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Metabolically efficient walking assistance using optimized timed forces at the waist

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1	Title: Pushing the boundaries of efficient walking assistance,
2	using timed forces at the center of mass
3	
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1 **Running head:** Assisting gait via center-of-mass forces

2

3 Abstract:

The metabolic rate of walking can be reduced by applying a constant forward force at the 4 center of mass (COM). The optimal constant force magnitude minimizes propulsion ground 5 6 reaction force at the expense of increased braking, which raises the question of whether selectively assisting propulsion could augment benefits. We show that it is possible to reduce the metabolic 7 rate by 48% with a greater efficiency ratio of metabolic cost reduction per unit of net aiding work 8 9 compared to other assistive robots using an optimized sinusoidal force profile. A model explains that the optimal profile accelerates the COM into the inverted pendulum movement during single 10 support. Contrary to the hypothesis, the optimal force timing did not entirely coincide with 11 propulsion. Wearable robots often use control algorithms that mimic biological kinetics. Such 12 bioinspired actuation is not necessarily optimal for reducing metabolic rate. 13

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15 **One sentence summary:**

16 Not mimicking biological ground reaction force but assisting center-of-mass acceleration

17 reduces the metabolic cost of walking by half.

1 Main text:

2 INTRODUCTION

Robots often incorporate anthropomorphic designs (1). In the subfield of rehabilitation 3 robotics, different groups develop biologically inspired exoskeletons that assist with moments at 4 the biological joints (2). While exoskeletons have appropriate applications, simple pendulum 5 6 models suggest that timed linear forces acting through the center of mass (COM) can actuate walking more efficiently than moments (3). Gottschall and Kram conducted foundational research 7 on the effects of linear forward aiding forces with constant magnitude at the COM and show that 8 9 the metabolic rate of walking in healthy adults can be reduced by up to 47% with a long elastic tether (4). They report that optimal aiding forces minimize propulsion at the expense of increased 10 braking ground reaction forces (GRFs) and call for further research on devices that could 11 specifically assist propulsion, perhaps without impeding braking. 12

There is increased interest in devices that can apply such non-constant force profiles during 13 specific phases of the gait cycle (5, 6). Bhat et al. (5) use stiff tethers to elicit cyclic force profiles 14 from the back-and-forth movements on a treadmill. They find smaller metabolic rate reductions 15 than those for constant forces (4, 7), which suggests that their force profiles are suboptimal. Penke 16 17 et al. (6) use a pulley system that connects the COM to one of the ankles to apply cyclic force profiles. This system reduces the metabolic rate of individuals poststroke by 12%, demonstrating 18 19 potential clinical impact. In healthy participants, they find relatively smaller reductions than those 20 for constant forces, which suggests that there is room for further improvements.

21 Certain clinical populations have large increases in metabolic rate during walking [e.g., 22 200 to 300% in cerebral palsy (8)]. Exoskeletons can currently reduce the metabolic rate by up to 23 one-fourth in healthy individuals (9, 10). Even if, potentially, similar reductions in metabolic rate

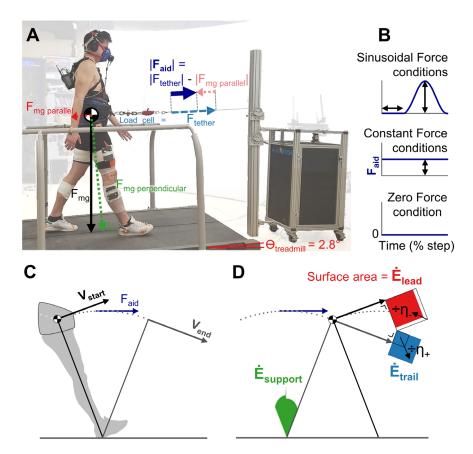
versus unassisted walking could be obtained in clinical populations, this might be insufficient to 1 bring the metabolic cost to normal levels and allow for sustained walking practice in certain 2 patients who have a metabolic cost that is higher compared to healthy individuals. In the current 3 study, we investigate different ways of assisting treadmill walking using timed forward forces at 4 the COM instead of exoskeletons. Simple pendulum models (3) and studies with exoskeletons (9-5 6 14) show that the parameters that define the shape of the actuation profile during the walking cycle can influence the metabolic rate. We aimed to investigate the effects of timing and magnitude for 7 non-constant force profiles at the COM in healthy individuals. This initial work is intended to 8 9 assess novel assistive strategies at the COM using a robotic tether system with the goal of reducing the metabolic rate of walking in healthy adults. Depending on the outcome, this could be a 10 preliminary step towards potentially testing such a system in clinical populations. Gottschall and 11 Kram (4) report that the metabolically optimal magnitude for constant forces minimizes 12 propulsion. Therefore, we expected a similar relationship between propulsion reduction and 13 14 metabolic rate reduction, and we hypothesized that non-constant force profiles in which the peak magnitude coincides with propulsion would optimally reduce propulsion and metabolic rate. 15

We conducted experiments with a cable robot that applied forces to a waist belt in ten 16 17 healthy participants (Fig. 1A and movie S1) (15, 16). A force control algorithm was used to apply 32 conditions with sinusoidal force profiles with desired magnitudes, timings, and durations 18 19 (Sinusoidal Force conditions; Fig. 1B) (17) as well as four conditions where the force had a 20 constant magnitude throughout the step cycle (Constant Force conditions). Since the tether acts symmetrically on both legs, we expressed timings in percent of the step cycle instead of percent 21 22 of the stride cycle. We cannot differentiate between actuating during the first or second double 23 support phase (stride-based terminology), but we can differentiate between actuating during 1 double support and single support (step-based terminology). We calculated the net aiding work

2 rate by multiplying the net aiding force by the COM velocity, and we estimated the metabolic rate

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using respiratory measurements.



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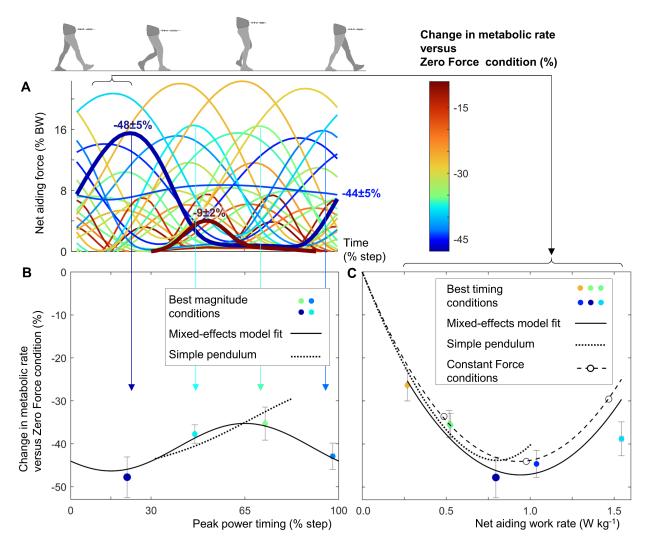
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Fig. 1. Methods. (**A**) Setup. A cable robot (HuMoTech, Pittsburgh, PA, USA) applied force profiles to a waist belt. To avoid the tether going slack during portions of the step cycle, a constant backward force was simulated by inclining the treadmill (*17*, *18*) by a 2.8° angle. The reported aiding forces are the forces measured with a load cell minus the parallel component of gravity. (**B**) Conditions. We used the cable robot to apply Sinusoidal Force conditions, Constant Force conditions, and a Zero Force condition. (**C**) Simple pendulum model. We modeled an inverted pendulum (*3*, *19*) with the mean mass and leg length of our participants. This model was used to simulate different aiding force timings and magnitudes. For each condition, we identified the initial COM velocity required to match the experimental step time and the velocity at the end of single support. Based on COM velocity, we estimated the metabolic rate (visualized as colored surfaces) of the trailing and leading legs during the step-to-step transition and the support leg during single support (*15*). The metabolic rate of the step-to-step transition was estimated by dividing the redirection work rate that the legs have to produce (*3*) by the assumed efficiencies (*20*).

1 **RESULTS**

2	The greatest reduction in metabolic rate compared to the condition with a net aiding force
3	of zero (Zero Force condition) was $47.8 \pm 4.7\%$ (all results are mean \pm standard error of the mean
4	(SEM) except when stated otherwise, $P < 0.001$, paired t test with Holm-Šidák correction, Fig.
5	2A). This reduction occurred in the Sinusoidal Force condition with a peak force of $15.0 \pm 0.5\%$
6	body weight (BW), a duration of $64.1 \pm 1.6\%$ of the step, and a peak timing during the double
7	support phase at 21.1 \pm 0.3% of the step. The smallest reduction was 9.1 \pm 2.0% (P = 0.006) in the
8	condition with a peak force of 4.7 \pm 0.1% BW, a duration of 27.0 \pm 0.4%, and a peak timing of
9	$50.9 \pm 0.4\%$ of the step.
10	A mixed-effects model analysis of the effects of the different actuation profile parameters
11	showed a significant effect of peak timing on metabolic rate ($P_{\text{Ppeak} \cdot \sin(\text{tpeak})} < 0.001$) with an
12	optimum at the middle of the double support phase (15.1 \pm 0.1% of step time, Fig. 2B) (15). As
13	expected, different conditions reduced metabolic rate by different amounts, but, surprisingly, there
14	were no conditions that increased the metabolic rate. Humans appear to be less sensitive to the
15	timing of assistance at the COM than that of joint-level assistance, as certain conditions in studies
16	with exoskeletons increase the metabolic rate $(11, 14)$.
17	We found a U-shaped effect of net aiding work rate (P_{Waid} and $P_{\text{Waid}}^2 < 0.001$) with an
18	optimum at 0.958 \pm 0.043 W kg ⁻¹ (Fig. 2C). There was no significant effect of net aiding power
19	duration. The best Sinusoidal Force condition produced a greater reduction in metabolic rate per
20	unit net aiding work rate than the best Constant Force condition (ratios were 2.06 ± 0.21 and 1.54
21	\pm 0.14, respectively, <i>P</i> = 0.032), but there was no significant difference in the metabolic rate. We
22	also used the fits of the mixed-effects model analysis equation to each participant to improve the
23	estimates of the individual optima. This analysis suggests that there was still a small metabolic rate

