

Open access • Posted Content • DOI:10.1101/2020.06.18.105163

# Predicting walking response to ankle exoskeletons using data-driven models — Source link [2]

Michael C. Rosenberg, Bora S. Banjanin, Samuel A. Burden, Katherine M. Steele Institutions: University of Washington Published on: 25 Aug 2020 - bioRxiv (Cold Spring Harbor Laboratory)

Topics: Exoskeleton and Gait (human)

#### Related papers:

- Predicting walking response to ankle exoskeletons using data-driven models.
- Ankle Joint Torque Estimation Using an EMG-Driven Neuromusculoskeletal Model and an Artificial Neural Network
   Model
- A PID Controller Approach to Explain Human Ankle Biomechanics across Walking Speeds
- · Predictive Simulations of Gait with Exoskeletons that Alter Energetics
- Testing Simulated Assistance Strategies on a Hip-Knee-Ankle Exoskeleton: a Case Study



1

# Predicting walking response to ankle exoskeletons using data-driven models

<sup>3</sup> Michael C. Rosenberg<sup>1</sup>, Bora S. Banjanin<sup>2</sup>, Samuel A. Burden<sup>2</sup>, Katherine M. Steele<sup>1</sup>

<sup>4</sup> <sup>1</sup> Department of Mechanical Engineering, University of Washington, Seattle, WA, USA

5 <sup>2</sup> Department of Electrical and Computer Engineering, University of Washington, Seattle, WA, USA

- 6 Correspondence: mcrosenb@uw.edu
- 7

1

2

8 I. KEYWORDS

9 Ankle exoskeleton; data-driven modeling; locomotion; prediction; joint kinematics; muscle activity

- 10
- 11 II. Abstract

12 Despite recent innovations in exoskeleton design and control, predicting subject-specific impacts of 13 exoskeletons on gait remains challenging. We evaluated the ability of three classes of subject-specific phase-14 varying models to predict kinematic and myoelectric responses to ankle exoskeletons during walking, without 15 requiring prior knowledge of specific user characteristics. Each model – phase-varying (PV), linear phasevarying (LPV), and nonlinear phase-varying (NPV) - leveraged Floquet Theory to predict deviations from a 16 17 nominal gait cycle due to exoskeleton torque, though the models differed in complexity and expected prediction accuracy. For twelve unimpaired adults walking with bilateral passive ankle exoskeletons, we 18 19 predicted kinematics and muscle activity in response to three exoskeleton torque conditions. The LPV 20 model's predictions were more accurate than the PV model when predicting less than 12.5% of a stride in the future and explained 49–70% of the variance in hip, knee, and ankle kinematic responses to torque. The 21 22 LPV model also predicted kinematic responses with similar accuracy to the more-complex NPV model. Myoelectric responses were challenging to predict with all models, explaining at most 10% of the variance 23 in responses. This work highlights the potential of data-driven phase-varying models to predict complex 24 25 subject-specific responses to ankle exoskeletons and inform device design and control.

#### 26 III. INTRODUCTION

Ankle exoskeletons are used to improve kinematics and reduce the energetic demands of locomotion in 27 unimpaired adults and individuals with neurologic injuries [1-5]. Customizing exoskeleton properties to 28 improve an individual's gait is challenging and accelerating the iterative experimental process of device 29 optimization is an active area of research [6, 7]. Studies examining the effects of exoskeleton properties – 30 sagittal-plane ankle stiffness or equilibrium ankle angle for passive exoskeletons and torque control laws for 31 powered exoskeletons – on kinematics, motor control, and energetics have developed design and control 32 33 principles to reduce the energetic demand of walking and improve the quality of gait [1, 6, 8, 9]. Predicting how an individual's gait pattern responds to ankle exoskeletons across stance may inform exoskeleton design 34 by enabling rapid evaluation of exoskeleton properties not tested experimentally. Additionally for powered 35 exoskeletons, which prescribe torque profiles using feedforward or feedback (e.g. kinematic or myoelectric) 36 control laws, predicting responses over even 10–20% of a stride may improve tracking performance or 37 transitions between control modes [4, 10-12]. However, predicting subject-specific responses to exoskeletons 38 remains challenging for unimpaired individuals and those with motor impairments [2, 12, 13]. 39

