

## **CADTH Reference List**

# Robotic-Assisted Gait Training for Children With Cerebral Palsy

July 2022

**Summary of Abstracts** 



Authors: Thyna Vu, Sarah C. McGill

Cite As: Robotic-Assisted Gait Training for Children With Cerebral Palsy. (CADTH reference list: summary of abstracts). Ottawa: CADTH; 2022 Jul.

**Disclaimer:** The information in this document is intended to help Canadian health care decision-makers, health care professionals, health systems leaders, and policy-makers make well-informed decisions and thereby improve the quality of health care services. While patients and others may access this document, the document is made available for informational purposes only and no representations or warranties are made with respect to its fitness for any particular purpose. The information in this document should not be used as a substitute for professional medical advice or as a substitute for the application of clinical judgment in respect of the care of a particular patient or other professional judgment in any decision-making process. The Canadian Agency for Drugs and Technologies in Health (CADTH) does not endorse any information, drugs, therapies, treatments, products, processes, or services.

While care has been taken to ensure that the information prepared by CADTH in this document is accurate, complete, and up to date as at the applicable date the material was first published by CADTH, CADTH does not make any guarantees to that effect. CADTH does not guarantee and is not responsible for the quality, currency, propriety, accuracy, or reasonableness of any statements, information, or conclusions contained in any third-party materials used in preparing this document. The views and opinions of third parties published in this document do not necessarily state or reflect those of CADTH.

CADTH is not responsible for any errors, omissions, injury, loss, or damage arising from or relating to the use (or misuse) of any information, statements, or conclusions contained in or implied by the contents of this document or any of the source materials.

This document may contain links to third-party websites. CADTH does not have control over the content of such sites. Use of third-party sites is governed by the third-party website owners' own terms and conditions set out for such sites. CADTH does not make any guarantee with respect to any information contained on such third-party sites and CADTH is not responsible for any injury, loss, or damage suffered as a result of using such third-party sites. CADTH has no responsibility for the collection, use, and disclosure of personal information by third-party sites.

Subject to the aforementioned limitations, the views expressed herein do not necessarily reflect the views of Health Canada, Canada's provincial or territorial governments, other CADTH funders, or any third-party supplier of information.

This document is prepared and intended for use in the context of the Canadian health care system. The use of this document outside of Canada is done so at the user's own risk.

This disclaimer and any questions or matters of any nature arising from or relating to the content or use (or misuse) of this document will be governed by and interpreted in accordance with the laws of the Province of Ontario and the laws of Canada applicable therein, and all proceedings shall be subject to the exclusive jurisdiction of the courts of the Province of Ontario, Canada.

The copyright and other intellectual property rights in this document are owned by CADTH and its licensors. These rights are protected by the Canadian *Copyright Act* and other national and international laws and agreements. Users are permitted to make copies of this document for non-commercial purposes only, provided it is not modified when reproduced and appropriate credit is given to CADTH and its licensors.

About CADTH: CADTH is an independent, not-for-profit organization responsible for providing Canada's health care decision-makers with objective evidence to help make informed decisions about the optimal use of drugs, medical devices, diagnostics, and procedures in our health care system.

Funding: CADTH receives funding from Canada's federal, provincial, and territorial governments, with the exception of Quebec.

Questions or requests for information about this report can be directed to requests@cadth.ca

## Key Messages

- No evidence was identified regarding the clinical effectiveness of robotic-assisted gait training (RAGT) versus no therapy for children with cerebral palsy.
- Eight randomized controlled trials and 1 non-randomized study were identified regarding the clinical effectiveness of RAGT training versus alternative therapies for children with cerebral palsy.
- No evidence was identified regarding the cost-effectiveness of RAGT versus no therapy for children with cerebral palsy.
- No evidence was identified regarding the cost-effectiveness of RAGT versus alternative therapies for children with cerebral palsy.

### **Research Questions**

- 1. What is the clinical effectiveness of robotic-assisted gait training (RAGT) versus no therapy for children with cerebral palsy?
- 2. What is the clinical effectiveness of RAGT versus alternative therapies for children with cerebral palsy?
- 3. What is the cost-effectiveness of RAGT versus no therapy for children with cerebral palsy?
- 4. What is the cost-effectiveness of RAGT versus alternative therapies for children with cerebral palsy?

## Methods

### Literature Search Methods

A limited literature search was conducted by an information specialist on key resources including Medline through Ovid, Scopus, the Cochrane Database of Systematic Reviews, the International HTA Database, and the websites of Canadian and major international health technology agencies, as well as a focused internet search. The search strategy comprised both controlled vocabulary, such as the National Library of Medicine's MeSH (Medical Subject Headings), and keywords. The main search concepts were robotic assisted gait training and cerebral palsy. No filters were applied to limit the retrieval by study type. Where possible, retrieval was limited to the human population. The search was completed on June 13, 2022, and limited to English-language documents published since January 1, 2015. Internet links were provided, where available.