1	benefit of $3.9 \pm 1.5\%$ of the optima of the Sinusoidal Force conditions compared to the optima of
2	the Constant Force conditions ($P = 0.030$). The highest reduction in the metabolic rate of the
3	Constant Force conditions (44.0 \pm 4.4 %) and the Constant Force level that minimized the
4	metabolic rate (7.9 \pm 0.03 % BW) were similar to other studies [47% with a net aiding force of
5	10% BW (4), 35% with a net aiding force of 8% BW (7) and 34% with a net aiding force of 8.4%
6	BW (6)]. Although the results show benefits of optimally timed forces, the effects of timing were
7	relatively small compared to the effect of aiding work rate. This could be considered as a limitation
8	given the fact that applying different timings is more challenging than applying different constant
9	magnitudes, which can be done with a passive tether system.



2 Fig. 2. Metabolic rate. (A) Net aiding forces. Lines represent means of 10 participants for Sinusoidal Force conditions 3 and the best Constant Force condition. The color scale indicates metabolic rate reduction. Thick lines indicate highest and lowest reductions. We applied the same profile during left and right steps; therefore, we plotted profiles versus 4 5 step instead of stride time. (B) Effect of peak timing. Dots and error bars represent means and SEMs of 10 participants 6 for conditions with different timings but approximately the same duration and net aiding work rate as the condition with 7 the highest reduction. Solid line represents the mixed-effects model fitted on all Sinusoidal Force conditions (-3.13 Waid + $1.67 \cdot \dot{W}_{aid^2}$ + $0.09 \cdot P_{peak} \cdot sin(\frac{t_{p_{peak}} + 60.16}{100} \cdot 2\pi)$; all variables are in W kg⁻¹ and %; *P*-values < 0.001; adjusted R² = 8 9 95.3%), evaluated at the mean magnitude of dots. Dotted line represents the estimation from the simple pendulum 10 simulated over the feasible range. (C) Effect of net aiding work rate. Dots are conditions with different net aiding work 11 rates within a timing range of ± 6% from the optimum (15.1%). Circles and the dashed line represent Constant Force 12 conditions. Independent variables of (B) and (C) were selected from different candidate metrics (15).

DISCUSSION

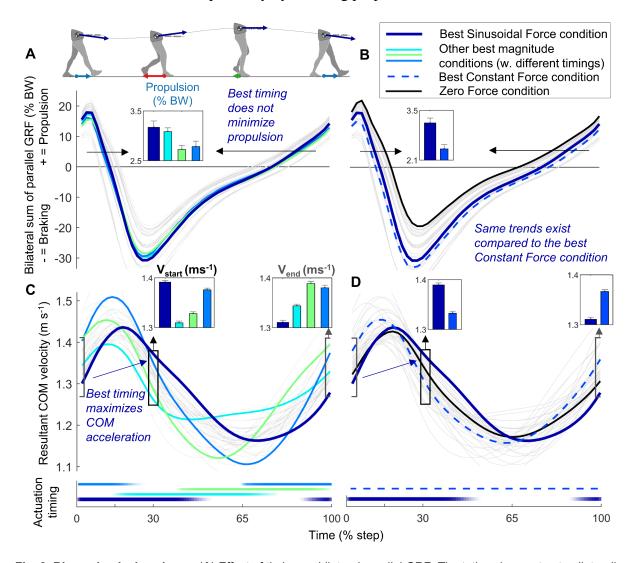
To understand the potential underlying mechanisms of the effects seen during 2 experimentation, we conducted an inverted pendulum simulation (3, 19) and analyzed relevant 3 biomechanical measurements (15). The simple pendulum model (Fig. 1C) reproduced the changes 4 observed in metabolic rates with increasing net aiding work rate but reached its optimum at a lower 5 work rate (0.796 W kg⁻¹; Fig. 2C). The model also closely reproduced the effect of assistance 6 timing over the feasible simulation range. The model could not simulate timings at the beginning 7 of the step because double support is instantaneous. Additionally, the model could not simulate 8 9 timings at the end of the step because the centripetal component from gravity minus the component from the tether force that pulls the COM away from the ground contact point (due to the alignment 10 between the tether and the leg) was not sufficient to sustain the radial acceleration (Fig. 2B). We 11 plotted the results from the model over the time range that corresponds to the single support phase 12 from the human experimental data (immediately after toe-off at 30% of the step cycle until the end 13 of the feasible simulation range). This simple model explained that aiding forces early in the step 14 cycle were optimal because they help accelerate the COM, thereby reducing the positive leg work 15 required to reach the apex of the inverted pendulum movement. Aiding force after midstance was 16 17 suboptimal because the COM spontaneously accelerates due to gravity.

We found significant effects of net aiding work rate and peak timing on the propulsion part of the bipedal sum of the GRF (all *P*-values < 0.001, mixed-effects model analysis; Fig. 3A, table S4). As expected, the peak timing that resulted in the greatest propulsion reduction fell inside the propulsion phase (mean \pm SEM of the peak timing that minimized propulsion was 74.7 \pm 0.2% of step, mixed-effects model analysis). However, the influence of timing on propulsion GRF was small (range between minimum and maximum propulsion GRFs of conditions with the same

1	magnitude in Fig. 3A is $0.6 \pm 0.1\%$ BW). Even force profiles that fell entirely during the braking
2	phase also reduced propulsion by up to $36.5 \pm 1.6\%$ at the expense of increased braking GRF and
3	reduced the metabolic rate by up to $37.7 \pm 2.1\%$ compared to the Zero Force condition (<i>P</i> -values
4	< 0.001, paired t tests with Holm-Šidák correction). Contrary to our hypothesis, the peak timing
5	that minimized metabolic rate based on the mixed-effects model analysis (15.1% of step time)
6	aligned with the transition from propulsion to braking (the first zero crossing occurred at 15.4 \pm
7	0.6% of step time, mean \pm SEM from all conditions; Fig. 3A) instead of aligning with the middle
8	of the propulsion phase. The simple pendulum model proved to better explain our results than the
9	GRF hypothesis. We found large effects of peak timing on the resultant COM velocity at the
10	beginning and end of single support (<i>P</i> -values < 0.001 ; Fig. 3B and table S4). The metabolically
11	optimal timing was close to the timing that maximized the COM velocity at the beginning of single
12	support (the peak timing that maximized initial COM velocity was $10.0 \pm 0.6\%$ of step time,
13	mixed-effects model analysis). The peak timings that minimized leading leg negative work and
14	trailing leg positive work all occurred in the first half of single support, as predicted by the model
15	(the peak actuation timings that minimized collision and push-off were 32.6 ± 1.3 and $54.0 \pm 0.5\%$
16	of step, respectively, the middle of single support was at 65%, all <i>P</i> -values for timing < 0.001 ;
17	table S4). These results confirmed the model prediction that assisting COM acceleration at the
18	beginning of the step reduces the required leg work (Fig. 4). The Sinusoidal Force Condition that
19	minimized the metabolic rate required $15.1 \pm 0.6\%$ greater propulsion than the best Constant Force
20	condition ($P < 0.001$, Fig. 3B), but increased COM acceleration compared to the best Constant
21	Force condition (4.6 \pm 0.2% greater COM velocity at the beginning of the single support and 6.1
22	\pm 0.2% smaller COM velocity at the end of the single support, all P-values < 0.001, Fig. 3C). This

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Constant Force condition, not primarily by reducing propulsion GRF.



shows that the Sinusoidal Force conditions reduce the metabolic rate in a different way than the

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Fig. 3. Biomechanical analyses. (**A**) Effect of timing on bilateral parallel GRF. The tether does not act unilaterally on one leg; therefore, we chose to analyze the bilateral GRF instead of the unilateral GRF. Colored lines represent conditions with similar actuation magnitudes and approximately the same net aiding work rate as the best condition from Fig. 2B but with different timings. The GRFs of the colored conditions appear offset because of the net aiding force that is not shown here. Gray lines represent all other conditions. Bar plots represent the average propulsion GRF averaged over the entire step duration. The thick line marks the condition that showed the highest metabolic rate reduction. (**B**) Comparison of bilateral parallel GRF between the condition with the highest reduction in metabolic rate (thick blue line), best Constant Force condition (dashed blue line), and Zero Force condition (black line). (**C**) Effect of timing on the resultant COM velocity. Bar plots represent the magnitude of the COM velocity vector at the beginning

and end of single support. Horizontal lines represent the actuation periods. The condition with the highest reduction in metabolic rate (dark blue bar) has the lowest COM velocity at the end of single support, the highest COM velocity at the beginning of single support, and thus the highest acceleration during double support. (D) Comparison of COM velocity between the condition with the highest reduction in metabolic rate (thick blue line), best Constant Force condition (dashed blue line), and Zero Force condition (black line).