40

41 Common physics-based models, including simple mechanical models and more physiologically-detailed musculoskeletal models, use principles from physics and biology to analyze and predict exoskeleton impacts 42 on gait. For example, one lower-limb mechanical walking model predicted that an intermediate stiffness in a 43 passive exoskeleton would minimize the energy required to walk, a finding that was later observed 44 experimentally in unimpaired adults [1, 14]. More physiologically-detailed musculoskeletal models have 45 been used to predict the impacts of exoskeleton design on muscle activity during walking in children with 46 cerebral palsy and running in unimpaired adults [15, 16]. While these studies identified hypothetical 47 relationships between kinematics and the myoelectric impacts of exoskeleton design parameters, their 48 49 predictions were not evaluated against experimental data.

Challenges to accurately predicting responses to ankle exoskeletons with physics-based models largely stem 51 from uncertainty in adaptation, musculoskeletal physiology, and motor control, which vary between 52 individuals and influence response to exoskeletons. While individuals explore different gait patterns to 53 54 identify an energetically-optimal gait, exploration does not always occur spontaneously, resulting in sub-55 optimal gait patterns for some users [17]. Popular physiologically-detailed models of human gait typically assume instantaneous and optimal adaptation, which do not reflect how experience and exploration may 56 influence responses to exoskeletons, possibly reducing the accuracy of predicted responses [18, 19]. 57 58 Additionally, when specific measurement sets are unavailable for model parameter tuning, population-59 average based assumptions about musculoskeletal properties and motor control are required [17, 20-22]. However, musculoskeletal properties and motor control are highly uncertain for individuals with motor 60 61 impairments, today's most ubiquitous ankle exoskeleton users [19, 20, 22, 23]. Musculotendon dynamics and motor complexity are known to explain unintuitive exoskeleton impacts on gait energetics, suggesting that 62 uncertain musculotendon parameters and motor control may limit the accuracy of predicted changes in gait 63 with ankle exoskeletons [19, 21, 24]. Predictions of exoskeleton impacts on gait using physiological models, 64 65 therefore, require accurate estimates of adaptation, musculotendon parameters, and motor control.

66

67 Conversely, data-driven approaches address uncertainty in user-exoskeleton dynamics by representing the system entirely from experimental data. For instance, human-in-the-loop optimization provides a model-free 68 69 alternative to physics-based prediction of exoskeleton responses by automatically exploring different 70 exoskeleton torque control strategies for an individual [6, 7]. This experimental approach requires no prior 71 knowledge about the individual: optimization frameworks identify torque control laws that decrease 72 metabolic rate relative to baseline for an individual using only respiratory data and exoskeleton torque measurements. However, experimental approaches to exoskeleton optimization require the optimal design to 73 be tested, potentially making the search for optimal device parameters time-intensive. Alternatively, machine 74 75 learning algorithms, such as the Random Forest Algorithm, have used retrospective gait analysis and clinical

exam data to predict changes in joint kinematics in response to different ankle-foot orthosis designs in 76 children with cerebral palsy [8]. This study reported good classification accuracy, though predictions may 77 not generalize to new orthosis designs. Unlike physiologically-detailed or physics-based models, human-in-78 79 the-loop optimization and many machine learning models are challenging to interpret, limiting insight into 80 how a specific individual's physiology influences response to exoskeleton torque. A balance between physiologically-detailed and model-free or black-box data-driven approaches may facilitate the prediction 81 and analysis of responses to ankle exoskeletons without requiring extensive knowledge of an individual's 82 83 physiology.

84

In this work, we investigated a subject-specific data-driven modeling framework – phase-varying models – that may fill the gap between physiologically-detailed model-based and model-free experimental approaches for predicting gait with exoskeletons. Phase-varying models typically have linear structure whose parameters are estimated from data, enabling both prediction and analysis of gait with exoskeletons [25, 26]. Unlike physiologically-detailed models, phase-varying models do not require knowledge of the physics or control of the underlying system. Unlike experimental approaches, the model-based framework enables prediction of responses to untested exoskeleton designs or control laws.