### **Selection Criteria and Summary Methods**

One reviewer screened literature search results (titles and abstracts) and selected publications according to the inclusion criteria presented in <u>Table 1</u>. Full texts of study publications were not reviewed. The overall summary of findings was based on information available in the abstracts of selected publications.



### **Table 1: Selection Criteria**

Criteria	Description
Population	Children with cerebral palsy (≤ 18 years old)
Intervention	Robotic-assisted gait training
Comparator	Q1 and Q3: No treatment Q2 and Q4: Alternative therapies (e.g., physiotherapy, manual gait training, water-based therapy)
Outcomes	Q1: Clinical effectiveness (e.g., gross motor function, gait parameters, walking endurance, independence) Q2: Cost-effectiveness (e.g., incremental cost per health benefit, quality-adjusted life-years gained)
Study designs	Health technology assessments, systematic reviews, randomized controlled trials, non-randomized studies, economic evaluations

### Results

Eight randomized controlled trials (RCTs)<sup>1-8</sup> and 1 non-randomized study<sup>9</sup> were identified regarding the clinical effectiveness of RAGT versus alternative therapies for children with cerebral palsy. No relevant health technology assessments, systematic reviews, or economic evaluations were identified.

Additional references of potential interest that did not meet the inclusion criteria are provided in <u>Appendix 1</u>.

## **Overall Summary of Findings**

Eight RCTs<sup>1-8</sup> and 1 non-randomized study<sup>9</sup> were identified regarding the clinical effectiveness of RAGT versus alternative therapies for children with cerebral palsy. A detailed summary of these studies can be found in <u>Table 2</u>.

Two RCTs<sup>1,2</sup> compared RAGT to usual or standard care: 1 RCT<sup>1</sup> found no change on Gross Motor Function Measure (GMFM) or timed walking tests after either treatment, while the other<sup>2</sup> found RAGT was associated with improved locomotor function and functional capabilities compared to standard care. The latter study also noted that functional improvements were mainly seen in children classified as level II to III on the Gross Motor Function Classification System (GMFCS) scale, compared to those classified as level IV.<sup>2</sup>

Three RCTs<sup>4-6</sup> compared RAGT to different types of treadmill-related training programs. One RCT<sup>4</sup> compared robotic-assisted treadmill exercise to partial body weight-supported treadmill exercise or anti-gravity treadmill exercise, and reported they did not differ significantly on speed, gait analysis, or GMFM scores. One RCT<sup>5</sup> compared RAGT to treadmill only, and reported that the RAGT group had significant improvements post-intervention but the treadmill-only group did not. Another RCT<sup>6</sup> compared RAGT to robotic treadmill training with controlled resistance for children classified as levels I to IV on the GMFCS scale; they found that outcomes did not change significantly after RAGT, but did improve significantly following robotic treadmill training with controlled resistance.

Two RCTs<sup>7,8</sup> and 1 non-randomized study<sup>9</sup> compared RAGT to physiotherapy; all reported significant improvements in the RAGT group on locomotor parameters,<sup>7</sup> kinetic data,<sup>7</sup> kinematic data,<sup>8</sup> and GMFM scores<sup>8,9</sup> when comparing pre- and post-intervention. The non-randomized study<sup>9</sup> reported that no differences were observed between RAGT or task-oriented physiotherapy. The 2 RCTs<sup>7,8</sup> did not report between-group comparisons in the abstract.

One RCT<sup>3</sup> compared RAGT to non-assisted gait training for children classified as levels I to III on the GMFCS scale, and reported that RAGT was associated with improved hip angle and limb symmetry, but not gait speed.

No evidence was identified specific to children under the age of 6 with cerebral palsy classified as level V on the GMFCS scale; therefore, no summary can be provided for this specific subgroup. In addition, no relevant literature was found regarding the clinical effectiveness of RAGT compared to no treatment, or the cost-effectiveness of RAGT compared to no treatments for children with cerebral palsy; therefore, no summary can be provided.

