (Onose, 2007; Onose et al., 2012a). However, during the last decade not enough progress has been made in the development and subsequent translational implementation of EAPs. We still need to improve the safety, controllability and energy supply for EAPs²⁵.

Another extremely challenging and still incompletely solved item, as mentioned above, is the problem of balance maintenance using advanced mechatronic wearable exoskeletons. Reliable balance is needed for prevention of falls during exoskeleton usage.

At least regarding the most difficult and demanding—in need for total assistance from the apparatus—but, at the same time, an essential category of potential beneficiaries, *the complete paraplegics*, all current devices require supplementary aids for securing the balance: walkers, crutches or—at



²⁵http://sbdi.co.kr/cart/data/info/IDTechEx_Electroactive_Polymers_and_Devices_2013-2018_Sample.pdf?ckattempt=1



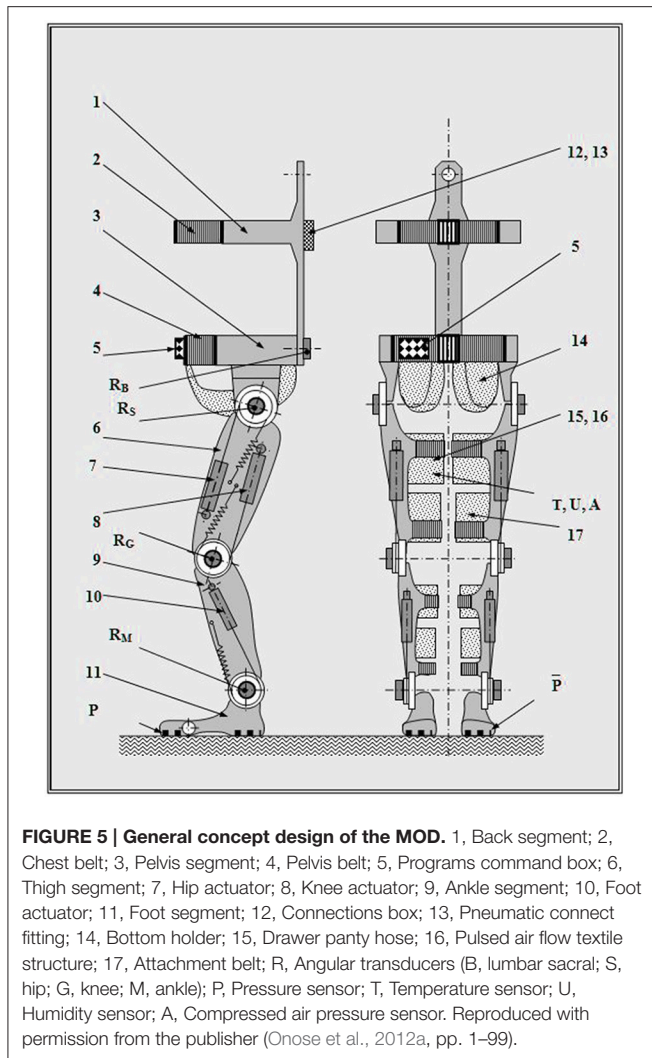


FIGURE 5 | General concept design of the MOD. 1, Back segment; 2, Chest belt; 3, Pelvis segment; 4, Pelvis belt; 5, Programs command box; 6, Thigh segment; 7, Hip actuator; 8, Knee actuator; 9, Ankle segment; 10, Foot actuator; 11, Foot segment; 12, Connections box; 13, Pneumatic connect fitting; 14, Bottom holder; 15, Drawer panty hose; 16, Pulsed air flow textile structure; 17, Attachment belt; R, Angular transducers (B, lumbar sacral; S, hip; G, knee; M, ankle); P, Pressure sensor; T, Temperature sensor; U, Humidity sensor; A, Compressed air pressure sensor. Reproduced with permission from the publisher (Onose et al., 2012a, pp. 1–99).

best—canes. In sum, the powered exoskeletons that assist over-ground walking require the user to maintain balance (Swinnen et al., 2014; Yoshimoto et al., 2015; Louie and Eng, 2016).