92

93 Phase-varying models leverage dynamical properties of stable gaits derived from Floquet Theory, which 94 ensures that the convergence of a perturbed trajectory to a stable limit cycle may be locally approximated 95 using time-varying linear maps [27]. Similar principles have been shown to generalize to limit cycles in nonsmooth or hybrid systems, such as human walking [28]. Moreover, phase-varying modeling principles have 96 97 been applied to biological systems, identifying linear phase-varying dynamics to investigate gait stability and predict changes in kinematics in response to perturbations [25, 26, 29-31]. Responses to ankle exoskeleton 98 99 torques may be similarly defined as perturbations off an unperturbed (*i.e.* zero torque) gait cycle, suggesting 100 that the principles of phase-varying models will generalize to walking with exoskeletons. To the best of our

knowledge, phase-varying models have never been used to study walking with exoskeletons and the extent
 to which the principles underlying phase-varying models of locomotion generalize to walking with
 exoskeletons is unknown.

104

To determine if phase-varying models represent useful predictive tools for locomotion with exoskeletons, the 105 purpose of this research was to evaluate the ability of subject-specific phase-varying models to predict 106 kinematic and myoelectric responses to ankle exoskeleton torque during walking. We predicted responses to 107 108 exoskeletons in unimpaired adults walking with passive ankle exoskeletons under multiple dorsiflexion 109 stiffness conditions. We focused on three related classes of phase-varying models with different structures, complexity, and expected prediction accuracies: a phase-varying (PV), a linear phase-varying (LPV), and a 110 111 nonlinear phase-varying (NPV) model. Since passive exoskeletons typically elicit small changes in joint kinematics and muscle activity, we expected the validity of Floquet Theory for human gait to extend to gait 112 113 with exoskeletons, indicating that the LPV model should accurately predict responses to passive exoskeleton 114 torque [1, 25-27, 29]. We, therefore, hypothesized that the LPV models would predict kinematic and 115 myoelectric responses to torque more accurately than the PV model and as accurately as the NPV model. To exemplify the potential utility of subject-specific phase-varying models in gait analysis with ankle 116 exoskeletons, we show how varying the length of model prediction time horizon may inform measurement 117 118 selection for exoskeleton design and control. To assess the viability of data-driven phase-varying models in 119 gait analysis settings, we evaluated the effect of limiting the size of the training dataset on prediction 120 accuracy.

121

122 IV. METHODS

123 A. Experimental protocol

We collected kinematic and electromyographic (EMG) data from 12 unimpaired adults (6 female / 6 male; age =  $23.9 \pm 1.8$  years; height =  $1.69 \pm 0.10$  m; mass =  $66.5 \pm 11.7$  kg) during treadmill walking with bilateral

6

passive ankle exoskeletons at a self-selected speed. Each participant performed two sessions on separate days
within a one week span. In the first session, we modified the exoskeletons for fit and comfort and performed
a 20-minute practice session. *Additional detail regarding experimental setup, input variable calculations, modeling algorithms, and statistical analyses can be found in Supplemental – S1.*

Data were collected during the second session. We monitored changes in kinematics using a modified Helen-131 Hayes marker set [32] and a 10-camera motion capture system (Qualisys AB, Gothenburg, SE), and measured 132 muscle activity using 14 wireless EMG sensors (Delsys Inc., Natick, USA). The EMG sensors were placed 133 bilaterally on the soleus, medial gastrocnemius, tibialis anterior, vastus medialis, rectus femoris, lateral 134 hamstrings, and gluteus medius following SENIAM guidelines [33]. Participants performed four randomized 135 136 trials on a split-belt instrumented treadmill (Bertec Corp., Columbus, USA) under different exoskeleton conditions (Fig. 1). Unlike many clinical exoskeletons (ankle-foot orthoses), whose torque profiles are 137 smooth functions of ankle angle, the passive exoskeletons used in this study generated ankle plantarflexion 138 torques as a piecewise-linear function of the user's ankle angle, and the exoskeleton's neutral angle and 139 140 rotational stiffness. The exoskeletons did not resist plantarflexion, similar to other experimental devices [1, 3]. The four exoskeleton conditions were set to sagittal-plane stiffness values:  $K_0$  (0 Nm/deg),  $K_1$  (1.17) 141 Nm/deg),  $K_2$  (3.26 Nm/deg), and  $K_3$  (5.08 Nm/deg), a range known to alter kinematics and myoelectric 142 143 signals during gait (Fig. 1) [1]. Participants walked for six minutes per trial, the last four of which were 144 recorded, and could rest between trials.