### **Table 2: Summary of Included Clinical Effectiveness Studies**

Study citation	Study design, population	Intervention and comparator(s)	Relevant outcome(s)	Authors' conclusions			
Randomized controlled trials							
Ammann-Reiffer et al. (2020) <sup>1</sup>	Study design: Randomized crossover trial Population: Children with spastic CP (mean age = 11.3) N = 16	Intervention: RAGT (3 sessions per week for 5 weeks) Comparator: Usual care (for 5 weeks)	Primary outcome: • GMFM Dimension E Secondary outcomes: • GMFM Dimension D • Timed walking tests before and after each treatment sequence and after a 5-week follow-up	No outcomes changed significantly after RAGT or usual care, and there were no period, follow-up, or carry-over effects observed.			
Jin et al. (2020) <sup>2</sup>	Study design: Single- centre, single-blinded randomized crossover trial Population: Children with CP (GMFCS II to IV, age range = 6.75 ± 2.15 years) N = 20	Intervention: RAGT (Walkbot-K system, 3 sessions per week for 6 weeks) Comparator: Standard care (2 to 4 sessions for 6 weeks)	<ul> <li>GMFM score</li> <li>WeeFIM</li> <li>COPM score</li> </ul>	RAGT led to benefits in locomotor function and functional capability for daily activities. Factors associated with functional improvements were mainly observed in children classified as levels II to III (GMFCS), compared to those at level IV.			

		Intervention and				
Study citation	Study design, population	comparator(s)	Relevant outcome(s)	Authors' conclusions		
Kawasaki et al. (2020) <sup>3</sup>	Study design: RCT Population: Children with spastic CP (GMFCS I to III) N = 10	Intervention: RAGT Comparator: Non- assisted gait training	<ul> <li>Maximum hip angle</li> <li>Limb symmetry</li> <li>Gait speed</li> </ul>	Limb symmetry improved significantly after RAGT but not in the comparator group. RAGT was also associated with improved maximum hip flexion and extension angle. There was no change in gait speed.		
Aras et al. (2019)⁴	Study design: RCT Population: Children with spastic CP (mean age = 9.3 ± 2.3; age range = 6 to 14) N = 29	Intervention: RATE Comparator: PBWSTE, ATE	<ul> <li>Three-dimensional gait analysis</li> <li>Open-circle indirect calorimeter</li> <li>6MWT</li> <li>GMFM scale</li> </ul>	The ATE and RATE groups had significantly improved oxygen consumption, but not the PBWSTE group. There was no statistically significant change compared to baseline on walking speed or gait analysis after treatment, or between groups on GMFM-D, GMFM-E, and 6MWT.		
Wu et al. (2017)⁵	Study design: RCT Population: Children with CP N = 23	Intervention: Robotic- assisted treadmill training (3 times per week for 6 weeks) Comparator: Treadmill training only (manual assistance provided as needed; 3 times per week for 6 weeks)	<ul> <li>Walking speed</li> <li>Six-minute walking distance</li> </ul>	A greater increase in 6-minute walking distance was seen after robotic training compared to treadmill-only training. The robotic group saw significant increases in walking speed and 6-minute walking distance. No significant change was seen after treadmill-only training.		
Wu et al. (2017)6	Study design: RCT Population: Children with spastic CP (mean age = 10.6; age range = 6 to 14; GMFCS I to IV) N = 23	Intervention: Robotic treadmill training with controlled assistance (3x per week for 6 weeks) Comparator: Robotic treadmill training with controlled resistance	<ul> <li>Overground walking speed</li> <li>Six-minute walk distance</li> <li>GMFM scores</li> </ul>	Overground gait speed and 6-minute walk distance did not change significantly after robotic assistance training; both outcomes improved significantly after resistance training, and was still significantly greater 8 weeks after training.		
Randomized controlled trials with between-group comparison not presented in abstract						
Wallard et al. (2018) <sup>7</sup>	Study design: RCT Population: Children with CP N = 36	Intervention: RAGT (Lokomat; 20 sessions) Comparator: Physiotherapy	<ul> <li>Locomotor parameters</li> <li>Kinetic data</li> </ul>	Significant differences were seen for the treatment group pre- and post- intervention.		

Study citation	Study design, population	Intervention and comparator(s)	Relevant outcome(s)	Authors' conclusions		
Wallard et al. (2017) <sup>8</sup>	Study design: RCT Population: Children with bilateral spastic CP N = 30	Intervention: RAGT (20 sessions with Lokomat) Comparator: Physiotherapy (daily)	<ul> <li>Kinematic data</li> <li>GMFM-D and GMFM-E</li> </ul>	Significant improvements were observed between the treatment group pre- and post-values for both outcomes.		
Non-randomized studies						
Peri et al. (2017) <sup>9</sup>	Study design: NRS Population: Children	Intervention: RAGT (4 per week for 10 weeks) • GMFM-88 • GMFM-E	• GMFM-88 • GMFM-E	No differences across the 4 protocols were highlighted.		
	with CP (age 4 to 17) N = 44	Comparator: • RAGT + TOP (2 + 2 per week for 10 weeks) • RAGT + TOP (5 + 5 sessions per week for 4 weeks) • TOP alone (4 per week for 10 weeks)	• GMFM-66	Mixed approaches (RAGT + TOP) did not show significant changes, while the RAGT- or TOP-only protocols saw significant improvements. RAGT seems to have similar effects to TOP over 10 weeks.		