As an illustration of the current state of this problem, it is worth mentioning a study that reported loss of balance (LOB) and falls as the primary safety outcomes (Kolakowsky-Hayner et al., 2013). Additionally, the EU FP7 “BALANCE” project (Veneman, 2014) made an observation that “none of the devices available on the market use the exoskeleton itself to support postural balance.” Therefore, the goal of the BALANCE project is to research the possibility of using the exoskeleton to maintain balance and prevent falls. The project is organized as a 4 year Specific Targeted Research Project (STREP)²⁶ and has the practical objectives to “create a human-cooperative robotic postural balance controller framework,” in order to “implement the human-cooperative postural balance controller on a real exoskeleton” and to

²⁶http://balance-fp7.eu/project_structure.php

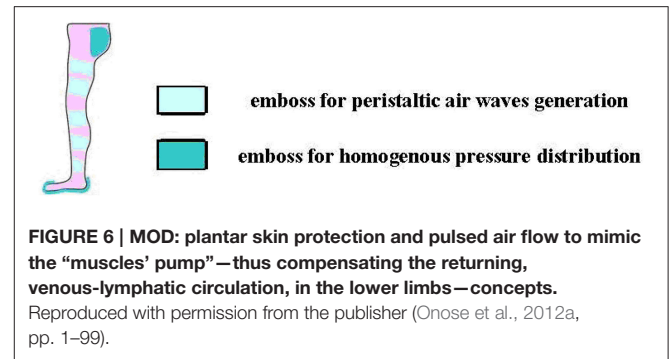


FIGURE 6 | MOD: plantar skin protection and pulsed air flow to mimic the “muscles’ pump”—thus compensating the returning, venous-lymphatic circulation, in the lower limbs—concepts.

Reproduced with permission from the publisher (Onose et al., 2012a, pp. 1–99).

“evaluate the developed concepts in subjects walking with the exoskeleton.”²⁷

Lastly, one more direction to improve the performances of a mechatronic wearable exoskeleton, is to empower it with brain’s voluntary motor commands using a brain-machine-interface (BMI) (Onose et al., 2012b; Lebedev, 2014). The feasibility of such a BMI has been demonstrated in an experiment conducted in rhesus monkeys (Fitzsimmons et al., 2009). In this study, monkeys were implanted with invasive multielectrode arrays in the leg representation of the sensorimotor cortex. The animals were trained to walk bipedally on a treadmill. Activity of several hundred cortical neurons was recorded and converted into the kinematics of lower limb movements. This BMI was able to extract lower limb kinematics when monkeys walked both forward and backward on the treadmill.

These results justified the foundation of the Walk Again Project with the goal of the advancement of BMI-controlled exoskeletons (Nicollelis and Lebedev, 2009). An European project, called Mindwalker (Figure 2A), declared the goal of controlling an exoskeleton with electroencephalographic (EEG) and electromyographic (EMG) recordings (Cheron et al., 2012). Decoding of leg movements from EEGs has been demonstrated in humans walking on a treadmill (Presacco et al., 2012). Moreover, a BMI that decoded steady-state visual evoked potentials (SSVEP) from EEGs was employed to operate a lower-limb exoskeleton by healthy human subjects (Kwak et al., 2015).

The progress in BMIs for bipedal walking opens the perspective of neural control of exoskeleton-assisted walking by a completely paralyzed subject. While a slow control of walking could be provided by EEG-based BMIs, invasive BMIs hold promise to enable faster and more accurate control. However, invasive recordings are risky, so there have been no studies of invasive BMIs for walking control in humans. Additionally, it is unclear whether patients could become proficient enough to efficiently control an exoskeleton through an invasive or noninvasive BMI (Onose et al., 2012b). In this respect, we are satisfied of having deployed a clinical trial in the motor imagery BMI domain, by enrolling nine chronic tetraplegics volunteers (the largest lot for such type of trial, at that time), to control

²⁷<http://balance-fp7.eu/objectives.php>

wirelessly a robotic arm by EEG commands (Onose et al., 2012b).

We have foreseen also the possibility of empowering MOD's actuating systems by voluntary commands extracted from EEGs (Onose et al., 2008). From this point of view, we have expressed our confidence in the success of the Walk Again²⁸ and Mindwalker projects²⁹ (Figure 2).

CONCLUSION

We suggest that the joint perspective of health professionals, manufacturers and consumers on the mechatronic wearable exoskeletons should contribute to a comprehensive agenda regarding the development of devices that assist and rehabilitate bipedal posture and walking to severely impaired people. The technological advances, together with collaborative endeavors of multidisciplinary teams of researchers, could foster the progress and pave the way to the long awaited optimized and practical exoskeleton.