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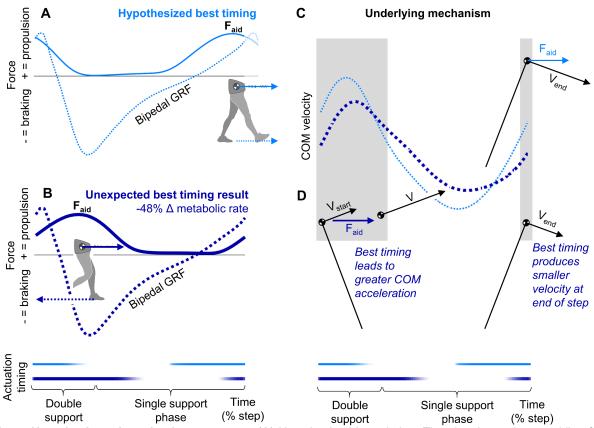
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7 8 Fig. 4. Hypothesis and mechanism summary. (A) Hypothesized best timing. The plot shows the net aiding force (solid line) and bipedal sum of the parallel GRF (dashed line) in the Sinusoidal Force condition where the net aiding force matches closest to the bipedal propulsion timing, plotted versus step time. (B) Best timing result. Condition with the highest reduction in metabolic rate. (C) COM velocity. Plot showing the COM velocity of conditions with the hypothesized best timing and the highest metabolic rate reduction. (D) Simple pendulum model. Stick figures show the higher COM acceleration at the beginning of the step and lower COM velocity at the end of the step in the condition with early peak timing compared to the condition with late peak timing. The pendulum model predicts that the higher 15 initial acceleration and lower final velocity will result in lower positive and negative leg work rates required to redirect 16 the COM.

1	Plotting reductions in metabolic rate versus net aiding work rate allows for comparing results with
2	devices that provide angular assistance such as exoskeletons (Fig. 5). This analysis confirmed that
3	applying constant aiding forces with tethers $(4, 7)$ has similar effects as downhill walking $(20, 21)$.
4	Non-constant force profiles obtained with passive tether systems that were tested in healthy adults
5	have peak timings around 85% of the step time $(5, 6)$, and this was close to the least optimal timing
6	according to our data (65.1%), which could explain their smaller observed reductions in healthy
7	participants. Reductions in metabolic rate in W kg ⁻¹ from the Sinusoidal Force conditions with the
8	best timing were greater than the best-in-class results for a wide range of devices, including
9	exoskeletons and prostheses (all <i>P</i> -values \leq 0.036, <i>t</i> tests) (4, 5, 16, 22, 23, 6, 7, 9–14), except for
10	one condition in a study (6) that uses constant aiding force in healthy participants ($P = 0.691$, one-
11	sample <i>t</i> test).

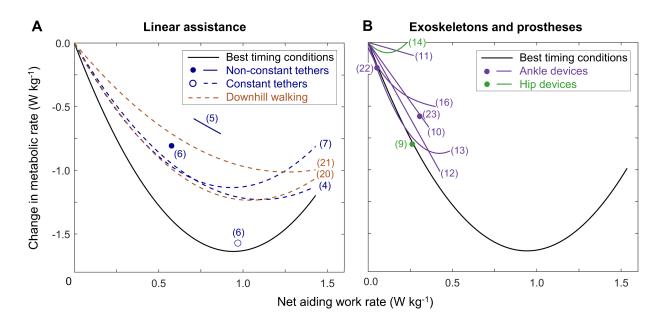




Fig. 5. Literature comparison. (A) Linear assistance. (B) Exoskeletons and prostheses. Reductions in metabolic rate compared to a Zero Force or no assistance condition versus net aiding work rate. The black line represents the effect of magnitude of conditions from the present study within 6% of the optimum peak timing (Fig. 2C). Colored lines represent trends from actuation magnitude parameter sweeps (4, 5, 7, 10–14, 16). Circles represent results from studies that do not use actuation magnitude sweep protocols (6, 9, 15, 22, 23).

The field of wearable robotics has evolved from sophisticated full-body exoskeletons (2) 1 toward simpler single-joint exoskeletons that first achieved metabolic rate reduction (9, 22, 23). 2 Our study suggests that a simpler strategy with linear forces applied at the COM can provide 3 further gains in stationary applications such as treadmill exercise therapy. Although robotic tethers 4 cannot assist with overground mobility similar to exoskeletons, the greater reductions could enable 5 6 treadmill exercise therapy applications. Clinical practice guidelines recommend that combining robotic assistance with higher intensity stepping is an area that remains to be investigated (24). 7 While the large effects of robotic waist tether assistance could allow for higher intensity stepping 8 9 training, it remains to be seen whether the effects observed in healthy participants transfer to patients. The effect of timing on the horizontal GRF was small, but we found large effects on the 10 COM velocity. Using their pulley mechanism that allowed for applying non-constant force 11 profiles, Penke et al. (6) found a 12% reduction in metabolic cost in individuals poststroke but no 12 reduction using constant forces. Thus, timing could potentially have a greater influence in 13 14 populations with non-constant walking velocity, such as individuals poststroke, but this remains to be tested. Evaluating the effect of timed assistance at the COM in clinical populations requires 15 an investigation of the targeted clinical population since there are different instances where the 16 17 effects of assistive devices in healthy participants did not translate to patients (25-28). In clinical populations or older adults, it is possible that the effects of non-constant forces on maintaining 18 19 balance eliminate any metabolic cost reduction benefit. 20 As an initial evaluation of the applicability of the waist tether in impaired gait, we

As an initial evaluation of the applicability of the waist tether in impaired gait, we conducted an experiment in a healthy participant in which we induced a non-constant COM velocity by unilaterally restricting plantarflexion. The results of this single-participant experiment suggest that it is possible to change the metabolic rate from +2 to -42% compared to the Zero Force

1	condition or +7 to -35% compared to a No Tether condition using different Sinusoidal Force
2	profiles after one habituation session (15) . We also conducted an experiment to evaluate the
3	feasibility of using the waist tether system as a method of exercise therapy in patients with limited
4	mobility due to a prevalent disease. Two patients with symptomatic peripheral artery disease
5	walked with the waist tether during two sessions, referred to as habituation and testing sessions.
6	During the testing session, we performed repeated comparisons between a Sinusoidal Force
7	condition and a No Tether condition, and we found reductions in metabolic rate of 12.0 and 9.7%
8	in one patient and 23.1 and 20.1% in the second patient (15). It is promising that these experiments
9	show that it is possible to reduce the metabolic cost of impaired walking using a robotic waist
10	tether. However, it is not yet possible to draw general conclusions from these small sample size
11	experiments. Further experiments are needed to evaluate if these results are reproducible in larger
12	samples of patients with impaired gait.
13	A number of wearable robots for clinical populations are designed to assist specifically

A number of wearable robots for clinical populations are designed to assist specifically during propulsion (29) or mimic biological kinematics or kinetics (30). We found that assisting during propulsion can reduce the metabolic rate but does not optimally reduce the metabolic rate, and even net aiding force profiles that occur entirely during braking reduce the metabolic rate. This finding indicates that bioinspired controls that mimic biological kinetics are not necessarily optimal for reducing the metabolic rate.

This work shows that it is possible to reduce the metabolic rate of walking by half by accelerating the COM. The observation that all force profiles with suboptimal timing still reduced the metabolic rate suggests that further reductions can be obtained with profiles that do not stay at zero during a part of the step cycle. Using a simple pendulum model, we predict that one could approach an 80% reduction by accelerating the COM during the first half of single support,

- 1 followed by decelerating the COM during the second half (with a backward force) (fig. S10) (15).
- 2 Such strategies could be implemented in cable robots for treadmill exercise therapy (31) or mobile
- 3 devices that assist via the trunk, such as motorized rollators (32).

2 **Participants**

MATERIALS AND METHODS

Ten healthy male participants (age: 28.0 ± 4.7 years, body mass: 83.2 ± 12.2 kg, height: 3 1.80 ± 0.05 m, leg length: 0.993 ± 0.036 m; mean \pm SD; table S5) took part in the study. 4 Participants were recruited using a convenience sampling strategy. We only included participants 5 6 without a previous history of musculoskeletal or neurological disorders. The sample size was selected to match the largest sample size from studies with waist tethers [n = 6 (5), n = 7 (6), n =7 10 (4, 7, 33)] and actuation timing and magnitude sweeps with wearable robots [n = 7 (12, 34), n]8 = 8 (11), n = 9 (22), n = 10 (13, 16, 35, 36)]. The researchers were not blinded to the data. The 9 participants could not be blinded to the conditions in which they were tested, but they were not 10 informed about any hypothesis regarding which conditions were optimal. All participants provided 11 written informed consent. The Institutional Review Board of the University of Nebraska Medical 12 Center approved the study. 13

14 **Protocol**

The study consisted of a habituation session and a testing session one week later. Each 15 session contained 1.8 hours of walking. The habituation session duration exceeded reported 16 durations to maximize benefits from different wearable robots [forty minutes for a hip exosuit (37), 17 105 minutes for an ankle exoskeleton (38)]. The testing session started with a five-minute standing 18 19 trial to measure resting metabolic rate (K5, Cosmed, Rome, Italy). To ensure accurate 20 measurement of the resting metabolic rate, we had the participant rest for at least fifteen minutes while we prepared the experiment, and we asked the participants to have fasted for at least five 21 22 hours, abstain from caffeine and alcohol overnight, and abstain from vigorous physical activity for 23 fourteen hours prior to the data collection (39). Both sessions had a ten-minute warm-up during

1	which we cycled through all conditions and tuned the gains. The desired treadmill speed was set
2	to 1.25 m s ⁻¹ (Bertec). Near the end of the warm-up, we determined each participant's preferred
3	step frequency under the Zero Force condition. This step frequency was used to pace the participant
4	via a metronome to avoid walking pace changes that would affect the metabolic rate (40).
5	Participants were able to keep their mean step time within 0.011 ± 0.008 s (mean \pm inter-participant
6	SD from all conditions) of their instructed step time $(0.562 \pm 0.021 \text{ s})$.