- 145
- 146

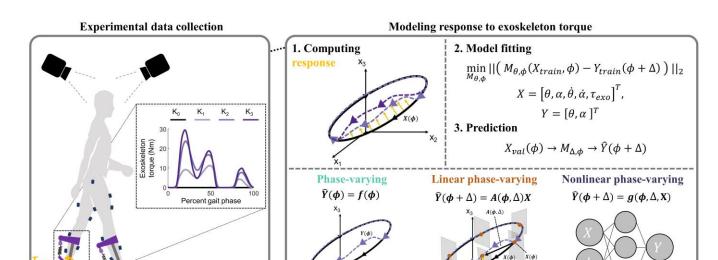


Fig. 1. Left box: Data were collected during treadmill walking with bilateral ankle exoskeletons that used linear springs to resist dorsiflexion. Increasing exoskeleton stiffness ( $K_0-K_3$ ) increased exoskeleton torque ( $\tau_{exo}$ , yellow). <u>Right box</u>: (1) Purple dashed arrows represent responses to exoskeleton torque, which were defined as deviations from the average zero-torque gait cycle ( $K_0$ ). (2) Response data from the training set were used to fit each model. Input variables included joint kinematics, muscle activity, their time derivatives, and exoskeleton torque. (3): Models were validated by predicting responses from the held-out torque condition using the models fit in (2). <u>Right box (bottom)</u>: The three phase-varying models were fit and evaluated on the same training and validation sets.

 $M_{\Delta,\phi}$  = generic model function of prediction horizon and phase; X = experimental inputs; Y = experimental outputs;  $\hat{Y}$  = predicted outputs;  $\phi$  = phase;  $\Delta$  = prediction horizon; A = linear function; f, g = nonlinear functions;  $\theta$  = joint kinematics;  $\alpha$  = muscle activation;  $\tau_{exo}$  = exoskeleton torque.

147

- 148 The marker trajectories were low-pass filtered at 6 Hz using a zero-lag fourth-order Butterworth filter [5].
- 149 We computed joint kinematics by scaling a generic 29 degree-of-freedom skeletal model to each participant's
- skeletal geometry and body mass using the inverse kinematics algorithm in OpenSim 3.3 to convert marker
- trajectories into joint kinematics [18, 34]. To compute linear EMG envelopes, we high-pass filtered the EMG
- data at 40 Hz, rectified the data, and low-pass filtered at 10 Hz [9]. Kinematic and EMG data were pre-
- 153 processed using custom scripts in MATLAB (MathWorks, Natick, USA).
- 154

#### 155 B. Gait phase and phase-varying models

156 Unlike the typical gait cycle definition – the percentage of time between successive foot contact events – we

- 157 used a gait phase based on kinematic posture, which we expected to improve predictions of a system's
- response to perturbations from the exoskeletons [35]. Using a posture-based gait phase groups kinematically-

similar measurements at a specific phase, reducing variance in the data at any point in the cycle, and ensuring that similar postures across exoskeleton conditions were used during model fitting and prediction. Moreover, Floquet Theory ensures that phase is well-defined using any periodically-varying measurements [27]. We used the *Phaser* algorithm, which estimates a system's phase using arbitrary input signals considered to be phase-locked, to generate gait phase estimates as a function of left and right leg hip flexion angles, similar to a phase variable proposed to control robotic prostheses [30, 35]. Following gait phase estimation, we modeled gait using three subject-specific models of response to exoskeletons:

166

#### 167 1) Phase-varying model

The phase-varying (PV) model was our simplest model and predicts outputs purely as a function of gait phase. Rather than taking exoskeleton torque as an input, PV model predictions are similar to guessing the average of the training data at a certain gait phase (Table I) [30, 36]. The PV model takes a phase estimate as an input and returns a prediction of the system's outputs,  $\hat{Y}_{\phi} \in \mathbb{R}^{M}$ , where *M* is the number of outputs. The PV model was parameterized using a seventh-order Fourier Series as a function of phase and served as a lower bound on prediction accuracy.