²⁸<http://virtualreality.duke.edu/project/walk-again-project/>

²⁹<https://www.utwente.nl/ctw/bw/research/projects/MINDWALKER/>

REFERENCES

- Arazpour, M., Bani, M. A., Hutchins, S. W., and Jones, R. K. (2013). The physiological cost index of walking with mechanical and powered gait orthosis in patients with spinal cord injury. *Spinal Cord* 51, 356–359. doi: 10.1038/sc.2012.162
- Arazpour, M., Hutchins, S. W., and Bani, M. A. (2015a). The efficacy of powered orthoses on walking in persons with paraplegia. *Prosthet. Orthot. Int.* 39, 90–99. doi: 10.1177/0309364613520031
- Arazpour, M., Samadian, M., Bahramizadeh, M., Joghtaei, M., Maleki, M., Bani, M. A., et al. (2015b). The efficiency of orthotic interventions on energy consumption in paraplegic patients: a literature review. *Spinal Cord* 53, 168–175. doi: 10.1038/sc.2014.227
- Asbeck, A. T., Dyer, R. J., Larusson, A. F., and Walsh, C. J. (2013). Biologically-inspired soft exosuit. *IEEE Int. Conf. Rehabil. Robot.* 2013:6650455. doi: 10.1109/icorr.2013.6650455
- Bar-Cohen, Y. (2005). Current and future developments in artificial muscles using electroactive polymers. *Expert Rev. Med. Devices* 2, 731–740. doi: 10.1586/s17434440.2.6.731
- Benson, I., Hart, K., Tussler, D., and van Middendorp, J. J. (2016). Lower-limb exoskeletons for individuals with chronic spinal cord injury: findings from a feasibility study. *Clin. Rehabil.* 30, 73–84. doi: 10.1177/0269215515575166
- Bortole, M., del Ama, A., Rocon, E., Moreno, J. C., Brunetti, F., and Pons, J. L. (2013). "A robotic exoskeleton for overground gait rehabilitation," in *Proceedings of 2013 IEEE International Conference on Robotics and Automation (Karlsruhe)*, 3356–3361.
- Bortole, M., Venkatakrishnan, A., Zhu, F., Moreno, J. C., Francisco, G. E., Pons, J. L., et al. (2015). The H2 robotic exoskeleton for gait rehabilitation after stroke: early findings from a clinical study. *J. Neuroeng. Rehabil.* 12:54. doi: 10.1186/s12984-015-0048-y
- Bryce, T. N., Dijkers, M. P., and Kozlowski, A. J. (2015). Framework for assessment of the usability of lower-extremity robotic exoskeletal orthoses. *Am. J. Phys. Med. Rehabil.* 94, 1000–1014. doi: 10.1097/PHM.0000000000000321
- Chen, G., Chan, C. K., Guo, Z., and Yu, H. (2013). A review of lower extremity assistive robotic exoskeletons in rehabilitation therapy. *Crit. Rev. Biomed. Eng.* 41, 343–363 doi: 10.1615/critrevbiomedeng.2014010453

AUTHOR CONTRIBUTIONS

All authors listed, have made substantial, direct and intellectual contribution to the work, and approved it for publication.

ACKNOWLEDGMENTS

The authors thank to: Florin Alexiuc, Ioana Andone, Aurelian Angheliescu, Cornel Baluță, Ana Maria Bumbea, Eftalea Cârpuș, Crenguța Coman, Cristina Daia, Alexandru Vlad Ciurea, Jean Ciurea, Radu Stefan Craciunoiu, Ionut Catalin Dimache, Constantin Dogariu, Angela Dorogan, Condurache Dumitru, Maria Gratiela Epureanu, Eva Gheorghită, Carmen Giuglea, Roxana Grejdanescu, Monica Haras, Gheorghe Huian, Noela Ionescu, Andrada Mirea, Marius Modreanu, Alexandru Moldovanu, Cristian Nicolau, Raluca Magdalena Niță, Liliana Onose, Horia Catalin Orasanu, Loredana Theodora Păun, Vergil Racicovschi, Dorin Roșu, Aura Spînu, Olga Surdu, Emilia Visileanu, and Tudor Ursu—for their contribution to the inter-/multi-disciplinary scientific research project that has resulted in the achievement of the MOD and/or to the afferent monography (Onose et al., 2012a), which constitutes a valuable background resource, including for this article.