6MWT = 6-minute walk test; ATE = anti-gravity treadmill exercise; COPM = Canadian occupational performance measure; CP = cerebral palsy; GMFCS = Gross Motor Function Classification System; GMFM = Gross Motor Function Measure; NRS = non-randomized study; PBWSTE = Partial body weight-supported treadmill exercise; RAGT = robotic-assisted gait training; RATE = Robotic-assisted treadmill exercise; RCT = randomized controlled trial; TOP = task-oriented physiotherapy; WeeFIM = pediatric Functional Independence Measure.



### References

#### Health Technology Assessments

No literature identified.

#### Systematic Reviews

No literature identified.

#### **Randomized Controlled Trials**

#### Comparator - Alternative Therapies

- 1. Ammann-Reiffer C, Bastiaenen CHG, Meyer-Heim AD, van Hedel HJA. Lessons learned from conducting a pragmatic, randomized, crossover trial on robot-assisted gait training in children with cerebral palsy (PeLoGAIT). J Pediatr Rehabil Med. 2020; 13(2): 137-148. PubMed
- 2. Jin LH, Yang SS, Choi JY, Sohn MK. The effect of robot-assisted gait training on locomotor function and functional capability for daily activities in children with cerebral palsy: a single-blinded, randomized cross-over trial. *Brain Sci.* Oct 30 2020; 10(11): 30. PubMed
- 3. Kawasaki S, Ohata K, Yoshida T, Yokoyama A, Yamada S. Gait improvements by assisting hip movements with the robot in children with cerebral palsy: a pilot randomized controlled trial. J Neuroengineering Rehabil. 2020; 17(1): 87. PubMed
- 4. Aras B, Yasar E, Kesikburun S, Turker D, Tok F, Yilmaz B. Comparison of the effectiveness of partial body weight-supported treadmill exercises, robotic-assisted treadmill exercises, and anti-gravity treadmill exercises in spastic cerebral palsy. *Turk J Phys Med Rehabil.* 2019 Jun;65(4):361-370. PubMed
- 5. Wu M, Kim J, Arora P, Gaebler-Spira DJ, Zhang Y. Effects of the integration of dynamic weight shifting training into treadmill training on walking function of children with cerebral palsy: a randomized controlled study. Am J Phys Med Rehabil. Nov 2017; 96(11): 765-772. PubMed
- 6. Wu M, Kim J, Gaebler-Spira DJ, Schmit BD, Arora P. Robotic resistance treadmill training improves locomotor function in children with cerebral palsy: a randomized controlled pilot study. Arch Phys Med Rehabil. 2017; 98(11): 2126-2133. PubMed

#### Between-Group Comparison Not Presented in Abstract

- 7. Wallard L, Dietrich G, Kerlirzin Y, Bredin J. Effect of robotic-assisted gait rehabilitation on dynamic equilibrium control in the gait of children with cerebral palsy. *Gait Posture*. 2018; 60: 55-60. PubMed
- 8. Wallard L, Dietrich G, Kerlirzin Y, Bredin J. Robotic-assisted gait training improves walking abilities in diplegic children with cerebral palsy. *Europ J Paediatr Neurol*. May 2017; 21(3): 557-564. PubMed

#### Non-Randomized Studies

#### Comparator – Alternative Therapies

9. Peri E, Turconi AC, Biffi E, et al. Effects of dose and duration of Robot-Assisted Gait Training on walking ability of children affected by cerebral palsy. *Technol Health Care*. 2017; 25(4): 671-681. PubMed

#### **Economic Evaluations**

No literature identified.

## **Appendix 1: References of Potential Interest**

#### **Previous CADTH Reports**

10. Upper and lower extremity robotic therapy and assistive devices for pediatric patients with complex developmental disabilities, brain injury, or complex pain disorders: clinical effectiveness, cost-effectiveness, and guidelines. (Rapid response report: summary of abstracts). Ottawa: CADTH, 2015. <a href="https://www.cadth.ca/sites/default/files/pdf/htis/june-2015/RB0875%20Upper%20and%20Lower%20Extremity%20Robotics%20Final.pdf">https://www.cadth.ca/sites/default/files/pdf/htis/june-2015/RB0875%20Upper%20and%20Lower%20Extremity%20Robotics%20Final.pdf</a> Accessed 2022 Jun 15.