174

## 175 2) Linear phase-varying model

The linear phase-varying (LPV) model is a discrete-time model that predicts system outputs at a future phase based on measurements at an initial phase (Table I). For any phase,  $\phi$ , from 0-100% of a stride and a prediction horizon,  $\Delta$ , the LPV model estimates a map  $A_{\phi,\Delta} \in \mathbb{R}^{M \times N+1}$ , from the initial phase to the final phase, where N + 1 denotes the number of input variables (N) plus a constant term. At 64 initial phases spaced equally over the gait cycle, we fit discrete maps between initial and final phases using weighted leastsquares regression [25, 26, 29]. We weighted each observation based on the proximity of its phase estimate to the prescribed initial phase using a Gaussian weighting scheme. For each prediction horizon, the LPV

model was represented as a continuously phase-varying function,  $F_{LPV,\Delta}(\phi) \approx A_{\phi,\Delta}$ , parametrized by a Fourier Series. We expected the LPV model's prediction accuracy to exceed that of the PV model [27].

185

## 186 3) Nonlinear phase-varying model

While the LPV model should approximate nonlinearities in the dynamics of response to torque, we selected a nonlinear phase-varying (NPV) model that serves as an upper bound on prediction accuracy. Specifically, we used a three-layer feedforward neural network – a universal function approximator (Table I) [37]. Neural networks are considered state-of-the-art predictors and are used in numerous domains, including image recognition and robotics [38]. The NPV model's parameters were tuned for each prediction horizon and included phase as an input. We expected the NPV model's prediction accuracy to meet or exceed that of the LPV model.

- 194
- 195 Table I: Summary of model structures and expected prediction accuracies.

Model	Functional form	Linear terms	Nonlinear terms	Expected prediction accuracy
Phase-varying (PV)	$\hat{Y}_{\phi} = F_{PV}(\phi)$	None	Phase	Low
Linear phase-varying (LPV)	$\hat{Y}_{\phi+\Delta} = F_{LPV,\Delta}(\phi) X_{\phi}$	Inputs	Phase	Moderate
Nonlinear phase-varying (NPV)	$\widehat{Y}_{\phi+\Delta} = G_{NPV,\Delta}(\phi, X_{\phi})$	None	Phase Inputs	Moderate-High

F = model functions parameterized by a Fourier Series; G = feedforward neural network model;  $\phi =$  phase;  $\Delta =$  prediction horizon; X = inputs;  $\hat{Y} =$  predicted outputs

196

## 197 C. Inputs and output variables

To reflect clinically-relevant measurements and the dynamics of the neuromusculoskeletal system, we selected input variables expected to encode musculoskeletal dynamics and motor control: 3D pelvis orientation and lower-limb and lumbar joint angles, processed EMG signals, and their time derivatives at an initial phase,  $\phi$  [1, 2, 39]. We appended ten time-history exoskeleton torque samples per leg – uniformly distributed between the initial and final phases – to the inputs, resulting in *N* = 80 inputs [6, 12]. Our decision

to use exoskeleton torque samples was motivated by Floquet Theory, according to which an individual's 203 posture at a future time is a linear function of their initial posture and the exoskeleton torque signal between 204 205 initial and final times [27]. Model outputs (M = 20) included right and left leg sagittal-plane hip, knee and ankle kinematics, and EMG signals from each muscle at a future phase,  $\phi + \Delta$ , offset from the initial phase 206 207 by prediction horizon  $\Delta$ . While phase-varying models may also predict joint moments, we omitted prediction 208 of kinetic outcomes due to the presence of sporadic poor force plate strikes for some gait cycles in our dataset. 209 We modeled response to exoskeleton torque as the deviation from the unperturbed gait cycle (*i.e.* the zero-210 torque,  $K_0$  condition) by subtracting the phase-averaged zero-torque gait cycle from each exoskeleton 211 condition [26, 29]. All data were de-meaned and scaled to unit variance of the training set. Additional detail 212 regarding the selection of torque as model inputs and experimental ground reaction forces can be found in 213 Supplemental – S2.

