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Human-in-the-loop optimization of a wearable robot using foot pressure sensors

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Abstract

**Background:** Individuals with below-knee amputation (BKA) experience increased physical effort when walking, and the use of a robotic ankle-foot prosthesis (AFP) can reduce such effort. Our prior study on a robotic AFP showed that walking effort could be reduced if the robot is personalized to the wearer. The personalization is accomplished using human-in-the-loop (HIL) optimization, in which the cost function is based on a real-time physiological signal indicating physical effort. The conventional physiological measurement, however, requires a long estimation time, hampering real-time optimization due to the limited experimental time budget. In addition, the physiological sensor, based on respiration uses a mask with rigid elements that may be difficult for the wearer to use. Prior studies suggest that a symmetry measure using a less intrusive sensor, namely foot pressure, could serve as a metric of gait performance. This study hypothesized that a function of foot pressure, the symmetric foot force-time integral, could be used as a cost function to rapidly estimate the physical effort of walking; therefore, it can be used to personalize assistance provided by a robotic ankle in a HIL optimization scheme.

**Methods:** We developed a new cost function derived from a well-known clinical measure, the symmetry index, by hypothesizing that foot force-time integral (FFTI) symmetry would be highly correlated with metabolic cost. We conducted experiments on human participants (N = 8) with simulated amputation to test the new cost function. The study consisted of a discrete trial day, an HIL optimization training day, and an HIL optimization data collection day. We used the discrete trial day to evaluate the correlation between metabolic cost and a cost function using symmetric FFTI percentage. During walking, we varied the prosthetic ankle stiffness while measuring foot pressure and metabolic rate. On the second and third days, HIL optimization was used to find the optimal stiffness parameter with the new cost function using symmetric FFTI percentage. Once the optimal stiffness parameter was found, we validated the performance with comparison to a weight-based stiffness and control-off conditions. We measured symmetric FFTI percentage during the stance phase, prosthesis push-off work, metabolic cost, and user comfort in each condition. We expected the optimized prosthetic ankle stiffness based on the newly developed cost function could reduce the energy expenditure during walking for the individuals with simulated amputation.

**Results:** We found that the cost function using symmetric foot force-time integral percentage presents a reasonable correlation with measured metabolic cost (Pearson’s R > 0.62). When we employed the new cost function in HIL ankle-foot prosthesis parameter optimization, 8 individuals with simulated amputation reduced their cost of walking by 15.9% (p = 0.01) and 16.1% (p = 0.02) compared to the weight-based and control-off conditions, respectively. The symmetric FFTI percentage for the optimal condition tended to be closer to the ideal symmetry value (50%) compared to weight-based (p = 0.23) and control-off conditions (p = 0.04).

**Conclusion:** This study suggests that foot force-time integral symmetry using foot pressure sensors can be used as a cost function when optimizing a wearable robot parameter.

**Keywords:** Human-in-the-loop; Foot pressure; Symmetry; Foot force-time integral; Bayesian Optimization; Prosthesis; Wearable robot
Background

A below-knee amputation (BKA) is one of the most common types of major amputation worldwide [1–3], yet it can be difficult to walk easily with an artificial limb [4–6]. As a result, individuals with BKA have expressed the desire for prosthetic devices that reduce the physical effort of walking [7]. One strategy to reduce physical effort is to prescribe a prosthetic foot with stiffness that is personalized to the wearer [8–11]. The stiffness is individually adjusted by clinical experts according to their observations, but this becomes difficult with an increasing number of prosthesis users [2]. The user’s body weight can also be used to adjust the stiffness [12–14]; however, recent studies suggest that a weight-based stiffness may not be the most metabolically economic [9, 14].

Human-in-the-loop (HIL) optimization has been developed for the task of identifying an optimal, personalized parameter that accounts for inter-subject variability in performance [15–19]. HIL optimization has been used to identify a user-specific assistance parameter in a wearable device and thus contributed to reducing physical effort during walking for healthy individuals [15, 19] and simulated amputees [9]. Individuals with BKA present higher inter-subject performance variability than able-bodied counterparts given assistance [20]. This increase in performance variability may be partially due to differences in residual limb tissue composition, geometry, and intended prosthetic components to be used distal to the socket [21]. Therefore, an individually tuned ankle-foot prosthesis through HIL optimization may improve assistance benefits by accounting for inter-subject performance variability.

In an HIL optimization scheme that uses metabolic cost as an indicator of physical effort, Bayesian optimization is used due to its sample-efficient and noise-tolerant characteristics [12, 15]. Bayesian optimization optimizes a posterior
distribution of metabolic cost over the control parameter space to minimize the user’s physical effort. The metabolic cost, used in the cost function, is the energy demand needed to perform a given task [22]. Its measurement, however, is challenging due to slow mitochondrial dynamics and noise in respiratory measurements [23]. As a result, it typically takes at least 5 min to obtain a reasonable estimate per testing condition. This estimation results in increased experimental time; thus, this optimization method has only been performed for walking and partial running [18] for a healthy individual. In addition, the respiratory measure for this metabolic cost estimation requires an uncomfortable and non-portable physiological sensor. These limitations have led to a search for alternative cost functions to be used with individuals with reduced physical strength, and they should be based on measures that are both time-efficient and comfortable.

Electromyography (EMG) has been used to estimate the metabolic cost of cycling [24] and joint moments [25], and this could be leveraged to estimate the metabolic cost of walking [13, 26]. However, the use of EMG demands exponentially higher computational costs [24]. For instance, the estimate of a joint moment with reinforcement learning [25] requires a relatively long learning time (a maximum of 6 hours), which limits the practical application of EMG on site. Another widely used cost function is a user-based subjective preference. Due to its subjective characteristics, this method often finds different optimum points for each trial, and the optimized assistance tends to show a low correlation with metabolic cost [27, 28].