#### Systematic Reviews

#### Mixed Population - Not Specific to Children

- 11. Conner BC, Remec NM, Lerner ZF. Is robotic gait training effective for individuals with cerebral palsy? A systematic review and meta-analysis of randomized controlled trials. *Clin Rehabil.* Jul 2022; 36(7): 873-882. PubMed
- 12. Bunge LR, Davidson AJ, Helmore BR, et al. Effectiveness of powered exoskeleton use on gait in individuals with cerebral palsy: a systematic review. *PLoS One*. 2021; 16(5): e0252193. PubMed
- 13. Selph SS, Skelly AC, Wasson N, et al. Physical activity and the health of wheelchair users: a systematic review in multiple sclerosis, cerebral palsy, and spinal cord injury. Arch Phys Med Rehabil. 12 2021; 102(12): 2464-2481.e33. PubMed

#### Unclear Intervention and Participant Age Group

- 14. Conner BC, Remec NM, Michaels CM, Wallace CW, Andrisevic E, Lerner ZF. Relationship between ankle function and walking ability for children and young adults with cerebral palsy: A systematic review of deficits and targeted interventions. *Gait Posture*. 2022; 91: 165-178. PubMed
- 15. Chiu HC, Ada L, Bania TA. Mechanically assisted walking training for walking, participation, and quality of life in children with cerebral palsy. Cochrane Database Syst Rev. 2020 Nov 18;11(11):CD013114. doi: 10.1002/14651858.CD013114.pub2. PubMed: ; https://www.cochranelibrary.com/cdsr/doi/10.1002/14651858.CD013114 .pub2/full Accessed 2022 Jun 16.

#### Unclear Comparator

- 16. Llamas-Ramos R, Sanchez-Gonzalez JL, Llamas-Ramos I. Robotic systems for the physiotherapy treatment of children with cerebral palsy: a systematic review. Int J Environ Res Public Health. 2022; 19(9): 22. PubMed
- 17. Cumplido C, Delgado E, Ramos J, et al. Gait-assisted exoskeletons for children with cerebral palsy or spinal muscular atrophy: a systematic review. *NeuroRehabilitation*. 2021; 49(3): 333-348. <u>PubMed</u>
- 18. Valè N, Gandolfi M, Vignoli L, et al. Electromechanical and robotic devices for gait and balance rehabilitation of children with neurological disability: a systematic review. Applied Sciences (Switzerland). 2021; 11(24).
- 19. Morsy SE, Salem EE, Elhadidy EI. Effect of automated locomotor training on gait in children with cerebral palsy: a systematic review and meta-analysis. World J Sport Sci. 2019. 14(2): 34-41. https://idosi.org/wjss/14(2)19/2.pdf Accessed 2022 Jun 15.

#### Before-After Comparison

20. Lefmann S, Russo R, Hillier S. The effectiveness of robotic-assisted gait training for paediatric gait disorders: systematic review. *J Neuroengineering Rehabil*. 2017; 14(1): 1. PubMed

#### Before-After Comparison and Mixed Population – Not Specific to Children

- 21. Volpini M, Aquino M, Holanda AC, Emygdio E, Polese J. Clinical effects of assisted robotic gait training in walking distance, speed, and functionality are maintained over the long term in individuals with cerebral palsy: a systematic review and meta-analysis. *Disabil Rehabil*. 2021: 1-11. PubMed
- 22. Carvalho I, Pinto SM, Chagas DDV, Praxedes Dos Santos JL, de Sousa Oliveira T, Batista LA. Robotic gait training for individuals with cerebral palsy: a systematic review and meta-analysis. Arch Phys Med Rehabil. 2017; 98(11): 2332-2344. PubMed

#### Full Text Published in Spanish

23. Colomera JA, Nahuelhual P. Effectiveness of robotic assistance for gait training in children with cerebral palsy. a systematic review [article in Spanish]. Rehabilitacion. 2020; 54(2): 107-115. PubMed

#### Before-After Comparison and Full Text Published in Spanish

24. Cañadas Martinez LE, Montero Mendoza S. Effectiveness of automatic walking systems in children with cerebral palsy: A systematic review [article in Spanish]. *Fisioterapia*. 2020; 42(2): 75-84.