214

215 We first computed each model's ability to predict responses to torque within the range of exoskeleton 216 stiffness levels used to train the models (interpolation) by training each model using the K<sub>0</sub>, K<sub>1</sub> and K<sub>3</sub> 217 datasets and validating by predicting outputs from the held-out K<sub>2</sub> dataset using inputs from the same dataset 218 at an initial gait phase. While "what-if" predictions – predicting responses to "untested" (held-out) torques 219 using nominal kinematics and EMG from a "tested" condition (e.g.  $K_0$ ) – are needed to for predictions to 220 inform passive exoskeleton design, we chose to predict using the held-out inputs to provide unambiguous 221 interpretation of each model's prediction accuracy. In "what-if" predictions, errors stem from both poor 222 model fit and poor matches between the "tested" and "untested" input data at the initial phase. By instead 223 predicting using "untested" inputs, our predictions errors reflect only the models' fits to each participant's 224 dynamics and provide upper bounds on the potential accuracy of "what-if" predictions. We selected the  $K_2$ 225 condition for validation in our experimental design because responses in this intermediate torque condition 226 should be encoded by the  $K_0$ ,  $K_1$ , and  $K_3$  conditions. During validation, experimental outputs from the  $K_2$ 227 condition were compared to the corresponding model predictions.

We quantified each model's prediction accuracy using the relative remaining variance (RRV) of model 229 predictions compared to the held-out experimental data [25]. The RRV is calculated as the ratio of the 230 variances of the prediction error and the experimental data. An RRV value of zero implies a perfect 231 prediction, while unity RRV values can be achieved by predicting the mean of the validation data. Since we 232 de-meaned the data and predicted deviations from the zero-torque condition, RRV values below unity 233 indicate that predictions are more accurate than guessing constant (e.g. zero) response to exoskeleton torque. 234 235 We computed RRV values for each output using a bootstrapping procedure with 200 iterations [25]. We 236 computed RRV values for each model over the entire validation time series of approximately 240 strides. During analysis, the right and left leg RRV values for each output variable were averaged, as we expected 237 238 nearly symmetric responses from our unimpaired participants.

239

228

We evaluated the LPV and NPV models' prediction accuracies for the K<sub>2</sub> condition over a range of prediction 240 horizons, in increments of 6.25% (1/16<sup>th</sup> of a stride), between 6.25 and 100% of a gait cycle. When optimizing 241 242 exoskeleton torque profiles, predicting responses using measurements at an initial phase (e.g. initial contact 243 of the foot with the ground) to achieve a desired outcome at some final phase may be of interest, such as 244 improving midstance knee kinematics in children with cerebral palsy [2, 3]. However, as the prediction 245 horizon increases, coherence between measurements at initial and final phases decreases due to nonlinearities 246 in musculoskeletal dynamics, resulting in prediction accuracies reducing to those of the average prediction 247 (*i.e.* the PV model), rather than a stride-specific prediction [39-41]. Identifying the maximum prediction horizon at which initial measurements improve predictions at a final phase may inform exoskeleton control 248 249 laws or design criteria. Therefore, we identified the largest prediction horizon lengths at which RRV values 250 were significantly less than those of the PV model, which were constant across prediction horizons.

The amount of data required to accurately predict response to exoskeletons will restrict the settings in which 252 phase-varying models are practical, such as in clinical gait analysis where datasets typically contain only a 253 few gait cycles [2, 8]. We quantified the impact of training set size on prediction accuracy by determining 254 255 the amount of training data needed for prediction accuracies of the K<sub>2</sub> condition to approach to their values when models were fit using the entire training set  $(RRV_{full})$ . We iteratively reduced the training set size by 256 10% of the full size (approximately 24 strides per exoskeleton condition), removing data from the end of 257 each torque condition in the training set, providing a range of 24-240 strides of training data per condition. 258 259 For all training set sizes, we evaluated models using the full-length validation set.