Another candidate to measure the cost of walking is the ground reaction force. Ground reaction force has been used to identify gait characteristics such as deviations from the center of mass [12, 29, 30], gait symmetry [31], and the energy relation [32] in individuals with BKA [33]. Furthermore, healthy individuals present closer to symmetric ground reaction forces between the left and right limbs during walking [32], which is not the case for BKA [27]. When walking intervention is provided by an assistive device, individuals with BKA can improve their ground reaction force symmetry through evidence of improved joint kinematics symmetry [34], which has been proven to reduce walking effort [27] through a decrease in the energy needed to maintain balance [13]. This result suggests that gait symmetry using a parameter of ground reaction forces such as foot force-time integral (FFTI) [35, 36] can be used as an objective function for optimizing assistance and can serve as an alternative measure of the metabolic cost. Also, the FFTI information can be quickly obtained using a portable and comfortable foot pressure sensor [31], suggesting that the time to estimate the cost function could be reduced.

In this study, we hypothesized that the foot force-time integral (FFTI) symmetry could be used to estimate the physical effort of walking as a fast, portable and comfortable measure, and the cost function using the estimated effort can be used in a rapid human-in-the-loop (HIL) optimization scheme. To test this hypothesis, we developed a cost estimation method using the FFTI and evaluated the performance of the algorithm with individuals with simulated amputation [13, 37] using an ankle-foot prosthesis (AFP) emulator as an experimental platform. The cost estimation
method was employed in HIL Bayesian optimization of the AFP stiffness parameter to identify subject-specific personalized assistance. The optimized assistance was compared with the baseline conditions. We expect that the results of this study will inform follow-up experiments among individuals with amputation, eventually leading to the design of prosthetic limbs to reduce walking effort.

Methods
We performed an experiment to test the hypothesis that the physical effort can be estimated using foot force-time integral (FFTI) symmetry, and therefore a function of FFTI symmetry can be used to optimize assistance. We conducted walking experiments to evaluate the performance of the physical effort estimation method using FFTI symmetry. The performance of estimating effort with FFTI symmetry was evaluated by conducting a correlation analysis between the measured and estimated metabolic cost, and we assessed the performance of the optimized assistance using the new cost function using FFTI symmetry.

Prosthesis control

Hardware platform
We used a tethered robotic ankle-foot prosthesis emulator to permit real-time adjustments of free control parameters such as stiffness and net push-off energy [10, 31, 38, 39]. The device provided active plantarflexion torque as a function of ankle angle using the control parameters while users walked with the device (Fig. 1), as described in detail in [9, 14]. The rear part of the toe was connected to the two off-board servomotors (Caplex, Humotech, Pittsburgh, PA), which provided power. Control was performed using a real-time control system (Performance Real-Time Target Machine, Speedgoat, Switzerland). The emulator demonstrated performance of 250 Nm plantarflexion peak torque, more than 10 Hz control bandwidth, 17 Hz disturbance rejection bandwidth, and less than 5 Nm error on average in both plantarflexion and dorsiflexion. Those values are well within human ankle torque ranges during typical walking (120 Nm for plantarflexion) [40]. This characteristic enables our device to optimize free control parameters according to continuous biofeedback in the HIL optimization scheme. We also further improved the ankle-foot prosthesis emulator to meet the robustness demand of HIL optimization given an extended experiment duration [9].

Prosthesis controller with a free parameter
We developed a controller composed of low-, mid-, and high-level controllers with a free control parameter, namely stiffness (Fig. 2 A). The stiffness parameter was selected before an experiment to examine the effect of stiffness on the symmetric FFTI percentage and measured metabolic cost, or it was selected in real-time in the high-level controller during the HIL optimization. The stiffness parameter was used in the mid-level controller for generating a desired trajectory during the stance phase. The low-level controller then conducted linear control to track the desired trajectory.

The low-level controller provided linear torque and position control. The controller calculated the actuation command based on the error between the desired and actual
values of ankle angle position and torque. Subsequently, the calculated signals were sent to each Humotech Caplex actuator unit. The desired torque and position, as well as the control mode, were received from the mid-level controller (Fig. 2E,F).

The mid-level controller sent control commands, desired torque and position, and the control mode to the low-level controller based on gait mechanics. For walking, the gait mechanics were divided into a swing and stance phase. The stance phase was further divided into dorsiflexion and plantarflexion. Once there was transition toward dorsiflexion from the swing phase, torque control was enabled. During the stance phase, the desired ankle torque was commanded based on an ankle-angle torque curve (Fig. 2A). The curve was adjusted by manipulating the stiffness parameter, which influenced the shape and peak torque of the curve. The stiffness was a free control parameter, which was optimized with the high-level controller (Fig. 2D). The swing phase acted to enable position control by holding each motor at the desired position. The stiffness parameter, therefore, had no influence during the swing phase.

The high-level controller optimized a control parameter of the mid-level controller, such as the stiffness of the ankle-angle torque curve. In our control scheme, HIL Bayesian optimization performed this high-level control action (Fig. 2D). Bayesian optimization is a sequential design strategy for near-global optimization of a parameterized black-box function and it is a sample-efficient and noise-tolerant method [31, 37]. This method is well suited to optimizing objective functions, which are expensive to evaluate under constraints of noisy physiological signals with a limited time budget [10, 41, 42]. Bayesian optimization was used here to optimize a posterior distribution of estimated cost using the foot force-time integral over the control parameter space (Fig. 2C). In this case, the

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**Figure 2** Overview of the human-in-the-loop optimization using the symmetry cost function. (A) Mid-level controller: the desired ankle torque was generated depending on ankle angle and stiffness, commanded from the high-level controller. (B) Emulation system: the torque was delivered through the ankle-foot prosthesis emulator to a participant while collecting foot pressure and respiratory data. (C) Symmetric foot force-time integral (FFTI) cost function: the collected foot pressure was summed in the left (SP<sub>left</sub>) and right side (SP<sub>right</sub>) to estimate the cost of walking. (D) High-level controller: Bayesian optimization updated the stiffness parameter to minimize the estimated cost of walking. (E–H) Functional block diagram of the system: this block diagram describes how the system works in (A–D).
control parameter, $x$, is the stiffness, which alters the ankle torque curve in the mid-level controller (Fig. 2A).

**HIL Bayesian optimization**

The HIL Bayesian optimization was divided into two phases, initialization and optimization. We initialized the HIL Bayesian optimization by evaluating the estimated metabolic cost for three iterations, which correspond to pseudo-randomly chosen stiffness parameters in the range of 0.5 to 1.8 Nm/rad to avoid myopic sampling and premature convergence [12, 15, 43]. The upper and lower bounds of the stiffness condition were adjusted based on the safety torque limit.