#### **Randomized Controlled Trials**

#### Mixed Population - Not Specific to Children

25. Klobucka S, Klobucky R, Kollar B. Effect of robot-assisted gait training on motor functions in adolescent and young adult patients with bilateral spastic cerebral palsy: a randomized controlled trial. *NeuroRehabilitation*. 2020; 47(4): 495-508. <u>PubMed</u>

#### Alternative Intervention – RAGT With Physiotherapy

- 26. Pool D, Valentine J, Taylor NF, Bear N, Elliott C. Locomotor and robotic assistive gait training for children with cerebral palsy. Dev Med Child Neurol. 2021; 63(3): 328-335. PubMed
- 27. Yaśar B, Atlcl E, Razaei DA, SaldIran TÇ. Effectiveness of robot-assisted gait training on functional skills in children with cerebral palsy. J Pediatr Neurol. 2021.
- 28. Mahgoub MSE, Amin WW, Zahran SS. Effect of locomotor training with a roboticgait orthosis (Lokomat) in spasticity modulation of spastic hemiplegic children: a randomized controlled trial. *Fizjoterapia Polska*. 2020; 20(4): 94-101.

#### Alternative Intervention - RAGT With Virtual Reality

29. Fu WS, Wu BA, Song YC, Qu CH, Zhao JF. Virtual reality combined with robot-assisted gait training to improve walking ability of children with cerebral palsy: a randomized controlled trial. Technol Health Care. May 23 2022. PubMed

#### Alternative Intervention – Motor-Assisted Cycle

 Damiano DL, Stanley CJ, Ohlrich L, Alter KE. Task-specific and functional effects of speed-focused elliptical or motor-assisted cycle training in children with bilateral cerebral palsy: randomized clinical trial. Neurorehabil Neural Repair. Aug 2017; 31(8): 736-745. PubMed

#### Non-Randomized Studies

#### Mixed Population – Not Specific to Cerebral Palsy

31. Hong J, Lee J, Choi T, et al. Feasibility of overground gait training using a joint-torque-assisting wearable exoskeletal robot in children with static brain injury. Sensors (Basel). May 19 2022; 22(10): 19. PubMed

#### Alternative Intervention – Robotic-Assisted Gait Therapy With Physiotherapy

- 32. Abidin N, Unlu Akyuz E, Cankurtaran D, Karaahmet OZ, Tezel N. The effect of robotic rehabilitation on posture and trunk control in non-ambulatory cerebral palsy. Assist Technol. May 05 2022; 1-7. PubMed
- 33. Yazici M, Livanelioglu A, Gucuyener K, Tekin L, Sumer E, Yakut Y. Effects of robotic rehabilitation on walking and balance in pediatric patients with hemiparetic cerebral palsy. Gait Posture. 2019; 70: 397-402. PubMed

#### Alternative Intervention – RAGT With Botulinum Therapy

34. Balgayeva M, Bulekbayeva S. Effectiveness of the combined use of robotic kinesiotherapy and botulinum therapy in the complex rehabilitation of children with cerebral palsy. Asian J Pharm Clin Res. 2018; 11(9): 360-364.

#### Single-Arm Studies

- 35. Klobucka S, Klobucky R, Kollar B. The effect of patient-specific factors on responsiveness to robot-assisted gait training in patients with bilateral spastic cerebral palsy. *NeuroRehabilitation*. 2021; 49(3): 375-389. PubMed
- 36. Manikowska F, Brazevic S, Krzyzanska A, Jozwiak M. Effects of robot-assisted therapy on gait parameters in pediatric patients with spastic cerebral palsy. Front Neurol. 2021; 12: 724009. PubMed
- 37. Manikowska F, Krzyzanska A, Chmara P, Chen BP, Jozwiak M. Baseline gross motor function affects the outcome of robot-assisted therapy in ambulatory individuals with spastic cerebral palsy. *Brain sci.* Nov 26 2021; 11(12): 26. PubMed
- 38. Petrarca M, Frascarelli F, Carniel S, et al. Robotic-assisted locomotor treadmill therapy does not change gait pattern in children with cerebral palsy. Int J Rehabil Res. 2021; 44(1): 69-76. PubMed
- Zarkovic D, Sorfova M, Tufano JJ, et al. Gait changes following robot-assisted gait training in children with cerebral palsy. *Physiol Res.* 2021; 70(S3): S397-S408. <u>PubMed</u>
- Cherni Y, Ballaz L, Lemaire J, Dal Maso F, Begon M. Effect of low dose robotic-gait training on walking capacity in children and adolescents with cerebral palsy. Neurophysiol Clin. Nov 2020; 50(6): 507-519. <u>PubMed</u>
- 41. Nakagawa S, Mutsuzaki H, Mataki Y, et al. Safety and immediate effects of Hybrid Assistive Limb in children with cerebral palsy: a pilot study. *Brain Dev.* Feb 2020; 42(2): 140-147. PubMed
- 42. Sucuoglu H. Effects of robot-assisted gait training alongside conventional therapy on the development of walking in children with cerebral palsy. J Pediatr Rehabil Med. 2020; 13(2): 127-135. PubMed
- 43. Zarkovic D, Sorfova M, Tufano JJ, et al. Effect of robot-assisted gait training on selective voluntary motor control in ambulatory children with cerebral palsy. Indian Pediatr. 10 15 2020; 57(10): 964-966. PubMed This pilot study investigated the efficacy of a 4-week robot-assisted gait training in 12 children with spastic diparesis. Short-term results and a 3-month follow-up showed statistically significantly increased selective motor control, walking farther distances, gross motor score, and decreased joint contractures.
- 44. Aycardi LF, Cifuentes CA, Munera M, et al. Evaluation of biomechanical gait parameters of patients with cerebral palsy at three different levels of gait assistance using the CPWalker. J Neuroengineering Rehabil. 2019; 16(1): 15. PubMed
- 45. Digiacomo F, Tamburin S, Tebaldi S, et al. Improvement of motor performance in children with cerebral palsy treated with exoskeleton robotic training: A retrospective explorative analysis. Restor Neurol Neurosci. 2019; 37(3): 239-244. PubMed