260

To test each model's generalizability across a range of exoskeleton torque conditions, we separately predicted 261 262 responses to torque in the K<sub>1</sub>, K<sub>2</sub>, and K<sub>3</sub> datasets, termed *held-out conditions*, at a 12.5% stride prediction horizon (1/8<sup>th</sup> of a stride). Predictions over these conditions evaluated both the models' ability to interpolate 263  $(K_1 \text{ and } K_2)$  and extrapolate  $(K_3)$  responses to exoskeleton torques included in the training set. For each held-264 out condition  $(K_1, K_2, or K_3)$ , we trained the models using kinematic, EMG, and exoskeleton torque inputs 265 266 from the zero-torque ( $K_0$ ) condition and the two non-zero-torque exoskeleton conditions not held out for 267 validation. We evaluated each model by predicting output variables from the held-out exoskeleton condition 268 using input data at an initial gait phase in the same condition. We compared prediction accuracies across 269 held-out conditions.

270

To compare differences in performance across the three models, we identified differences in the models' prediction accuracies using repeated-measures analysis of variance tests at a significance level of  $\alpha = 0.05$ . When significant differences between models emerged, we identified pair-wise differences between models using post-hoc paired t-tests ( $\alpha = 0.05$ ) and a Holm-Sidak step-down correction for multiple comparisons [9, 42]. We report percent reductions in RRV values compared to the PV model and percent differences between the LPV and NPV models.

### 278 V. RESULTS

The ankle exoskeletons had the largest impact on ankle kinematics, smaller impacts on knee and hip kinematics, and variable impacts on muscle activity (Fig. 2). Compared to the K<sub>0</sub> condition, the peak ankle dorsiflexion angle during single-limb support decreased significantly in the K<sub>2</sub> (36.7%) and K<sub>3</sub> (40.0%) conditions (p < 0.020). Average integrated EMG increased slightly, but not significantly in the hamstrings and tibialis anterior (p > 0.066) in the K<sub>2</sub> and K<sub>3</sub> conditions compared to the K<sub>0</sub> condition.

284

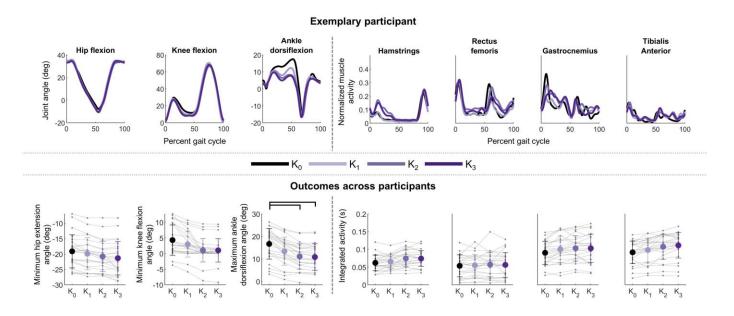


Fig. 2. <u>Top</u>: Average kinematic (left) and EMG (right) data for one participant who exhibited large, repeatable responses to exoskeleton torque and high model prediction accuracies (P03). Black lines show the zero-torque condition (K<sub>0</sub>) that was subtracted from all conditions to reflect responses to exoskeleton torque. <u>Bottom</u>: Average (±1SD) kinematic and myoelectric responses for all participants in each torque condition. Brackets denote significant differences between exoskeleton conditions according to post-hoc paired t-tests ( $\alpha = 0.05$ ) and a Holm-Sidak step-down correction. Thin gray lines represent individual legs.