After initialization, Bayesian optimization was iteratively performed over two steps using the estimated cost function (Fig. 2C): first estimating the posterior distribution of the cost as a function of ankle-foot prosthesis stiffness using a Gaussian process (Fig. 2D) [17, 44], and then selecting the next ankle-foot prosthesis stiffness, $x_{n+1}$, to evaluate using the expected improvement (Fig. 2D) [17]. This stiffness parameter was sent to the mid-level controller to regulate the torque as a function of ankle angle (Fig. 2A), and assistance was provided to a participant (Fig. 2B). Bayesian optimization was terminated if the experiment time budget, 15 min, was reached or three consecutive parameters were selected.

The Gaussian process calculated the estimated metabolic cost response surface, which is represented using the mean, $\mu_x$, the covariance, $k(x, x')$. As a standard practice [41], we used zero mean. For the covariance function, we selected a squared exponential kernel ($k(x_i, x_j)$) as shown in Eq. (1) [9, 12]:

$$k(x_i, x_j) = \sigma_f^2 \cdot e^{-\frac{(x_i - x_j)^2}{2l^2}}$$  \hspace{1cm} (1)

where, $\sigma_f^2$ is the signal variance of the estimated cost (estimated metabolic rate using foot force-time integral symmetry) variance, and $l$ is the length scale parameter (stiffness). The $\sigma_f$ and $l$ are hyperparameters, and we optimized the hyperparameters at each iteration to maximize the log marginal likelihood of the data, $D = x, y; x = [x_1 \ldots x_n]; y = [y_1 \ldots y_n]$.

The samples of estimated metabolic cost using foot force-time integral symmetry ($f(x)$) are assumed to have an additive, independent, and identically distributed noise,

$$y(x) = f(x) + \epsilon, \epsilon \sim N(0, \sigma_{noise}^2)$$

where $\sigma_{noise}^2$ is the noise variance and is a hyperparameter. Given the Gaussian process and data, $D$, the posterior estimated metabolic distribution was computed for a stiffness parameter, $x_*$, as $y(x_*) \equiv y^* \sim N(\mu_*, \sigma_*^2)$,

$$\mu_* = k_*^T (K + \sigma_{noise}^2 I)^{-1} y_{1:n}$$

$$\sigma_* = k(x_*, x_*) + k_*^T (K + \sigma_{noise}^2 I)^{-1} k_*$$  \hspace{1cm} (2)
where the $K$ and $k$ were calculated using:

$$k_s = [k(x_s, x_1), \ldots, k(x_s, x_n)]^T$$

$$K = \begin{pmatrix} k(x_1, x_1) & \cdots & k(x_1, x_n) \\ \vdots & \ddots & \vdots \\ k(x_n, x_1) & \cdots & k(x_n, x_n) \end{pmatrix}$$

where $x_s$ is discrete stiffness parameter, $x_1, \ldots, x_n$ are the stiffness parameter for previous $n$ iterations, and $y_1, \ldots, y_n$ are estimated metabolic cost using the foot force-time integral from the previous $n$ iterations.

To acquire the next stiffness parameter, we used the expected improvement (EI), which balanced between minimum predictive points and high uncertainty. EI selected the next parameter by calculating the expected reduction in the estimated metabolic cost over the stiffness previously evaluated using Gaussian process posterior distribution using Eq. (3):

$$EI[x_s] = (y_{best} - \mu_*) \cdot CDF(\mu_*) + \sigma_* \cdot PDF(u_*)$$

where $y_{best}$ is given as $\min_{1 \leq i \leq n} E[y(x_i)]$, $u_*$ is $(y_{best} - \mu_*)/\sigma_*$, and $CDF$ and $PDF$ corresponds to cumulative distribution function and probability distribution function of the posterior function (Gaussian process). EI value was set to zero when $\sigma_*$ was zero. The next parameter was then calculated using Eq. (4):

$$x_{n+1} = \arg\max_{x_s} (EI[x_s])$$

where the $\arg\max$ function identified stiffness corresponding to the maximum EI value in the parameter range ($x_s$). This newly selected parameter is then passed to the mid-level controller.

**New cost function to estimate metabolic cost**

We developed a cost function ($f(x)$) to be used in HIL Bayesian optimization based on the symmetry index [45]. Several symmetric indexes have been used with temporal, pressure, and force features [46]. The combination of the ground reaction force and time (force-time integral) had shown the lowest standard deviation compared to other temporal- and force-parameters-only methods [47]. In this study, we calculated the force-time integral using an F-scan insole pressure sensor (Tekscan, MI, USA). We first summed the pressure on each foot to estimate the foot force and then integrated the force through the stance phase. This force-time integral was then used to estimate the metabolic cost of walking.

The F-scan insole sensors were placed at the insole of the lift shoe (right) and the insole of the cast boot (left) (Fig. 1). Each F-scan sensor was connected to the port hub through an ethernet cable. We calibrated both the left and right sensors using the F-scan step calibration function. In this calibration, a subject was asked to stand on the opposite-side leg for 5 seconds and then switch to leg to be calibrated for a remaining 15 seconds. Using the pressure information from the sensor cells and a subject’s body weight, a calibration file was generated. The sensor has 25
sensel per square inch, and we replaced sensors when we saw a 10% drop in the sensing cells. For the real-time streaming, we used F-scan’s Matlab SDK to extract the magnitude of all the sensors in a single frame at 100Hz.

We first obtained the symmetric foot force-time integral (FFTI) percentage, focusing on the limb which is assisted by the ankle-foot prosthesis (AFP), left side:

\[
\text{FFTI} = \frac{SP_{\text{left}}}{SP_{\text{left}} + SP_{\text{right}}} \times 100
\]

where \(SP_{\text{left}}\) is the summed pressure in the left side and \(SP_{\text{right}}\) is the summed pressure in the right side. The sum of the pressure (\(SP\)) for each foot is obtained by adding the pressure in each cell over the stance phase of the gait. Similar to the force-time integral measure [47], \(SP\) captures the sum of force applied during the stance time.