Participants walked under thirty-six different force profiles that were randomized and 7 grouped into three blocks, each separated by ten minutes of rest (fig. S12A). Thirty-two Sinusoidal 8 9 Force conditions were combinations of three desired durations (33, 66, and 99% of step time), four desired onset timings (0, 25, 50, and 75%), and different desired peak net aiding force magnitudes 10 (ranging from 4 to 24% of BW; all force levels are reported as net aiding forces, and the tether 11 forces were 5% higher to offset the parallel gravity component from the treadmill inclination). As 12 with other devices (41, 42), our system had a specific bandwidth limit [3 dB bandwidth = 10 Hz 13 (17)]. This required us to use lower peak force ranges at shorter durations (fig. S12B). We tested 14 three conditions where the desired net aiding force remained at a constant level (4, 8, and 12% 15 BW) throughout the step cycle (Constant Force conditions). The second-highest constant force 16 17 level was chosen based on the mean of the optima of earlier studies [7% (7) and 9% (4)]. We compared the effects of all the Sinusoidal and Constant Force conditions to a baseline condition 18 19 where the desired net aiding force was set to 0% BW (Zero Force condition). Because of potential 20 metabolic drift (43-45), we repeated the Zero Force condition, similar to other studies (12, 23). We also repeated one Sinusoidal Force condition that was randomly selected for each participant 21 22 in each block, resulting in a total of forty conditions. The first and last conditions of each block 23 lasted five minutes to ensure that the metabolic rate could reach steady state. We switched between

the other conditions every two minutes using estimation methods from the literature (10, 46) to
obtain the steady-state metabolic rate.

3 Measurements and data processing

Oxygen consumption and carbon dioxide production were measured using indirect 4 calorimetry (K5, Cosmed, Rome, Italy) during the entire protocol. We calibrated the indirect 5 calorimetry system before every session using a gas container with a known oxygen and carbon 6 dioxide concentration and a 3 L fixed-volume calibration syringe that delivers simulated breath 7 volume. The experiments occurred in a large and well-ventilated room. Breath-by-breath 8 measurements were converted to W kg⁻¹ using the Brockway equation (47). The respiratory 9 exchange ratios were lower than 1 (0.768 \pm 0.053, mean \pm inter-participant SD of all conditions, 10 *P*-values < 0.001, one-sample t tests), which confirmed that the intensity level was within the 11 aerobic range where it is appropriate to use the Brockway equation (48). We estimated the steady-12 state metabolic rate of the resting trial and the conditions at the beginning of each block by 13 averaging the breath-by-breath data in the final two minutes. For all other conditions, the steady-14 state metabolic rates were estimated by fitting the breath-by-breath data from immediately after 15 the transition to each new condition until right before the change to the next condition with an 16 17 exponential function and estimating the asymptote (10, 46) (fig. S13A).

For each participant, we identified the time constant that minimized the squared error of the exponential fits versus the breath-by-breath metabolic rate measurements following guidelines from Selinger and Donelan (46). Based on the properties of our data (breath frequency of $19.8 \pm$ 2.04 breaths min⁻¹, inter-breath SD of 0.656 ± 0.189 W kg⁻¹, average change between conditions of 0.664 ± 0.193 W kg⁻¹), the recommended minimum number of conditions to approximate time constants with a confidence interval of 95% is 49. Data from both sessions (80 conditions) were

1	used to meet this recommendation for time constant identification, and the resulting time constants
2	of our participants (46.1 \pm 15.7 s, mean \pm inter-participant SD) were close to the time constants
3	reported by Selinger and Donelan (46) (41.8 \pm 12.1 s). To evaluate the metabolic rate estimation
4	accuracy, we compared estimations based on two minutes of data to estimations that used five
5	minutes from the final conditions of each block. The mean absolute error was $4.24 \pm 2.60\%$ of the
6	Zero Force condition, and this result is on the lower end of a range of errors in similar methods
7	[from 4.3% (10) up to 12.4% (49)].
8	We calculated the net metabolic rate by subtracting the resting metabolic rate. To evaluate
9	metabolic drift due to the long protocol, we fit a linear trend through the net metabolic rate of the
10	Zero Force conditions over time (fig. S13 B and C). We chose a linear fit since, according to the
11	literature, the metabolic drift trends look predominantly linear (44, 45). The average slope of this
12	trend during the testing session was higher than zero ($P = 0.0256$, one-sample t test). Studies show

trend during the testing session was higher than zero (P = 0.0256, one-sample t test). Studies show 12 that prolonged downhill running can evoke a linear drift in oxygen consumption (44, 45). Thus, 13 the fact that walking with net aiding forces is mechanically similar to downhill locomotion might 14 explain why a drift occurred in our experiment. To correct for the metabolic drift, we calculated 15 all W kg⁻¹ reductions in metabolic rate from the testing session versus the linear drift trend. To 16 avoid the metabolic drift affecting normalization of the percentage reduction in metabolic rate, we 17 divided the drift-adjusted reductions by the intercept value of the linear fit, which represented the 18 Zero Force metabolic rate before fatigue. Analysis of the repeated Sinusoidal Force conditions 19 20 showed that the drift correction improved the intraclass correlation of the repeated conditions from 0.629 to 0.766. This is considered good to excellent for intraclass correlation and is of a similar 21 order of magnitude as reported intraclass correlations for within-session repeatability of VO2 22 23 measurements [0.87 (50)].

1	We measured the GRF and load cell data during the last minute of each condition.
2	Crossover steps between belts were removed, and ~ 50 steps per condition per participant were
3	used. The force treadmill (Bertec) was calibrated using an instrumented pole (C-motion,
4	Germantown, MD) (51) with an accuracy threshold of \pm 5 mm for the center of pressure. We
5	minimized signal drift by zeroing between walking blocks and subtracting the median of the swing
6	phases. The GRF and load cell (Futek) signals were smoothed with a 10 Hz low-pass filter. We
7	chose the cut-off based on the typical frequency content of walking [6 Hz (52)] and the robotic
8	tether bandwidth [10 Hz (17)]. We calculated the parallel and perpendicular GRFs by performing
9	a coordinate transformation over the treadmill inclination. This inclination was verified using a
10	bubble level and a motion capture system (VICON Vero), which was calibrated to an accuracy of
11	0.6 mm. There was a strong linear relationship between the mean parallel GRF and net aiding force
12	with a slope coefficient of -1.004 \pm 0.005 (mean \pm inter-participant SD), which confirmed that the
13	calibration of both devices was consistent. To further minimize errors due to the load cell offset,
14	we subtracted the mean sum of the parallel components of the GRF, gravity, and the load cell force
15	based on the fact that the sum of all forces must be in equilibrium, on average.
16	We calculated the COM acceleration from the GRF, tether force, and gravity using
17	Newton's second law (53). The total mass was measured by the force treadmill (Bertec), and we
18	verified that this value corresponded to the body mass plus the added mass (~4.9 kg for the waist
19	belt, calorimetry unit, shoes, and other small equipment). We calculated the velocity and position
20	of the COM by integrating the COM acceleration (54). To obtain the COM velocity relative to a

treadmill velocity (1.26 m s⁻¹) was obtained by recording motion capture markers attached to the treadmill belt (55). To calculate the net aiding power, we multiplied the net aiding force by the

coordinate system that moves with the treadmill belt, we added the treadmill velocity. The actual

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1 COM velocity in the direction of the force and relative to the treadmill belt coordinate system. We 2 calculated individual leg COM power by taking the dot product of the COM velocity and the 3 individual (right) GRF (*56*). GRF recordings in three of the 400 trials from the testing session 4 failed due to equipment malfunctions. For these trials, we calculated the net aiding power by 5 assuming that the parallel COM velocity was equal to the treadmill velocity. The GRF variables 6 of these trials were treated as missing values.