- 46. Lee SJ, Jin D, Kang SH, Gaebler-Spira D, Zhang LQ. Combined ankle/knee stretching and pivoting stepping training for children with cerebral palsy. IEEE Trans Neural Syst Rehabil Eng. 2019; 27(9): 1743-1752. PubMed
- 47. Weinberger R, Warken B, Konig H, et al. Three by three weeks of robot-enhanced repetitive gait therapy within a global rehabilitation plan improves gross motor development in children with cerebral palsy a retrospective cohort study. *Europ J Paediatr Neurol*. Jul 2019; 23(4): 581-588. PubMed
- 48. Bayon C, Martin-Lorenzo T, Moral-Saiz B, et al. A robot-based gait training therapy for pediatric population with cerebral palsy: goal setting, proposal and preliminary clinical implementation. J Neuroengineering Rehabil. 2018; 15(1): 69. PubMed
- 49. Kang J, Martelli D, Vashista V, Martinez-Hernandez I, Kim H, Agrawal SK. Robot-driven downward pelvic pull to improve crouch gait in children with cerebral palsy. Science Robotics. 2017;2(8). PubMed
- 50. Bayon C, Lerma S, Ramirez O, et al. Locomotor training through a novel robotic platform for gait rehabilitation in pediatric population: short report. *J Neuroengineering* Rehabil. 2016; 13(1): 98. PubMed
- 51. Wu M, Landry JM. Toward flexible assistance for locomotor training: design and clinical testing of a cable-driven robot for stroke, spinal cord injury, and cerebral palsy. In: Reinkensmeyer D, Dietz V, eds. Neurorehabilitation Technology 2nd ed. Cham (Switzerland): Springer; 2016.