Then, we constructed a cost function based on a symmetry index (SI) with a hypothesis that the metabolic cost would be minimized when a participant loaded equal force between the left and right feet during the stance phase [31, 45] as suggested by the simplified dynamic walking model [48]. Hence, we aim to minimize a function of symmetry index, \(SI = |SP_{\text{left}} - SP_{\text{right}}|/(0.5(SP_{\text{left}} + SP_{\text{right}}))\):

\[
f(x) = \alpha \cdot SI^2 + \beta = A(\text{FFTI} - 50)^2 + B
\]

where \(A = \alpha/100\), and \(B = \beta\). The detailed derivation can be found in the appendix.

**Experimental methods**

We conducted human subject experiments to evaluate the developed cost function by investigating the correlation between symmetric FFTI percentage and metabolic cost and then by employing the new cost function to optimize the ankle-foot prosthesis (AFP) stiffness parameter in human-in-the-loop (HIL) Bayesian optimization.

**Participants**

Eight healthy male adults (age 28.1 ± 3.3 years, weight 74.7 ± 9.1 kg, height 174.8 ± 6.8 cm) participated in this study. The experimental protocol was approved by the University of Illinois at Chicago Institutional Review Board. All subjects provided written informed consent in accordance with the Declaration of Helsinki.

For the experiment with individuals with simulated amputation, the ankle-foot prosthesis (AFP) end effector was modified by attaching a cast boot. The intent of walking similarly to amputation was simulated by immobilizing the non-amputated individual’s ankle, which effectively restricts the wearer’s ankle range of motion. The cast boot (Fig. 1) allowed a non-amputated individual with an intact lower extremity to safely interface with the AFP via a pyramidal adapter receptacle attached to the sole of the cast boot [37, 49, 50]. Subsequently, the non-amputated individual was raised above the opposite limb’s ground reference point, thus requiring the need to wear a lift shoe. The lift shoe is a boot consisting of an elevated sole manufactured from composite foam with a height of approximately 0.1 m.
**Experimental protocol**

Participants experienced three days of experimental protocol: discrete trials of eight AFP stiffness conditions, HIL optimization training, and HIL optimization data collection (Fig. 3). We provided an additional training day for the new three participants [51].

During the discrete trial day (Day1), participants first experienced a quiet standing condition for 3 min, which served as a baseline for the metabolic cost and foot pressure. Then, the subjects went through 8 stiffness conditions, 0.5 - 1.8 Nm/rad in a random order while they walked on a treadmill at a walking speed of 1.25 m/s for 5 min for each condition with a 5 min sitting break in between. If a participant expressed extreme discomfort and was almost unable to walk, we terminated the condition and excluded the stiffness during the following optimization.

On Day2, the participants experienced a HIL optimization training study to become familiarized with the long experimental time (15 min) while walking on a treadmill during multiple stiffness conditions without a break. After a minimum 24-hour rest, the subjects participated in the same protocol for the data collection on Day3.

For the HIL optimization training (Day2) and data collection days (Day3), participants initially experienced 3 min of standing to measure the base metabolic cost and foot pressure, followed by a 3 min sitting break. The HIL optimization started after 2 min of warm-up, and then the Bayesian optimization occurred over a maximum of 15 min while the subject walked on a treadmill at 1.25 m/s. The Bayesian optimization was said to be converged if three consecutive parameters were selected. After optimization, the participants experienced a 30 min sitting break. Then, the participants experienced the control-off condition, the weight-based condition, and the optimal condition in a random order for 5 min each. Subjects had a 5 min sitting break in-between each condition. For the control-off condition, we fixed the motor position; therefore, the participants walked while experiencing compliance from the Bowden cable. The weight-based condition provided assistance using a stiffness parameter based on the participant’s weight. For this experiment, we selected 1% of subject weight in
Figure 4 Cost function using symmetric foot force-time integral (FFTI) percentage evaluation results. (A, B) the correlation between FFTI symmetry cost and normalized measured metabolic cost at time intervals of 60-90s (A) and 210-270s (B). The measured and estimated cost using foot pressure symmetry (gray dots) was fitted with a linear regression (dark curve); also plotted are the confidence bounds of the data (light shade). For both time intervals, each dataset shows a statistically significantly high correlation.

kg/deg [52]. The control-off condition and the weight-based condition served as a baseline [53] to evaluate the performance of optimized assistance from HIL optimization using the foot force-time integral based cost function.

We collected respiratory rate (Cosmed, Rome, Italy), foot pressure (Tekscan, Boston, Massachusetts, USA), and user feedback to score comfort on a scale of 1 to 10, where 10 was the most comfortable score, and perceived effort on a scale of 6 to 20, where 20 was the highest rating of perceived effort [42, 54].

Data analysis

Foot force-time integral symmetry and metabolic cost

Using the data from the discrete trial day, we examined our hypothesis that symmetric gait, shown by symmetric foot force-time integral, could be used to minimize metabolic cost. Due to pressure sensor failure, two subjects’ pressure data were not used. We conducted a correlation analysis between estimated metabolic cost using the foot force-time integral and measured metabolic cost [55, 56]. We first calculated the symmetric foot force-time integral (FFTI) percentage using Eq. (5). The symmetric FFTI was further normalized with the min-max method to transform the data range from 0 to 1 (Fig. 4). Outliers were removed with the criterion of three standard deviations from the mean. Then, we obtained the symmetry cost using the Eq. (6). The measured, steady-state metabolic cost was calculated by taking the last 2 min data from the respiratory measure using the Brockway equation [57]. Then, we normalized the measured metabolic cost for each subject by subtracting the resting metabolic measure obtained from the standing condition and dividing it by the weight of the subject [12, 15, 58]. The measured metabolic cost was further normalized in the same manner as the estimated metabolic cost using symmetric FFTI by employing the min-max method and removing outliers.
The correlation between the estimated and measured metabolic costs was examined using a linear Pearson correlation analysis [59] (Fig. 4). We calculated the p-value and Pearson coefficient, $R$, for two different time intervals: 60-90s, and 210-270s. The 60-90s time period was chosen to represent the minimum possible adaptation period to update the symmetry cost. The 210-270s time period was chosen to indicate a consistent measure of the correlation between metabolic cost and symmetry cost throughout the 5 min walking period.