We segmented each time series into steps and strides based on heel strike detection using 7 the vertical GRF. For each participant and condition, we calculated the median of the strides (data 8 9 S3). For the Repeated Sinusoidal Force conditions and the Zero Force condition, we averaged three repetitions of these conditions. The metabolic rate and kinetic time series were normalized relative 10 to body mass. The peak values, step averages, and averages of positive and negative portions of 11 tether and GRF variables were calculated from the time-normalized data (table S6). We also 12 calculated the net aiding force duration and net aiding power duration by evaluating the length of 13 the net aiding force and net aiding power profile that was higher than 1% BW or 0.15 W kg⁻¹. 14 When calculating the peak force timing and peak power timing, we avoided creating artifacts in 15 inter-participant variability from step segmentation by converting peak timings that were close to 16 17 0 or 100% to a percentage below 0% or above 100% in cases where the majority of the peak timings in the same condition occurred at the opposite end of the step. For example, if the peak 18 timing of a certain condition was 99% of the step in one participant and 1% in all the other 19 participants, then the peak timing of 99% was converted to a value of -1%, which has the same 20 meaning as a peak timing of 99%. 21

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1 Statistical analyses

To determine the effects of the actuation profile shape parameters on the dependent 2 parameters, we used mixed-effects model analyses with participants as random factors (22, 57, 3 58). To avoid overfitting, terms that did not significantly contribute were removed using a stepwise 4 elimination procedure whereby the least significant terms were removed until only significant 5 6 terms remained (59–61). Since earlier research shows that the metabolic rate follows a U-shaped trend versus aiding force magnitude (4, 7), we included first- and second-order terms for the 7 actuation magnitude parameter. There are no prior data on the effect of actuation duration, and in 8 9 light of this, we began using both first- and second-order terms for this parameter. Since the timing was varied over the entire step, we expected a continuous trend between the end of the step cycle 10 and the beginning of the step cycle. In other words, even if the step cycle were defined differently 11 (e.g., from toe-off to toe-off instead of from heel strike to heel strike), we would expect to see a 12 similar (but shifted) continuous trend. Therefore, a periodic term was included for timing. We 13 identified the mean phase shift that minimized the sum of squared errors between the model and 14 the measurements for each participant. As there cannot be an effect of duration or timing when the 15 peak magnitude is zero, the duration and timing terms were multiplied by the peak magnitude (13, 16 17 60). Since the actuation magnitude and reduction in metabolic rate were zero in the Zero Force condition, we did not include an intercept. Finally, because we aimed to compare our results to 18 those of devices that provide joint work and since mechanical work is a component of muscle 19 20 energetic cost (62), we initially chose to express all parameters based on work rate and power instead of the average force and peak force, resulting in the following initial statistical model: 21

22
$$c_1 \cdot \dot{W}_{aid} + c_2 \cdot \dot{W}_{aid}^2 + c_3 \cdot P_{peak} \cdot \Delta t_P + c_4 \cdot P_{peak} \cdot \Delta t_P^2 + c_5 \cdot P_{peak} \cdot \sin\left(\frac{t_{p_{peak}} + c_6}{100} \cdot 2\pi\right)$$
(1)

1	where \dot{W}_{aid} is the net aiding work rate obtained by averaging the net aiding power over the step
2	duration, P_{peak} is the peak net aiding power, Δt_P is the duration of the power burst, and t_{Ppeak} is the
3	timing of the peak power. The metabolic rate, \dot{W}_{aid} , and P_{peak} are in W kg ⁻¹ . Δt_P and t_{Ppeak} are
4	expressed in % of step time.
5	To evaluate these choices, we compared the estimated and actual condition
6	averages for several alternative models (15). Since we eliminated non-significant terms, all terms
7	in the final equations were significant ($P < 0.05$). We used the adjusted R ² values to assess the
8	added value of different numbers of predictors. To analyze the Constant Force conditions, we used
9	a model that included only the net aiding work rate terms.
10	
11	$c_1 \cdot \dot{W}_{aid} + c_2 \cdot \dot{W}_{aid}^2 \tag{2}$
12	Final equation:
13	$-3.25 \cdot \dot{W}_{aid} + 1.76 \cdot \dot{W}_{aid}^2$ Adjusted R ² : 99.8%
14	
15	The controller did not keep the aiding work rate constant between conditions with different
16	timings as this would have required real-time COM velocity measurement. However, the utilized
17	statistical method of linear mixed-effects model analysis does not require varying the different
18	
	independent parameters in isolation (e.g., unlike approaches such as repeated measures ANOVA).
19	
	independent parameters in isolation (e.g., unlike approaches such as repeated measures ANOVA).
19	independent parameters in isolation (e.g., unlike approaches such as repeated measures ANOVA). To determine the effects of timing isolated from work, we evaluated the mixed-effects model
19 20	independent parameters in isolation (e.g., unlike approaches such as repeated measures ANOVA). To determine the effects of timing isolated from work, we evaluated the mixed-effects model analysis equation over the range of timings while keeping the magnitude parameters (\dot{W}_{aid} and
19 20 21	independent parameters in isolation (e.g., unlike approaches such as repeated measures ANOVA). To determine the effects of timing isolated from work, we evaluated the mixed-effects model analysis equation over the range of timings while keeping the magnitude parameters (\dot{W}_{aid} and P_{peak}) constant. To determine the effects of the net aiding work rate, we evaluated the equation

1	the inter-participant variability, we fit the significant terms of the equation to each participant's
2	data using the <i>fminsearch</i> function in MATLAB, and we determined the metabolic rate, peak
3	timing, and net aiding work rate at the optimum of each participant. This interpolated optimum
4	was defined as the Optimum timing and magnitude, and the condition with the best mean reduction
5	in metabolic rate was defined as the Best timing and magnitude. To evaluate differences in the
6	metabolic rate, propulsion GRF, COM velocity, and COM power between conditions, paired t tests
7	with Holm-Šidák correction were used for multiple comparisons (63). For statistical tests that
8	relied on the normality assumption, we verified that the data followed a normal distribution using
9	the Jarque-Bera test (table S6).

References and Notes:

- B. R. Duffy, Anthropomorphism and the social robot. *Rob. Auton. Syst.* 42, 177–190 (2003).
- A. J. Young, D. P. Ferris, State-of-the-art and Future Directions for Robotic Lower Limb Robotic Exoskeletons. *IEEE Trans. Neural Syst. Rehabil. Eng.* 25, 171–182 (2017).
- A. D. Kuo, Energetics of actively powered locomotion using the simplest walking model.
 J. Biomech. Eng. 124, 113–120 (2002).
- J. S. Gottschall, R. Kram, Energy cost and muscular activity required for propulsion during walking. *J. Appl. Physiol.* 94, 1766–1772 (2003).
- S. G. Bhat, S. Cherangara, J. Olson, S. Redkar, T. G. Sugar, Analysis of a Periodic Force Applied to the Trunk to Assist Walking Gait. 2019 Wearable Robot. Assoc. Conf. WearRAcon 2019, 68–73 (2019).
- K. Penke, K. Scott, Y. Sinskey, M. D. Lewek, Propulsive Forces Applied to the Body's Center of Mass Affect Metabolic Energetics Poststroke. *Arch. Phys. Med. Rehabil.* 100, 1068–1075 (2019).
- C. A. Zirker, B. C. Bennett, M. F. Abel, Changes in Kinematics, Metabolic Cost, and External Work During Walking with a Forward Assistive Force. *J. Appl. Biomech.* 29, 481–489 (2013).
- J. Rose, J. G. Gamble, A. Burgos, J. Medeiros, W. L. Haskell, Energy expenditure index of walking for normal children and for children with cerebral palsy. *Dev. Med. Child Neurol.* 32, 333–40 (1990).
- 9. J. Lee, K. Seo, B. Lim, J. Jang, K. Kim, H. Choi, Effects of assistance timing on metabolic cost, assistance power, and gait parameters for a hip-type exoskeleton. *IEEE Int. Conf.*

Rehabil. Robot., 498–504 (2017).

- J. Zhang, P. Fiers, K. A. Witte, R. W. Jackson, K. L. Poggensee, C. G. Atkeson, S. H. Collins, Human-in-the-loop optimization of exoskeleton assistance during walking. *Science*. 356, 1280–1284 (2017).
- 11. R. W. Jackson, S. H. Collins, An experimental comparison of the relative benefits of work and torque assistance in ankle exoskeletons. *J. Appl. Physiol.* **119**, 541–557 (2015).
- B. T. Quinlivan, S. Lee, P. Malcolm, D. M. Rossi, M. Grimmer, C. Siviy, N. Karavas, D. Wagner, A. Asbeck, I. Galiana, C. J. Walsh, D. Wagner, C. Siviy, M. Grimmer, S. Lee, B. T. Quinlivan, P. Malcolm, C. J. Walsh, D. M. Rossi, Assistance magnitude versus metabolic cost reductions for a tethered multiarticular soft exosuit. *Sci. Robot.* 2, eaah4416 (2017).
- S. Galle, P. Malcolm, S. H. Collins, D. De Clercq, Reducing the metabolic cost of walking with an ankle exoskeleton: interaction between actuation timing and power. *J. Neuroeng. Rehabil.* 14, 1–16 (2017).
- I. Kang, H. Hsu, A. Young, The Effect of Hip Assistance Levels on Human Energetic Cost Using Robotic Hip Exoskeletons. *IEEE Robot. Autom. Lett.* 4, 430–437 (2019).
- Supplementary Materials and Methods, Supplementary Discussion, Figures, Tables, Movie and Data are available as supplementary materials.
- 16. J. M. Caputo, S. H. Collins, Prosthetic ankle push-off work reduces metabolic rate but not collision work in non-amputee walking. *Sci. Rep.* **4**, 7213 (2014).
- A. M. Gonabadi, P. Antonellis, P. Malcolm, A System for Simple Robotic Walking Assistance With Linear Impulses at the Center of Mass. *IEEE Trans. Neural Syst. Rehabil. Eng.* 28, 1353–1362 (2020).