#### Single-Arm Studies and Mixed Population – Not Specific to Children

- 52. Fang Y, Orekhov G, Lerner ZF. Adaptive ankle exoskeleton gait training demonstrates acute neuromuscular and spatiotemporal benefits for individuals with cerebral palsy: a pilot study. *Gait Posture*. 2022 Jun;95:256-263. PubMed
- 53. Fang Y, Lerner ZF. Feasibility of augmenting ankle exoskeleton walking performance with step length biofeedback in individuals with cerebral palsy. *IEEE Trans Neural* Syst Rehabil Eng. 2021; 29(): 442-449. PubMed
- 54. Klobucka S, Klobucky R, Kollar B. The effect of patient-specific factors on responsiveness to robot-assisted gait training in patients with bilateral spastic cerebral palsy. *NeuroRehabilitation*. 2021; 49(3): 375-389. PubMed
- 55. Mataki Y, Mutsuzaki H, Kamada H, et al. Effect of the hybrid assistive limb on the gait pattern for cerebral palsy. Medicina (Kaunas). Dec 07 2020; 56(12): 07. PubMed
- 56. Orekhov G, Fang Y, Luque J, Lerner ZF. Ankle exoskeleton assistance can improve over-ground walking economy in individuals with cerebral palsy. *IEEE Trans Neural* Syst Rehabil Eng. 2020 02;28(2):461-467. PubMed
- 57. Lerner ZF, Harvey TA, Lawson JL. A battery-powered ankle exoskeleton improves gait mechanics in a feasibility study of individuals with cerebral palsy. Ann Biomed Eng. 2019 Jun;47(6):1345-1356. PubMed
- 58. Ueno T, Watanabe H, Kawamoto H, et al. Feasibility and safety of Robot Suit HAL treatment for adolescents and adults with cerebral palsy. *J Clin Neurosci*. Oct 2019; 68(): 101-104. PubMed
- 59. Lerner ZF, Gasparri GM, Bair MO, et al. An untethered ankle exoskeleton improves walking economy in a pilot study of individuals with cerebral palsy. *IEEE Trans Neural* Syst Rehabil Eng. 2018 10;26(10):1985-1993. PubMed
- 60. Matsuda M, Iwasaki N, Mataki Y, et al. Robot-assisted training using Hybrid Assistive Limb R for cerebral palsy. Brain Dev. 2018; 40(8): 642-648. PubMed
- 61. Matsuda M, Mataki Y, Mutsuzaki H, et al. Immediate effects of a single session of robot-assisted gait training using Hybrid Assistive Limb (HAL) for cerebral palsy. J Phys Ther Sci. 2018; 30(2): 207-212. PubMed
- 62. Takahashi K, Mutsuzaki H, Mataki Y, et al. Safety and immediate effect of gait training using a Hybrid Assistive Limb in patients with cerebral palsy. J Phys Ther Sci. 2018; 30(8): 1009-1013. PubMed
- 63. Lerner ZF, Damiano DL, Bulea TC. The Effects of Exoskeleton Assisted Knee Extension on Lower-Extremity Gait Kinematics, Kinetics, and Muscle Activity in Children with Cerebral Palsy. Sci. 2017 10 18;7(1):13512. PubMed
- 64. van Hedel HJ, Meyer-Heim A, Rusch-Bohtz C. Robot-assisted gait training might be beneficial for more severely affected children with cerebral palsy. *Dev Neurorehabil*. 2016; 19(6): 410-415. PubMed

#### Single-Arm Studies and Alternative Intervention – RAGT With Physical Therapy

65. Beretta E, Storm FA, Strazzer S, et al. effect of robot-assisted gait training in a large population of children with motor impairment due to cerebral palsy or acquired brain injury. Arch Phys Med Rehabil. 2020; 101(1): 106-112. PubMed

#### Single-Arm Studies and Alternative Intervention – Robotic Resistance Training

- 66. Conner BC, Schwartz MH, Lerner ZF. Pilot evaluation of changes in motor control after wearable robotic resistance training in children with cerebral palsy. *J Biomech.* 2021;126:110601. PubMed
- 67. Conner BC, Remec NM, Orum EK, Frank EM, Lerner ZF. Wearable adaptive resistance training improves ankle strength, walking efficiency and mobility in cerebral palsy: a pilot clinical trial. *IEEE Open J Eng Med Biol.* 2020;1:282-289. PubMed

#### Single-Arm Studies and Unclear Intervention

68. Lerner ZF, Damiano DL, Bulea TC. A lower-extremity exoskeleton improves knee extension in children with crouch gait from cerebral palsy. Sci Transl Med. 2017; 9(404): 23. PubMed

69. Wu M, Kim J, Arora P, Gaebler-Spira DJ, Zhang Y. Kinematic and EMG responses to pelvis and leg assistance force during treadmill walking in children with cerebral palsy. *Neural Plast*. 2016;2016:5020348. PubMed

#### **Review Articles**

#### Meta-Analyses - Unclear if Based on Systematic Review

70. Olmos-Gomez R, Gomez-Conesa A, Calvo-Munoz I, Lopez-Lopez JA. Effects of robotic-assisted gait training in children and adolescents with cerebral palsy: a network meta-analysis. J Clin Med. 2021; 10(21): 24. PubMed

#### Additional References

#### Conference Abstracts or Proceedings

- 71. Cherni Y, Ballaz L, Girarin-Vignola G, Begon M. P 175 Robotic-assisted locomotion training improves walking abilities in children with bilateral cerebral palsy [conference abstract]. Gait Posture. 2018; 65: 530-531.
- 72. Kerr L. Robotic-assisted backward gait result in functional mobility changes in two children diagnosed with cerebral palsy [conference abstract]. Arch Phys Med Rehabil. 2017;98(10):e106-e106.

The purpose is to investigate the utilization of robotic-assisted backward walking to provide high-level repetitions in task specific practice for 2 children diagnosed with cerebral palsy (CP). Would this training program of intensive and repetitive backward walking result in positive functional mobility changes in children with CP?

73. Peri E, Biffi E, Maghini C, et al. An ecological evaluation of the metabolic benefits due to robot-assisted gait training. Annu Int Conf IEEE Eng Med Biol Soc. Aug 2015; 2015: 3590-3. PubMed