**HIL optimization using the new cost function**

We calculated the normalized steady-state metabolic cost, symmetric FFTI percentage, and net ankle push-off work for the optimized, weight-based, and control-off conditions from the validation trials. We used the steady-state metabolic cost [57], divided by body weight and with standing steady-state metabolic cost subtracted. The symmetric FFTI percentage was obtained using the Eq. (5). The net ankle push-off work was calculated by first thresholding the data to extract each step within the stance phase. An average of ankle torque and ankle angle was then taken for the total amount of extracted steps. The ankle push-off work was calculated using trapezoidal numerical integration of the ankle torque curve [44].

**Statistical analysis**

We compared the optimal condition to the control-off and the weight-based conditions for the metabolic cost, symmetric FFTI, and net push-off work. We first tested normality tests with the Kolmogorov–Smirnov test. If normality was confirmed, we conducted the paired t-test. The significance levels for statistical analyses were defined at $p < 0.05$.

**Results**

The measured and predicted metabolic cost showed a statistically significantly high correlation ($R = 0.64$ for time interval 60-90 s and $R = 0.63$ for 210-270 s, $p < 0.001$) (Fig. 4).

The optimal personalized stiffness condition reduced the metabolic cost by 15.9% and 16.1% compared to the weight-based condition and the control-off condition, respectively (paired t-test, $p < 0.02$) (Fig. 5A).

The summed foot pressure was more symmetric for the optimal condition (symmetric foot force-time integral (FFTI) percentage $= 47.6\% \pm 8.4\%$), compared to the control off condition (symmetric FFTI percentage $= 43.2\% \pm 5.5\%$) (paired t-test, $p = 0.042$) (Fig. 5B). The optimal condition also appeared to increase the symmetry of the foot force-time integral compared to the weight-based condition (symmetric FFTI percentage $= 44.8\% \pm 8.7\%$), but it was not statistically significant (paired t-test, $p = 0.231$).

The net push-off work energy tended to be maintained for the optimal and weight-based conditions (paired t-test, $p = 0.618$). The mean values for push-off work were $-0.01 \pm 0.05$ and $0.01 \pm 0.07 \, J/kg$ for the weight-based and the optimal conditions, respectively.

Bayesian optimization identified the subject-specific optimal parameters between 0.5 and 1.8 $Nm/rad$ which was scaled between 1 and 100 (Fig. 5C,D). The
optimization ran for approximately 10 iterations and converged within 15 min. For a standard HIL optimization using metabolic cost, the optimization would take a minimum of 24 min, including exploration periods [9, 12, 15, 18, 60].

**Discussion**

In this study, we developed a new cost function using a portable and comfortable mechanical sensor, namely a foot pressure sensor, for use in a human-in-the-loop (HIL) optimization. The cost function using the foot force-time integral (FFTI) also has an important benefit: fast cost estimation. When this cost function was used in the HIL optimization, the optimal stiffness parameter was found within 15 min. The optimized assistance resulted in a more symmetric FFTI and reduced the metabolic cost of walking (Fig. 5A). The optimal stiffness parameter was varied depending on the participant, and each participant presented a
subject-specific estimated metabolic cost response surface (Fig. 5C, D). The FFTI-based cost estimation requires a shorter time to find an optimal condition; hence, it has the great potential to enable the application of this personalization method to individuals with limited physical strength. Also, with the portable measurement feature, HIL optimization using a pressure sensor can be used in a natural environment setting.

The optimized assistance found using the symmetric FFTI helped reduce the metabolic cost of walking. Evidence suggests that an asymmetrical gait caused by varying step length or step frequency could be energetically less optimal [48, 61]. In addition, such a gait may result in failure to stabilize the body during the transition between the swing phase and stance phases [62]. Body stabilization to mitigate asymmetrical gait has been shown to increase metabolic cost [32, 63, 64]. For example, when participants experienced asymmetrical gait while walking on a split-belt treadmill where each belt moved at a different speed, the participants showed higher metabolic cost compared to symmetrical gait as this necessitated changing the step length of each foot. Similarly, our results showed that when participants had symmetrical gait while walking with a wearable robot (i.e., symmetric FFTI percentage when walking as a result of optimal stiffness), the metabolic cost was statistically significantly lower than with asymmetrical walking.

Our optimization result suggests that HIL optimization using foot pressure can be applied to individuals with amputation as it can find the optimal stiffness in a relatively short time. People with neurological diseases (e.g., stroke, Parkinson’s disease, multiple sclerosis, etc.) and amputation typically present impaired gait and robotic exoskeletons have been developed to improve their gait performance [27, 51, 65]. In particular, personalized assistance has been developed to improve gait performance using HIL optimization based on metabolic cost [15, 18, 19]. The measurement of the metabolic cost requires approximately 5 min of walking for each condition due to noise and slow mitochondrial dynamics [23]. This estimation results in increased experimental time; thus, the optimization method has only been performed for walking and partial running [18] by an able-bodied counterpart and was limited regarding the application of patients (e.g., individuals with neurological diseases or amputation having reduced physical strength). The fast estimation and optimization of our study may enable the use of HIL optimization to the individuals with reduced physical strength.

We hypothesized that the symmetric foot force-time integral would result in metabolically efficient walking. It is possible that a slightly asymmetric gait is energetically optimal [66]. One participant may have an intrinsically asymmetric gait due to long term adaptations to their particular bio-mechanics [67]. With our newly developed symmetry cost function, we could shift our ideal location of the global minimum to suit an individual’s intrinsic asymmetrical gait. Future work could explore these intrinsic properties to see how an individual’s neural system may adapt to a perfectly symmetrical assistive condition while walking with an ankle exoskeleton and whether providing a slightly asymmetric condition as the optimal parameter could further reduce the metabolic cost of walking.

Further studies are required to test the applicability of symmetric pressure optimization to individuals with amputation. There are numerous differences
between individuals with simulated amputation and those with below-knee amputation, such as the training duration for wearing prostheses and sensory-motor control pathways. Perhaps due to these differences, researchers previously observed different outcomes between these populations [49, 65]. For the purpose of our study, these differences become less concerning because similar to a previous study [12], we used the same factors of mass, height, and alignment, and these factors are unlikely to interact with the summed foot pressure symmetry cost function. While our results are promising, an experiment needs to be conducted with individuals with amputation to draw a conclusion regarding the effect of this HIL optimization using foot force-time integral symmetry. In addition, the investigation of our cost function using foot force-time integral symmetry may also provide insight into the user adaptation to the device and may lead to an efficient method for considering human-robot co-adaptation.