- S. J. Abram, J. C. Selinger, J. M. Donelan, Energy optimization is a major objective in the real-time control of step width in human walking. *J. Biomech.* 91, 85–91 (2019).
- G. A. Cavagna, F. P. Saibene, R. Margaria, External work in walking. J. Appl. Physiol. 18, 1–9 (1963).
- R. Margaria, Positive and negative work performances and their efficiencies in human locomotion. *Int. Zeitschrift für Angew. Physiol. Einschließlich Arbeitsphysiologie.* 25, 339–351 (1968).
- 21. L. C. Hunter, E. C. Hendrix, J. C. Dean, The cost of walking downhill: Is the preferred gait energetically optimal? *J. Biomech.* **43**, 1910–1915 (2010).
- S. H. Collins, M. B. Wiggin, G. S. Sawicki, M. Bruce Wiggin, G. S. Sawicki, Reducing the energy cost of human walking using an unpowered exoskeleton. *Nature*. 522, 5–8 (2015).
- 23. L. M. Mooney, E. J. Rouse, H. M. Herr, Autonomous exoskeleton reduces metabolic cost of human walking. *J. Neuroeng. Rehabil.* **11**, 151 (2014).
- T. G. Hornby, D. S. Reisman, I. G. Ward, P. L. Scheets, A. Miller, D. Haddad, E. J. Fox, N. E. Fritz, K. Hawkins, C. E. Henderson, K. L. Hendron, C. L. Holleran, J. E. Lynskey, A. Walter, Clinical Practice Guideline to Improve Locomotor Function Following Chronic Stroke, Incomplete Spinal Cord Injury, and Brain Injury. *J. Neurol. Phys. Ther.* 44, 49–100 (2020).
- A. D. Segal, K. E. Zelik, G. K. Klute, D. C. Morgenroth, M. E. Hahn, M. S. Orendurff, P. G. Adamczyk, S. H. Collins, A. D. Kuo, J. M. Czerniecki, The effects of a controlled energy storage and return prototype prosthetic foot on transtibial amputee ambulation. *Hum. Mov. Sci.* 31, 918–931 (2012).

- R. E. Quesada, J. M. Caputo, S. H. Collins, Increasing ankle push-off work with a powered prosthesis does not necessarily reduce metabolic rate for transtibial amputees. *J. Biomech.* 49, 3452–3459 (2016).
- J. A. Norris, K. P. Granata, M. R. Mitros, E. M. Byrne, A. P. Marsh, Effect of augmented plantarflexion power on preferred walking speed and economy in young and older adults. *Gait Posture*. 25, 620–627 (2007).
- 28. Y. L. Kerkum, A. I. Buizer, J. C. Van Den Noort, J. G. Becher, J. Harlaar, M. A. Brehm, The effects of varying ankle foot orthosis stiffness on gait in children with spastic cerebral palsy who walk with excessive knee flexion. *PLoS One.* 10, e0142878 (2015).
- K. Z. Takahashi, M. D. Lewek, G. S. Sawicki, A neuromechanics-based powered ankle exoskeleton to assist walking post-stroke: A feasibility study. *J. Neuroeng. Rehabil.* 12, 1– 13 (2015).
- G. M. Gasparri, J. Luque, Z. F. Lerner, Proportional Joint-Moment Control for Instantaneously Adaptive Ankle Exoskeleton Assistance. *IEEE Trans. neural Syst. Rehabil. Eng.* 27, 751–759 (2019).
- V. Vashista, X. Jin, S. K. Agrawal, Active Tethered Pelvic Assist Device (A-TPAD) to study force adaptation in human walking. *Proc. - IEEE Int. Conf. Robot. Autom.*, 718–723 (2014).
- K. H. Seo, J. J. Lee, The development of two mobile gait rehabilitation systems. *IEEE Trans. Neural Syst. Rehabil. Eng.* 17, 156–166 (2009).
- A. H. Dewolf, Y. P. Ivanenko, R. M. Mesquita, F. Lacquaniti, P. A. Willems, Neuromechanical adjustments when walking with an aiding or hindering horizontal force. *Eur. J. Appl. Physiol.* 120, 91–106 (2020).

- 34. P. Malcolm, D. M. Rossi, C. Siviy, S. Lee, B. T. Quinlivan, M. Grimmer, C. J. Walsh, Continuous sweep versus discrete step protocols for studying effects of wearable robot assistance magnitude. *J Neuroeng Rehabil.* 14, 72 (2017).
- P. Malcolm, R. E. Quesada, J. M. Caputo, S. H. Collins, The influence of push-off timing in a robotic ankle-foot prosthesis on the energetics and mechanics of walking. *J Neuroeng Rehabil.* 12, 21 (2015).
- P. Malcolm, W. Derave, S. Galle, D. De Clercq, A Simple Exoskeleton That Assists Plantarflexion Can Reduce the Metabolic Cost of Human Walking. *PLoS One.* 8 (2013), doi:10.1371/journal.pone.0056137.
- F. A. Panizzolo, G. M. Freisinger, N. Karavas, A. M. Eckert-Erdheim, C. Siviy, A. Long,
 R. A. Zifchock, M. E. LaFiandra, C. J. Walsh, Metabolic cost adaptations during training with a soft exosuit assisting the hip joint. *Sci. Rep.* 9, 1–10 (2019).
- 38. M. Bruce Wiggin, thesis, North Carolina State University (2014).
- C. Compher, D. Frankenfield, N. Keim, L. Roth-Yousey, Evidence Analysis Working Group, Best practice methods to apply to measurement of resting metabolic rate in adults: a systematic review. *J. Am. Diet. Assoc.* 106, 881–903 (2006).
- M. Y. Zarrugh, C. W. Radcliffe, Predicting metabolic cost of level walking. *Eur. J. Appl. Physiol. Occup. Physiol.* 38, 215–23 (1978).
- S. N. Simha, J. D. Wong, J. C. Selinger, J. M. Donelan, A Mechatronic System for Studying Energy Optimization During Walking. *IEEE Trans. Neural Syst. Rehabil. Eng.* 27, 1416–1425 (2019).
- 42. G. Brown, M. M. Wu, F. C. Huang, K. E. Gordon, Movement augmentation to evaluate human control of locomotor stability. *Proc. Annu. Int. Conf. IEEE Eng. Med. Biol. Soc.*

EMBS, 66–69 (2017).

- 43. R. a. Robergs, D. Dwyer, T. Astorino, Recommendations for improved data processing from expired gas analysis indirect calorimetry. *Sport. Med.* **40**, 95–111 (2010).
- K. C. Westerlind, W. C. Byrnes, R. S. Mazzeo, A comparison of the oxygen drift in downhill vs. level running. *J. Appl. Physiol.* 72, 796–800 (1992).
- 45. R. W. Dick, P. R. Cavanagh, An explanation of the upward drift in oxygen uptake during prolonged sub-maximal downhill running. *Med. Sci. Sports Exerc.* **19**, 310–7 (1987).
- J. C. Selinger, J. M. Donelan, Estimating instantaneous energetic cost during non-steadystate gait. J. Appl. Physiol. 117, 1406–1415 (2014).
- J. M. Brockway, Derivation of formulae used to calculate energy expenditure in man. *Hum. Nutr. Clin. Nutr.* 41, 463–71 (1987).
- S. Kipp, W. C. Byrnes, R. Kram, Calculating metabolic energy expenditure across a wide range of exercise intensities: the equation matters. *Appl. Physiol. Nutr. Metab.* 43, 639-642 (2018).
- 49. K. A. Witte, P. Fiers, A. L. Sheets-Singer, S. H. Collins, Improving the energy economy of human running with powered and unpowered ankle exoskeleton assistance. *Sci. Robot.* 5, eaay9108 (2020).
- R. Duffield, B. Dawson, H. C. Pinnington, P. Wong, Accuracy and reliability of a Cosmed K4b2 portable gas analysis system. *J. Sci. Med. Sport.* 7, 11–22 (2004).
- S. H. Collins, P. G. Adamczyk, D. P. Ferris, A. D. Kuo, A simple method for calibrating force plates and force treadmills using an instrumented pole. *Gait Posture*. 29, 59–64 (2009).
- 52. D. A. Winter, Biomechanics and Motor Control of Human Movement (John Wiley &

Sons, Inc., Hoboken, NJ, USA, 2009; http://doi.wiley.com/10.1002/9780470549148).

- 53. I. Newton, *Newton's Principia: The Mathematical Principles of Natural Philosophy* (1687).
- 54. G. A. Cavagna, Force platforms as ergometers. J. Appl. Physiol. 39, 174–9 (1975).
- 55. Savelberg, Vorstenbosch, Kamman, van de Weijer JG, Schambardt, Intra-stride belt-speed variation affects treadmill locomotion. *Gait Posture*. **7**, 26–34 (1998).
- J. M. Donelan, R. Kram, A. D. Kuo, Simultaneous positive and negative external mechanical work in human walking. *J Biomech.* 35, 117–124 (2002).
- 57. A. Gałecki, T. Burzykowski, (2013; http://link.springer.com/10.1007/978-1-4614-3900-4_13), pp. 245–273.
- W. P. Vispoel, Book Reviews: Statistical Reasoning for the Behavioral Sciences (2nd ed.) Richard J. Shavelson Needham Heights, MA: Allyn and Bacon, 1988. viii + 744 pp. J. Educ. Stat. 15, 179–183 (1990).
- B. X. W. Liew, S. Morris, K. Netto, The effects of load carriage on joint work at different running velocities. *J. Biomech.* 49, 3275–3280 (2016).
- 60. P. Antonellis, S. Galle, D. De Clercq, P. Malcolm, Altering gait variability with an ankle exoskeleton. *PLoS One.* **13**, e0205088 (2018).
- P. Antonellis, C. M. Frederick, A. M. Gonabadi, P. Malcolm, Modular footwear that partially offsets downhill or uphill grades minimizes the metabolic cost of human walking. *R. Soc. Open Sci.* 7, 191527 (2020).
- 62. B. R. Umberger, K. G. M. Gerritsen, P. E. Martin, A model of human muscle energy expenditure. *Comput. Methods Biomech. Biomed. Engin.* **6**, 99–111 (2003).
- 63. Glantz SA, Primer of Biostatistics (McGraw-Hill, New York, NY, USA, 2005).