- Cheron, G., Duvinage, M., De Saedeleer, C., Castermans, T., Bengoetxea, A., Petieau, M., et al. (2012). From spinal central pattern generators to cortical network: integrated BCI for walking rehabilitation. *Neural Plast.* 2012:375148. doi: 10.1155/2012/375148
- Esquenazi, A., Talaty, M., Packel, A., and Saulino, M. (2012). The ReWalk powered exoskeleton to restore ambulatory function to individuals with thoracic-level motor-complete spinal cord injury. *Am. J. Phys. Med. Rehabil.* 91, 911–921. doi: 10.1097/PHM.0b013e318269d9a3
- Fineberg, D. B., Asselin, P., Harel, N. Y., Agranova-Breyter, I., Kornfeld, S. D., Bauman, W. A., et al. (2013). Vertical ground reaction force-based analysis of powered exoskeleton-assisted walking in persons with motor-complete paraplegia. *J. Spinal Cord Med.* 36, 313–321. doi: 10.1179/2045772313Y.0000000126
- Fitzsimmons, N. A., Lebedev, M. A., Peikon, I. D., and Nicolelis, M. A. L. (2009). Extracting kinematic parameters for monkey bipedal walking from cortical neuronal ensemble activity. *Front. Integr. Neurosci.* 3:3. doi: 10.3389/fnro.07.003.2009
- Frankel, H. L., Hancock, D. O., Hyslop, G., Melzak, J., Michaelis, L. S., Ungar, G. H., et al. (1969). The value of postural reduction in the initial management of closed injuries of the spine with paraplegia and tetraplegia. I. *Paraplegia* 7, 179–192. doi: 10.1038/sc.1969.30
- Guyton, A. C., and Hall, J. E. (eds.). (2006). *Textbook of Medical Physiology 11th (International) Edition – Chap. 15 (Vascular Distensibility and Functions of the Arterial and Venous Systems) and 20 (Cardiac Output, Venous Return, and Their Regulation)*. Philadelphia, PA: Elsevier Inc.
- Kalache, A., Fu, D., and Yoshida, S. (2007). *WHO Global Report on Falls Prevention in Older Age*. Geneva: WHO Press.
- Kannape, O. A., and Lenggenhager, B. (2016). Engineered embodiment: Comment on "The embodiment of assistive devices—from wheelchair to exoskeleton" by M. Pazzaglia and M. Molinari Oliver. *Phys. Life Rev.* 16, 181–183. doi: 10.1016/j.plrev.2016.01.011
- Kolakowsky-Hayner, S. A., Crew, J., Moran, S., and Shah, A. (2013). Safety and feasibility of using the Ekso™ bionic exoskeleton to aid ambulation after spinal cord injury. *J. Spine* S4:003. doi: 10.4172/2165-7939.S4-003
- Kwak, N. S., Müller, K. R., and Lee, S. W. (2015). A lower limb exoskeleton control system based on steady state visual evoked potentials. *J. Neural Eng.* 12:056009. doi: 10.1088/1741-2560/12/5/056009