Conclusion

By embedding foot pressure sensors into the HIL optimization, we were able to develop a new cost function to estimate the metabolic energy expenditure of walking. We developed the symmetric foot force-time integral (FFTI) percentage, considering that absolute symmetry may be the most energetically optimal for individuals while walking. The symmetry cost calculated using symmetric FFTI percentage yielded a high correlation with metabolic cost, which confirms that walking experiments may now explore a more comfortable and less intrusive measure for the cost of walking. Effectively, FFTI is also a pseudo measure for ground reaction forces and other gait characteristics; therefore, the potential to further explore the evaluation of walking optimization with varying cost measurements may be asserted. Another benefit of HIL optimization with the use of foot pressure is the rapid estimation time (60–90s), which is especially important for individuals with below-knee amputation (BKA) during walking as they are more prone to fatigue than individuals without BKA. The fast estimation time may eventually lead to a more comfortable and fatigue-tolerant procedure to optimize prosthesis parameters for individuals with BKA. Future work should investigate whether our newly developed symmetry cost function can be effectively used to reduce the metabolic cost of walking for individuals with BKA.

Appendix

Derivation of the cost function using symmetric index

The symmetric index for walking is given as follows [45]:

$$SI = \frac{|X_l - X_r|}{\frac{1}{2}(X_l + X_r)}$$

where X is a gait-related parameter such as stance time, step length, or foot force-time integral (FFTI). We obtained the FFTI using the sum of pressure (SI) during the stance phase [45]:

$$SI = \frac{|SP_{left} - SP_{right}|}{\frac{1}{2}(SP_{left} + SP_{right})}$$
where \( SP_{\text{left}} \) and \( SP_{\text{right}} \) are the sum of pressure of the left and right foot, respectively. Then, we obtained a cost function by squaring each side and multiplying and adding constants [45]:

\[
f(x) = \alpha \cdot 4 \cdot \left( \frac{|SP_{\text{left}} - SP_{\text{right}}|}{SP_{\text{left}} + SP_{\text{right}}} \right)^2 + \beta
\]

Simplifying further,

\[
f(x) = \alpha \cdot 4 \cdot \left( \frac{SP_{\text{left}}}{SP_{\text{left}} + SP_{\text{right}}} - \left( 1 - \frac{SP_{\text{left}}}{SP_{\text{left}} + SP_{\text{right}}} \right) \right)^2 + \beta
\]

\[
= A \cdot \left( \frac{SP_{\text{left}}}{SP_{\text{left}} + SP_{\text{right}}} - 50 \right)^2 + B
\]

where, \( A \), \( B \) are the coefficients of the function defined by \( A = \frac{\alpha}{100} \), \( B = \beta \). We obtained the coefficients \( A \) and \( B \) to minimize the mean square error between the measured metabolic cost from the respiratory measure and estimated metabolic cost using symmetric FFTI percentage data.

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Abbreviations
AFP: Ankle Foot prothesis
BKA: Below-Knee Amputation
EMG: Electromyography
FFTI: Foot Force-Time Integral
HIL: Human-in-the-loop optimization
SDK: Software Development Kit
SI: Symmetric Index
SP: Sum of Pressure

Availability of data and materials
Data and code will made available by the corresponding author upon reasonable request.

Ethics approval and consent to participate
The authors affirm that human research participants provided informed consent for publication under University of Illinois at Chicago, IRB-2019-0087.

Competing interests
The authors declare that they have no competing interests.

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Authors’ contributions
M.K., J.R., J.P., and H.C. conceived and oversaw the study. M.J., P.K., and H.J. performed the data collection, M.J., P.K., and X.Z. analyzed the data and M.K., M.J., and P.K. prepared the manuscript. All authors proofread the manuscript and approved the final manuscript.

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### References

1. Nigg, S., Vienneau, J., Maurer, C., Nigg, B.M.: Development of a symmetry index using discrete variables. Gait & Posture 38(1), 115–119 (2013). doi:10.1016/j.gaitpost.2012.10.024

2. Lener A, Soundry M: Armed Conflict Injuries to the Extremities. Armed Conflict Injuries to the Extremities (2011). doi:10.1007/978-3-642-16155-1

3. Ziegler-Graham, K., MacKenzie, E.J., Ephraim, P.L., Travison, T.G., Brookmeyer, R.: Estimating the Prevalence of Limb Loss in the United States: 2005 to 2050. Archives of Physical Medicine and Rehabilitation 89(3), 422–429 (2008). doi:10.1016/j.apmr.2007.11.005

4. Pagliarulo, M.A., Waters, R., Hlosp, H.J.: Energy Cost of Walking of Below-Knee Amputees Having No Vascular Disease. Physical Therapy 59(5), 538–542 (1979). doi:10.1093/PTJ/59.5.538

5. Silverman, A.K., Neptune, R.R.: Differences in whole-body angular momentum between below-knee amputees and non-amputees across walking speeds. Journal of biomechanics 44(3), 379–385 (2011). doi:10.1016/J.JBIOMECH.2010.10.027

6. Whyte, A.S., Carroll, L.J.: A preliminary examination of the relationship between employment, pain and disability in an amputee population. Disability and rehabilitation 24(9), 462–470 (2002). doi:10.1080/09638280110105213

7. Legro, M.W., Reiber, G., Michael Del Aguil, ., Ajax, M.J., Boone, D.A., Larsen, J.A., Smith, D.G., Sangeorzan, B.: Issues of importance reported by persons with lower limb amputations and prostheses. Technical Report 3 (1999)

8. Fey, N.P., Klute G. K., Neptune, R.R.: Optimization of prosthetic foot stiffness to reduce metabolic cost and intact knee loading during below-knee amputee walking: a theoretical study 134, 111005 (2012)

9. Tin-Chun Wen Xingyuan Zhou, M.J.S.S.M.K.: Development of a Bayesian Optimization Controller to Minimize the Metabolic Cost of Walking for Human-in-the-Loop Optimization of Ankle-Foot Prosthesis (AFP) Stiffness. In: in Review