- 64. B. B. Lloyd, R. M. Zacks, The mechanical efficiency of treadmill running against a horizontal impeding force. *J. Physiol.* **223**, 355–63 (1972).
- 65. Y. H. Chang, R. Kram, Metabolic cost of generating horizontal forces during human running. *J. Appl. Physiol.* **86**, 1657–1662 (1999).
- M. Saunders, V. Inman, H. Eberhart, The major determinants in normal and pathological gait. J. Bones Jt. Surg. 35, 543–558 (1953).
- 67. G. A. Pratt, M. M. Williamson, Series elastic actuators. *IEEE Int. Conf. Intell. Robot. Syst.*1, 399–406 (1995).
- J. Zhang, S. H. Collins, The Passive Series Stiffness That Optimizes Torque Tracking for a Lower-Limb Exoskeleton in Human Walking. *Front. Neurorobot.* 11, 1–16 (2017).
- S. Qian, B. Zi, W. W. Shang, Q. S. Xu, A review on cable-driven parallel robots. *Chinese J. Mech. Eng. (English Ed.* 31 (2018), doi:10.1186/s10033-018-0267-9.
- J. Zhang, C. C. Cheah, S. H. Collins, in 2015 IEEE International Conference on Robotics and Automation (ICRA) (IEEE, Seattle, 2015; http://ieeexplore.ieee.org/document/7139980/), pp. 5584–5589.
- S. Mochon, T. A. McMahon, Ballistic walking: an improved model. *Math. Biosci.* 52, 241–260 (1980).
- R. M. Alexander, Simple Models of Human Movement. *Appl. Mech. Rev.* 48, 461–470 (1995).
- 73. M. Garcia, A. Chatterjee, A. Ruina, M. Coleman, The simplest walking model: stability, complexity, and scaling. *J. Biomech. Eng.* **120**, 281–8 (1998).
- 74. T. McGeer, Passive Dynamic Walking. Int. J. Rob. Res. 9, 62–82 (1990).
- 75. M. Srinivasan, Fifteen observations on the structure of energy-minimizing gaits in many

simple biped models. J. R. Soc. Interface. 8, 74–98 (2011).

- S. Faraji, A. R. Wu, A. J. Ijspeert, A simple model of mechanical effects to estimate metabolic cost of human walking. *Sci. Rep.* 8, 1–12 (2018).
- 77. R. M. Alexander, Optimum Muscle Design for Oscillatory Movements. J. Theor. Biol.
 184, 253–259 (1997).
- 78. K. E. Zelik, A. D. Kuo, Human walking isn't all hard work: evidence of soft tissue contributions to energy dissipation and return. *J. Exp. Biol.* **213**, 4257–64 (2010).
- 79. A. J. Greene, P. J. Sinclair, M. H. Dickson, F. Colloud, R. M. Smith, Relative shank to thigh length is associated with different mechanisms of power production during elite male ergometer rowing. *Sport. Biomech.* 8, 302–17 (2009).
- 80. A. E. Minetti, C. Moia, G. S. Roi, D. Susta, G. Ferretti, Energy cost of walking and running at extreme uphill and downhill slopes. *J. Appl. Physiol.* **93**, 1039–46 (2002).
- L. M. Mooney, E. J. Rouse, H. M. Herr, Autonomous exoskeleton reduces metabolic cost of human walking during load carriage. *J Neuroeng Rehabil.* 11 (2014).
- J. Kim, G. Lee, R. Heimgartner, D. A. Revi, N. Karavas, D. Nathanson, I. Galiana, A. Eckert-Erdheim, P. Murphy, D. Perry, N. Menard, D. K. Choe, P. Malcolm, C. J. Walsh, Reducing the metabolic rate of walking and running with a versatile, portable exosuit. *Science (80-.).* 365, 668–672 (2019).
- R. C. Browning, J. R. Modica, R. Kram, A. Goswami, The effects of adding mass to the legs on the energetics and biomechanics of walking. *Med Sci Sport. Exerc.* 39, 515–525 (2007).
- J. B. de V. Weir, New methods for calculating metabolic rate with special reference to protein metabolism. J. Physiol. 109, 1–9 (1949).

- C. J. Wutzke, G. S. Sawicki, M. D. Lewek, The influence of a unilateral fixed ankle on metabolic and mechanical demands during walking in unimpaired young adults. *J. Biomech.* 45, 2405–2410 (2012).
- 86. I. I. Pipinos, A. R. Judge, J. T. Selsby, Z. Zhu, S. A. Swanson, A. A. Nella, S. L. Dodd, The myopathy of peripheral arterial occlusive disease: Part 1. Functional and histomorphological changes and evidence for mitochondrial dysfunction. *Vasc. Endovascular Surg.* 41, 481–489 (2008).
- 87. I. I. Pipinos, A. R. Judge, J. T. Selsby, Z. Zhu, S. A. Swanson, A. A. Nella, S. L. Dodd, The myopathy of peripheral arterial occlusive disease: Part 2. Oxidative stress, neuropathy, and shift in muscle fiber type. *Vasc. Endovascular Surg.* 42, 101–112 (2008).
- F. G. R. Fowkes, D. Rudan, I. Rudan, V. Aboyans, J. O. Denenberg, M. M. McDermott,
 P. E. Norman, U. K. A. Sampson, L. J. Williams, G. A. Mensah, M. H. Criqui,
 Comparison of global estimates of prevalence and risk factors for peripheral artery disease
 in 2000 and 2010: A systematic review and analysis. *Lancet.* 382, 1329–1340 (2013).
- B. H. Goodpaster, S. W. Park, T. B. Harris, S. B. Kritchevsky, M. Nevitt, A. V. Schwartz,
 E. M. Simonsick, F. A. Tylavsky, M. Visser, A. B. Newman, The loss of skeletal muscle strength, mass, and quality in older adults: The Health, Aging and Body Composition
 Study. *Journals Gerontol. Ser. A Biol. Sci. Med. Sci.* 61, 1059–1064 (2006).
- 90. S. Rossi, A. Colazza, M. Petrarca, E. Castelli, P. Cappa, H. I. Krebs, Feasibility Study of a Wearable Exoskeleton for Children: Is the Gait Altered by Adding Masses on Lower Limbs? *PLoS One.* 8, e73139 (2013).
- S. Patel, B. L. Patritti, J. Nikitczuk, B. Weinberg, U. D. Croce, C. Mavroidis, P. Bonato,
 Effects on normal gait of a new active knee orthosis for hemiparetic gait retraining. *Annu.*

Int. Conf. IEEE Eng. Med. Biol. - Proc. 2006, 1232–1235 (2006).

- S. M. Cain, K. E. Gordon, D. P. Ferris, Locomotor adaptation to a powered ankle-foot orthosis depends on control method. *J. Neuroeng. Rehabil.* 4, 48 (2007).
- 93. D. Torricelli, J. Gonzalez-Vargas, J. F. Veneman, K. Mombaur, N. Tsagarakis, A. J. Del-Ama, A. Gil-Agudo, J. C. Moreno, J. L. Pons, Benchmarking Bipedal Locomotion: A Unified Scheme for Humanoids, Wearable Robots, and Humans. *IEEE Robot. Autom. Mag.* 22, 103–115 (2015).
- 94. L. N. Awad, J. Bae, P. Kudzia, A. Long, K. Hendron, K. G. Holt, K. O'Donnell, T. D. Ellis, C. J. Walsh, Reducing Circumduction and Hip Hiking During Hemiparetic Walking Through Targeted Assistance of the Paretic Limb Using a Soft Robotic Exosuit. *Am. J. Phys. Med. Rehabil.* 96, S157–S164 (2017).
- 95. D. R. Carrier, C. Anders, N. Schilling, The musculoskeletal system of humans is not tuned to maximize the economy of locomotion. *Proc. Natl. Acad. Sci.* 108, 18631–18636 (2011).
- 96. A. Grabowski, C. T. Farley, R. Kram, Independent metabolic costs of supporting body weight and accelerating body mass during walking. *J. Appl. Physiol.* **98**, 579–583 (2005).
- J. Markowitz, H. Herr, Human Leg Model Predicts Muscle Forces, States, and Energetics during Walking. *PLOS Comput. Biol.* 12, e1004912 (2016).
- B. R. Umberger, Stance and swing phase costs in human walking. J. R. Soc. Interface. 7, 1329–1340 (2010).
- A. M. Gonabadi, P. Antonellis, P. Malcolm, Differences between joint-space and musculoskeletal estimations of metabolic rate time profiles. *PLoS Comput. Biol.* 16, e1008280 (2020).