- Lajeunesse, V., Vincent, C., Routhier, F., Careau, E., and Michaud, F. (2015). Exoskeletons' design and usefulness evidence according to a systematic review of lower limb exoskeletons used for functional mobility by people with spinal cord injury. *Disabil. Rehabil. Assist. Technol.* 4, 1–13. doi: 10.3109/17483107.2015.1080766
- Lebedev, M. (2014). Brain-machine interfaces: an overview. *Transl. Neurosci.* 5, 99–110. doi: 10.2478/s13380-014-0212-z
- Louie, D. R., and Eng, J. J. (2016). Powered robotic exoskeletons in post-stroke rehabilitation of gait: a scoping review. *J. Neuroeng. Rehabil.* 13, 53. doi: 10.1186/s12984-016-0162-5
- Louie, D. R., Eng, J. J., Lam, T., and Spinal Cord Injury Research Evidence (SCIRE) Research Team (2015). Gait speed using powered robotic exoskeletons after spinal cord injury: a systematic review and correlational study. *J. Neuroeng. Rehabil.* 12, 82. doi: 10.1186/s12984-015-0074-9
- Mirfakhrai, T., Madden, J. D. W., and Baughman, R. H. (2007). Polymer artificial muscles. *Mater. Today* 10, 30–38. doi: 10.1016/S1369-7021(07)70048-2
- Morley, J. E. (2012). Sarcopenia in the elderly. *Fam. Pract.* 29(Suppl. 1), i44–i48. doi: 10.1093/fampra/cm063
- Muresanu, D. F., Buzoianu, A., Florian, S. I., and von Wild, T. (2012). Towards a roadmap in brain protection and recovery. *J. Cell. Mol. Med.* 16, 2861–2871. doi: 10.1111/j.1582-4934.2012.01605.x
- Nicolelis, M. A., and Lebedev, M. A. (2009). Principles of neural ensemble physiology underlying the operation of brain-machine interfaces. *Nat. Rev. Neurosci.* 10, 530–540. doi: 10.1038/nrn2653
- Onose, G. (2007). *Manual de Recuperare, Medicină Fizică și Balneoclimatologie – Noțiuni de Baza și Actualități*, Vol. I. București: Editura Medicală.
- Onose, G., Cârdei, V., Ciurea, A. V., Ciurea, J., Onose, L., Angheliescu, A., et al. (2008). "Achievement of an experimental mechatronic orthotic device to assist/rehabilitate orthostatism and walk in patients with complete paraplegia and other specific severe disabling conditions (short form)," in *Proceedings of the 7th Mediterranean Congress of Physical and Rehabilitation Medicine* (Portorose; Torino: Edizioni Minerva Medica), 40–42.
- Onose, G., Cârdei, V., Crăciunoiu, Ș T., and Avramescu, V. (eds.). (2012a). *Mechatronic Orthotic Device. Concept, Sumptions and Functional-Construction Aspects*. București: Editura ICTCM.
- Onose, G., Grozea, C., Angheliescu, A., Daia, C., Sinescu, C. J., Ciurea, A. V., et al. (2012b). On the feasibility of using motor imagery EEG-based brain-computer interface in chronic tetraplegics for assistive robotic arm control: a clinical test and long-term post-trial follow-up. *Spinal Cord* 50, 599–608. doi: 10.1038/sc.2012.14
- Presacco, A., Forrester, L. W., and Contreras-Vidal, J. L. (2012). Decoding intralimb and inter-limb kinematics during treadmill walking from scalp electroencephalographic (EEG) signals, *IEEE Trans. Neural Syst. Rehabil. Eng.* 20, 212–219. doi: 10.1109/TNSRE.2012.2188304
- Salive, M. E. (2013). Multimorbidity in older adults. *Epidemiol. Rev.* 35, 75–83. doi: 10.1093/epirev/mxs009
- Swinnen, E., Beckwée, D., Meeusen, R., Baeyens, J. P., and Kerckhofs, E. (2014). Does robot-assisted gait rehabilitation improve balance in stroke patients? A systematic review. *Top Stroke Rehabil.* 21, 87–100. doi: 10.1310/tsr2102-87
- Talley Watts, L., Long, J. A., Manga, V. H., Huang, S., Shen, Q., and Duong, T. Q. (2015). Normobaric oxygen worsens outcome after a moderate traumatic brain injury. *J. Cereb. Blood. Flow. Metab.* 35, 1137–1144. doi: 10.1038/jcbfm.2015.18
- Uebelhart, D., Bernard, J., Hartmann, D. J., Moro, L., Roth, M., Uebelhart, B., et al. (2000). Modifications of bone and connective tissue after orthostatic bedrest. *Osteoporos. Int.* 11, 59–67. doi: 10.1007/s001980050007
- van der Kooij, H., Veneman, J. F., Ekkelenkamp, R., (2006). "Compliant actuation of exoskeletons," in *Robots Towards New Applications*, ed A. Lazinicca (Source: ISBN 978-3-86611-314-5, pp. 784, ARS/pLV. I-Tech Education and Publishing; DOI: 10.5772/33). Available online at: <https://core.ac.uk/download/files/363/11473533.pdf>
- Veneman, J. F. (2014). "Chapter: exoskeletons supporting postural balance – the balance project," in *Replace, Repair, Restore, Relieve – Bridging Clinical and Engineering Solutions in Neurorehabilitation. Vol. 7 of the Series Biosystems and Biorobotics*, eds W. Jensen, O. K. Andersen, and M. Akay (San Sebastian: Springer International Publishing Switzerland), 203–208.
- Yoshimoto, T., Shimizu, I., Hiroi, Y., Kawaki, M., Sato, D., and Nagasawa, M. (2015). Feasibility and efficacy of high-speed gait training with a voluntary driven exoskeleton robot for gait and balance dysfunction in patients with chronic stroke: nonrandomized pilot study with concurrent control. *Int. J. Rehabil. Res.* 38, 338–343. doi: 10.1097/MRR.0000000000000132
- Zeilig, G., Weingarden, H., Zwecker, M., Dudkiewicz, I., Bloch, A., and Esquenazi, A. (2012). Safety and tolerance of the ReWalk™ exoskeleton suit for ambulation by people with complete spinal cord injury: a pilot study. *J. Spinal Cord Med.* 35, 96–101. doi: 10.1179/2045772312Y.0000000003

Disclosures: GO, VC, ȘC, and VA have been the main researchers involved in achieving the aforementioned Romanian original experimental prototype of mechatronic orthotic device (MOD), but as it could be determined throughout this article, there hasn't been any intention at all to favor in description this model of device, including with the fact that it has been honestly acknowledged in the body text that currently, there are a series of other such models significantly closer to reach the optimal, desired form.

Conflict of Interest Statement: The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Copyright © 2016 Onose, Cârdei, Crăciunoiu, Avramescu, Opreș, Lebedev and Constantinescu. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) or licensor are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.