10. Shepherd, M.K., Azocar, A.F., Major, M.J., Rouse, E.J.: Amputee perception of prosthetic ankle stiffness during locomotion. Journal of neuroengineering and rehabilitation 15(1), 99 (2018)

11. Witte, K.A., Fiers, P., Sheets-Singer, A.L., Collins, S.H.: Improving the energy economy of human running with powered and unpowered ankle exoskeleton assistance. Science Robotics 5(40), 9180 (2020). doi:10.1126/scirobotics.aaay1018

12. Kim, M., Ding, Y., Malcolm, P., Speeckaert, J., Siivy, C.J., Walsh, C.J., Kuindersma, S.: Human-in-the-loop Bayesian optimization of wearable device parameters. PLoS ONE 12(9), 6–8 (2017). doi:10.1371/journal.pone.0184054

13. Kim, M., Collins, S.H.: Once-per-step control of ankle-foot prosthesis push-off work reduces effort associated with balance during walking 12(1), 43 (2015)

14. Kim, M., Chen, T., Chen, T., Collins, S.H.: An ankle-foot prosthesis emulator with control of plantarflexion and inversion-eversion torque. IEEE Transactions on Robotics 34(5), 1183–1194 (2018)

15. Ding, Y., Kim, M., Kuindersma, S., Walsh, C.J.: Human-in-the-loop Bayesian optimization of wearable device parameters. PLoS ONE 12(9), 6–8 (2017). doi:10.1371/journal.pone.0184054

16. Felt, W., Selinger, J.C., Donelan, J.M., Remy, C.D.: “Body-in-the-loop”: Optimizing device parameters using measures of instantaneous energetic cost. PLoS ONE 10(8), 1–21 (2015). doi:10.1371/journal.pone.0135342

17. Kim, M., Liu, C., Kim, J., Lee, S., Meguid, A., Walsh, C.J., Kuindersma, S.: Bayesian Optimization of Soft Exosuits Using a Metabolic Estimator Stopping Process. International Conference on Robotics and Automation, 9173–9179 (2019)

18. Zhang, J., Fiers, P., Witte, K.A., Jackson, R.W., Poggensee, K.L., Atkeson, C.G., Collins, S.H.: Human-in-the-loop optimization of exoskeleton assistance during walking. Science 356(6344), 1280–1283 (2017). doi:10.1126/science.aal5054

19. Zhang, X., Kamgarpour, M., Georgiou, A., Goulart, P., Lygeros, J.: Robust optimal control with adjustable uncertainty sets. Automatica 75, 249–259 (2017). doi:10.1016/j.automatica.2016.09.016

20. Quesada, R.E., Caputo, J.M., Collins, S.H.: Increasing ankle push-off work with a powered prosthesis does not necessarily reduce metabolic rate for transfemoral amputees. 49(14), 3452–3459 (2016)

21. Petron, A., Duval, J.F., Herr, H.: Multi-indentener device for in vivo biomechanical tissue measurement. IEEE Transactions on Neural Systems and Rehabilitation Engineering 25(5), 426–435 (2017). doi:10.1109/TNSRE.2016.2572168

22. Givoni, B., Goldman, R.F.: Predicting metabolic energy cost. Journal of applied physiology 30(3), 424–433 (1971). doi:10.1152/jappl.1971.30.3.429

23. Wai, T., Langer, T.: Mitochondrial Dynamics and Metabolic Regulation. Elsevier Inc. (2016). doi:10.1016/j.tem.2015.12.001

24. Blake, O.M., Wakeling, J.M.: Estimating changes in metabolic power from EMG. SpringerPlus 2(1), 1–7 (2013). doi:10.1186/2193-1801-2-229

25. Wu, W., Saul, K., Huang, H.H.: Using Reinforcement Learning to Estimate Human Joint Moments via EMG Signals or Joint Kinematics: An Alternative Solution to Musculoskeletal-Based Biomechanics. Journal of Biomechanical Engineering (2020). doi:10.1115/1.4049333

26. Koelweijn, A.D., Heinrich, D., van den Bogert, A.J.: Metabolic cost calculations of gait using musculoskeletal energy models, a comparison study. PLOS ONE 14(9), 0222037 (2019). doi:10.1371/journal.pone.0222037

27. Mattes, S.J., Martin, P.E., Royer, T.D.: Walking symmetry and energy cost in persons with unilateral transfemoral amputations: Matching prosthetic and intact limb inertial properties. Archives of Physical Medicine and Rehabilitation 81(5), 561–568 (2000). doi:10.1053/mr.2000.3851

28. Welker, G., Voloshina, A.S., Chiu, V.L., Collins, S.H.: Shortcomings of human-in-the-loop optimization for an ankle-foot prosthesis: a case study. bioRxiv, 2020–1017343970 (2020). doi:10.1101/2020.10.17.343970