- K. M. Newell, Y. T. Liu, G. Mayer-Kress, Time scales in motor learning and development. *Psychol. Rev.* 108, 57–82 (2001).
- 101. S. Galle, P. Malcolm, W. Derave, D. De Clercq, Adaptation to walking with an exoskeleton that assists ankle extension. *Gait Posture*. **38**, 495–499 (2013).
- 102. J. R. Koller, D. A. Jacobs, D. P. Ferris, C. D. Remy, Learning to walk with an adaptive gain proportional myoelectric controller for a robotic ankle exoskeleton. *J. Neuroeng. Rehabil.* 12, 97 (2015).
- 103. A. J. Young, J. Foss, H. Gannon, D. P. Ferris, Influence of Power Delivery Timing on the Energetics and Biomechanics of Humans Wearing a Hip Exoskeleton. *Front. Bioeng. Biotechnol.* 5, 1–11 (2017).
- 104. Y. Ding, F. A. Panizzolo, C. Siviy, P. Malcolm, I. Galiana, K. G. Holt, C. J. Walsh, Effect of timing of hip extension assistance during loaded walking with a soft exosuit. J. *Neuroeng. Rehabil.* 13, 87 (2016).
- 105. P. Antonellis, A. M. Gonabadi, P. Malcolm, Effects of timing and magnitude of forward forces at the waist on the metabolic cost of walking. *Am. Soc. Biomech.* (2019).
- 106. R. Schroeder, J. Croft, J. Bertram, The Role of Entrainment in Human Walking: Energy Minimization in Oscillating Environments. *Sci. Rep.* (2021), doi:10.21203/rs.3.rs-120108/v1.
- A. M. Jones, J. H. Doust, A 1% treadmill grade most accurately reflects the energetic cost of outdoor running. *J. Sports Sci.* 14, 321–7 (1996).
- K. E. Bijker, G. De Groot, A. P. Hollander, Delta efficiencies of running and cycling. *Med. Sci. Sports Exerc.* 33, 1546–1551 (2001).
- 109. H.-M. Maus, S. W. Lipfert, M. Gross, J. Rummel, A. Seyfarth, Upright human gait did not

provide a major mechanical challenge for our ancestors. Nat. Commun. 1, 70 (2010).

- 110. J. Doke, Mechanics and energetics of swinging the human leg. J. Exp. Biol. 208, 439–445 (2005).
- S. H. Collins, P. G. Adamczyk, A. D. Kuo, Dynamic arm swinging in human walking. Proc. R. Soc. B Biol. Sci. 276, 3679–3688 (2009).
- J. M. Donelan, D. W. Shipman, R. Kram, A. D. Kuo, Mechanical and metabolic requirements for active lateral stabilization in human walking. *J. Biomech.* 37 (2004), pp. 827–835.
- 113. G. M. Bryan, P. W. Franks, S. C. Klein, R. J. Peuchen, S. H. Collins, A hip-knee-ankle exoskeleton emulator for studying gait assistance. *Int. J. Rob. Res.* **40**, 722–746 (2021).
- 114. D. Vega, C. J. Arellano, Using a simple rope-pulley system that mechanically couples the arms, legs, and treadmill reduces the metabolic cost of walking. *J. Neuroeng. Rehabil.* 18, 96 (2021).
- 115. S. A. Graham, C. P. Hurt, D. A. Brown, Minimizing postural demands of walking while still emphasizing locomotor force generation for nonimpaired individuals. *IEEE Trans. Neural Syst. Rehabil. Eng.* 26, 1003–1010 (2018).
- 116. C. A. Camillo, C. R. Osadnik, C. Burtin, S. Everaerts, M. Hornikx, H. Demeyer, M. Loeckx, F. M. Rodrigues, K. Maes, G. Gayan-Ramirez, W. Janssens, T. Troosters, Effects of downhill walking in pulmonary rehabilitation for patients with COPD: a randomised controlled trial. *Eur. Respir. J.* 56 (2020), doi:10.1183/13993003.00639-2020.
- 117. S. Carda, M. Invernizzi, A. Baricich, G. Cognolato, C. Cisari, Does altering inclination alter effectiveness of treadmill training for gait impairment after stroke? A randomized controlled trial. *Clin. Rehabil.* 27, 932–938 (2013).

- 118. Sloan M. Zimmerman, thesis, The Ohio State University (2016).
- J. Kerestes, T. G. Sugar, Enhanced Running Using a Jet Pack. Proc. ASME 2014 Int. Des. Eng. Tech. Conf. Comput. Inf. Eng. Conf., 1–7 (2014).
- M. Finn-Henry, A. Baimyshev, M. Goldfarb, in *Proceedings of the IEEE RAS and EMBS International Conference on Biomedical Robotics and Biomechatronics* (IEEE, 2020; https://ieeexplore.ieee.org/document/9224425/), vols. 2020-Novem, pp. 611–616.
- W. C. Miller, M. Speechley, B. Deathe, The prevalence and risk factors of falling and fear of falling among lower extremity amputees. *Arch. Phys. Med. Rehabil.* 82, 1031–1037 (2001).
- K. C. Bairapareddy, P. Jirange, K. Vaishali, in *9.2 Physiotherapists* (European Respiratory Society, 2016; http://erj.ersjournals.com/lookup/doi/10.1183/13993003.congress-2016.OA4820), p. OA4820.
- 123. M. F. Reelick, M. B. van Iersel, R. P. C. Kessels, M. G. M. Olde Rikkert, The influence of fear of falling on gait and balance in older people. *Age Ageing*. 38, 435–440 (2009).
- 124. A. Schinkel-Ivy, E. L. Inness, A. Mansfield, Relationships between fear of falling, balance confidence, and control of balance, gait, and reactive stepping in individuals with subacute stroke. *Gait Posture*. 43, 154–159 (2016).
- 125. T. G. Sugar, A. Bates, M. Holgate, J. Kerestes, M. Mignolet, P. New, R. K. Ramachandran, S. Redkar, C. Wheeler, Limit cycles to enhance human performance based on phase oscillators. *J. Mech. Robot.* 7 (2015), doi:10.1115/1.4029336.
- R. Matsuzaki, Y. Fujimoto, in *IECON Proceedings (Industrial Electronics Conference)* (IEEE, 2013; http://ieeexplore.ieee.org/document/6700220/), pp. 6581–6586.
- 127. D. Li, H. Vallery, in International Workshop on Advanced Motion Control, AMC (IEEE,

2012; http://ieeexplore.ieee.org/document/6197144/), pp. 1-6.

- 128. S. Jabeen, A. Berry, T. Geijtenbeek, J. Harlaar, H. Vallery, Assisting gait with free moments or joint moments on the swing leg. *IEEE Int. Conf. Rehabil. Robot.* 2019-June, 1079–1084 (2019).
- T. L. Brown, J. P. Schmiedeler, Reaction Wheel Actuation for Improving Planar Biped Walking Efficiency. *IEEE Trans. Robot.* 32, 1290–1297 (2016).
- 130. J. A. M. Haarman, E. Maartens, H. van der Kooij, J. H. Buurke, J. Reenalda, J. S. Rietman, Manual physical balance assistance of therapists during gait training of stroke survivors: characteristics and predicting the timing. *J. Neuroeng. Rehabil.* 14, 125 (2017).

Acknowledgements: 1 The authors would like to thank A. Harp and M. Fritton, and A. Dzewaltowski for assistance with 2 pilot testing and data processing; A. Skiadopoulos for assistance with calibrations and additional 3 analyses; and B. Senderling, T. Vanderheyden and HuMoTech for technical support. We would 4 also like to thank G. Lee, J. Kim, J. Caputo, K. Takahashi, N. Hunt, M. Donelan, and the members 5 6 of the SFU Locomotion Laboratory for suggestions for improving the manuscript and R. Kram for thoughtful conversations on the experiments. 7 8 9 **Funding:** This work was supported by the Center for Research in Human Movement Variability of the 10 University of Nebraska at Omaha, the National Institutes of Health grant number P20GM109090 11 to P.M., Nebraska EPSCOR First grant number OIA-1557417 to P.M., and a Graduate Research 12 and Creative Activity (GRACA) grant from the University of Nebraska at Omaha and an AMTI 13 14 Force and Motion Foundation award to P.A. This work received additional support from the National Institutes of Health grant numbers R01HD090333 to S.A.M., R01AG034995 and 15 R01AG049868 to I.I.P., and the United States Department of Veterans Affairs Rehabilitation 16 17 Research and Development Service grant number I01RX003266 to S.A.M. 18 19 **Author contributions:** 20 P.A. and P.M. conceived the study concept and experimental methods. A.M.G. conceived and implemented the control and pulley hardware. 21

- 22 P.A. and A.M.G. conducted the main experiments.
- 23 P.A., A.M.G., S.A.M., and I.I.P. conducted the supplementary analyses.

1	A.M.G. and P.M. conducted the simple pendulum model analyses.
2	P.A. and A.M.G. processed the data.
3	P.A. and P.M. prepared the manuscript.
4	All authors revised and approved the final manuscript.
5	
6	Competing interests:
7	S.A.M. serves on the advisory board and as a consultant for DigiTrans LLC. The other authors
8	declare that they have no competing interests.
9	
10	Data and materials availability:
11	The data supporting the main conclusions of the manuscript are included in the manuscript and
12	supplementary materials.
13	
14	Supplementary Materials:
15	Supplementary Materials and Methods
16	Supplementary Discussion
17	Figs. S1 to S13
18	Tables S1 to S6
19	List of Abbreviations, Terminology and Symbols
20	Movie S1. Experimental setup.
21	Data S1. 3D visualization of the pulley system.
22	Data S2. Controller code.
23	Data S3. Source data.

1 ccess and process the data.