29. Kuo, A.D.: Stabilization of lateral motion in passive dynamic walking 18(9), 917–930 (1999)

30. Kuo, A.D.: A simple model predicts the step length-speed relationship in human walking 123, 264–269 (2001)
31. Su, B.L., Song, R., Guo, L.Y., Yen, C.W.: Characterizing gait asymmetry via frequency sub-band components of the ground reaction force. Biomedical Signal Processing and Control 18, 56–60 (2015). doi:10.1016/j.bspc.2014.11.008
32. Ellis, R.G., Howard, K.C., Kram, R.: The metabolic and mechanical costs of step time asymmetry in walking. Proceedings of the Royal Society B: Biological Sciences 280(1756) (2013). doi:10.1098/rspb.2012.2784
33. Seliktar, R., Harwin, W.S.: Some gait characteristics of below-knee amputees and their reflection on the ground reaction forces. Engineering in Medicine 15(1), 27–34 (1986). doi:10.1243/EMED_JOUR_1986_0
34. Kaufman, K.R., Frittole, S., Frigo, C.A.: Gait asymmetry of transfemoral amputees using mechanical and microprocessor-controlled prosthetic knees. Clinical Biomechanics 27(5), 460–465 (2012). doi:10.1016/j.clinbiomech.2011.11.011
35. Vitezova, S., Kutilek, P., Svoboda, Z., Szabo, Z.: Gait symmetry measures: A review of current and prospective methods. Elsevier Ltd (2018). doi:10.1016/j.bspc.2018.01.013
36. Wearing, S.C., Smeethers, J.E., Urry, S.R.: The effect of plantar fasciitis on vertical foot-ground reaction force. Clinical Orthopaedics and Related Research (409), 175–185 (2003). doi:10.1097/01.blo.0000057989.41099.d8
37. Caputo, J.M., Collins, S.H.C.: A universal ankle-foot prosthesis emulator for experiments during human locomotion 136, 35002 (2014)
38. Kim, M., Collins, S.H.: Step-to-step ankle inversion/eversion torque modulation can reduce effort associated with balance. Frontiers in Neurorobotics 11(NOV) (2017). doi:10.3389/fnbot.2017.00062
39. Caputo, J.M., Collins, S.H.: The effect of ankle-foot prosthesis push-off work on walking kinetics and overall effort. In: World Congress of Biomechanics, p. 1 (2014)
40. Hunt, A.E., Smith, R.M., Torode, M.: Extrinsic Muscle Activity, Foot Motion and Ankle Joint Moments During the Stance Phase of Walking 22(1), 31–41 (2001)
41. Brochu, V.M.C., De Freitas, N.: A Tutorial on Bayesian Optimization of Expensive Cost Functions, with Application to Active User Modeling and Hierarchical Reinforcement Learning (2010)
42. Fang, Y., Lerner, Z.F.: Feasibility of Augmenting Ankle Exoskeleton Walking Performance With Step Length Manipulation on gait symmetry during walking. Journal of NeuroEngineering and Rehabilitation 12(1) (2015). doi:10.1186/s12984-015-0027-3
43. Hunt, A.E., Smith, R.M., Torode, M.: Extrinsic Muscle Activity, Foot Motion and Ankle Joint Moments During the Stance Phase of Walking 22(1), 31–41 (2001)
44. Kushner, H.J.: A new method of locating the maximum point of an arbitrary multipeak curve in the presence of noise. B6(1), 97–106 (1964)
45. Robinson, R., Herzog, W., Nigg, B.: Use of force platform variables to quantify the effects of chiropractic manipulation on gait symmetry. undefined (1987)
46. Rosenbaum, D., Becker, H.-p., Rosenbaum, D., Wilhelms-, W.: Planter pressure distribution measurements. Technical background and clinical applications. Foot and Ankle Surgery 3(1), 1–14 (1997). doi:10.1064/J.1460-9584.1997.00043.X
47. Herzog, W., Nigg, B.M., Read, L.J., Olsson, E.: Asymmetries in ground reaction force patterns in normal human gait. Medicine and science in sports and exercise 21(1), 110–114 (1989). doi:10.1249/00005768-198902000-00020
48. Kuo, A.D., Donelan, J.M., Ruina, A.: Energetic consequences of walking like an inverted pendulum: step-to-step transitions 33, 66–97 (2005)
49. Zelik, K.E., Collins, S.H., Adamczyk, P.G., Segal, A.D., Klute, G.K., Morgenroth, D.C., Hahn, M.E., Orendurff, M.S., Czerniecki, J.M., Kuo, A.D.: Systematic variation of prosthetic foot parameter affects center-of-mass mechanics and metabolic cost during walking 19, 411–419 (2011)
50. Scott VanZant, R., McPoil, T.G., Cornwall, M.W.: Symmetry of Plantar Pressures and Vertical Forces in Healthy Subjects During Walking. Journal of the American Podiatric Medical Association 91(7), 337–342 (2001). doi:10.7547/87507315-91-7-337
51. Zelik, K.E., Collins, S.H., Adamczyk, P.G., Segal, A.D., Klute, G.K., Morgenroth, D.C., Hahn, M.E., Orendurff, M.S., Czerniecki, J.M.: The effects of a controlled energy storage and return prototype prosthetic foot on transtibial amputee ambulation 31, 918–931 (2012)
60. Selinger, J.C., O’Connor, S.M., Wong, J.D., Donelan, J.M.: Humans Can Continuously Optimize Energetic Cost during Walking. Current Biology 25(18), 2452–2456 (2015). doi:10.1016/j.cub.2015.08.016

61. Sánchez, N., Simha, S.N., Donelan, J.M., Finley, J.M.: Taking advantage of external mechanical work to reduce metabolic cost: the mechanics and energetics of split-belt treadmill walking. The Journal of physiology 597(15), 4053–4068 (2019). doi:10.1113/JP277725

62. Zulkifli, S.S., Loh, W.P.: A state-of-the-art review of foot pressure. Foot and ankle surgery : official journal of the European Society of Foot and Ankle Surgeons 26(1), 25–32 (2020). doi:10.1016/J.FAS.2018.12.005

63. Roper, J.A., Stegemöller, E.L., Tillman, M.D., Hass, C.J.: Oxygen consumption, oxygen cost, heart rate, and perceived effort during split-belt treadmill walking in young healthy adults. European journal of applied physiology 113(3), 729–734 (2013). doi:10.1007/S00421-012-2477-7

64. Sánchez, N., Simha, S.N., Donelan, J.M., Finley, J.M.: Taking advantage of external mechanical work to reduce metabolic cost: the mechanics and energetics of split-belt treadmill walking. bioRxiv (2018). doi:10.1101/500835

65. Kim, M.: Ankle controller design for robotic ankle-foot prostheses to reduce balance-related effort during walking using a dynamic walking approach. PhD thesis, Carnegie Mellon University (2015)

66. McCain, E.M., Berno, M.E., Libera, T.L., Lewek, M.D., Sawicki, G.S., Saul, K.R.: Reduced joint motion supersedes asymmetry in explaining increased metabolic demand during walking with mechanical restriction. Journal of Biomechanics 126, 110621 (2021). doi:10.1016/J.JBIOMECH.2021.110621

67. Seminati, E., Nardello, F., Zamparo, P., Ardigo, L.P., Fazioli, N., Minetti, A.E.: Anatomically Asymmetrical Runners Move More Asymmetrically at the Same Metabolic Cost. PLOS ONE 8(9), 74134 (2013). doi:10.1371/JOURNAL.PONE.0074134