285

286

When validating on the held-out K<sub>2</sub> condition, all three models predicted kinematic but not myoelectric responses to exoskeleton torque (Fig. 3, dashed lines). At a prediction horizon of  $\Delta = 12.5\%$  of a stride, the LPV model's prediction accuracy at the ankle – where the largest responses to torque were observed – was  $41.6 \pm 16.0\%$  more accurate than the PV model (p < 0.001) but not the NPV model (p = 0.130; Fig. 4;

Table II). Similarly, the LPV model's prediction accuracy at the hip was  $41.7 \pm 12.7\%$  better than the PV 292 model (p < 0.001). However, as prediction horizon increased, the average LPV and NPV model prediction 293 accuracies of all outputs except the ankle approached those of the PV model. Changes in knee and hip 294 kinematics were predicted more accurately than the baseline PV model for prediction horizons shorter than 295  $\Delta = 18.75\%$  of a stride (p < 0.001) in the LPV model and  $\Delta = 12.5\%$  of a stride (p < 0.001) in the NPV model 296 (Fig. 5). At the ankle, the LPV model predicted kinematics 29.1–60.0% more accurately than the PV model 297 298 for all prediction horizons (p < 0.001). The NPV model's predictions were significantly more accurate than those of the PV model for all prediction horizons except 25.0% and 75.5–81.3% of a stride (p < 0.001). 299 300

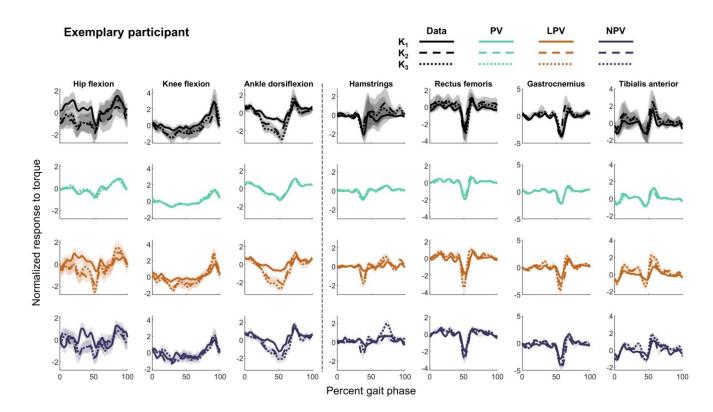


Fig. 3. Kinematic and myoelectric experimental (black) and predicted (colors) responses to torque for one participant who exhibited large responses to the exoskeletons (P03). Predictions are shown for a prediction horizon of 12.5% of a stride for the PV (green), LPV (orange), and NPV (purple) models. The three held-out conditions are denoted with solid (K<sub>1</sub>), dashed (K<sub>2</sub>), and dotted (K<sub>3</sub>) lines. Lines represent the average (±1SD; shaded region) data and predictions over all gait cycles in the corresponding validation dataset. The experimental data show that the K<sub>2</sub> and K<sub>3</sub> responses to torque were more similar to each other than to the K<sub>1</sub> response. The PV model predictions were similar across held-out conditions, while the LPV and NPV models scaled with exoskeleton torque. Full joint trajectories may be reproduced by rescaling and adding the average unperturbed gait cycle to the predictions. All comparisons used paired t-tests ( $\alpha = 0.05$ ) with a Holm-Sidak step-down correction for multiple comparisons.

15



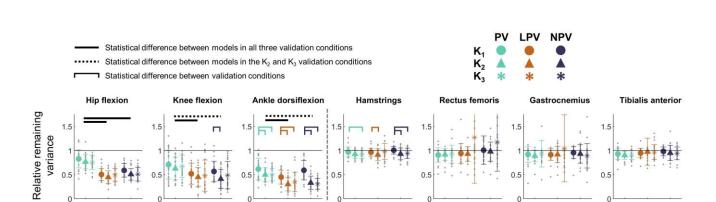


Fig. 4. Average ( $\pm 1$ SD) prediction accuracies for all participants and held-out conditions at a prediction horizon of 12.5% of a stride. Gray dots represent individual legs. Colored brackets denote statistically significant differences between held-out conditions for each model. Black horizontal bars denote significant differences between models across all three (solid) or two (dashed) held-out conditions. The large variance in the LPV model's predictions of rectus femoris and gastrocnemius responses in the held-out K<sub>3</sub> condition were due to bad predictions (RRV > 2) in a small number of legs. All comparisons used paired t-tests ( $\alpha = 0.05$ ) with a Holm-Sidak step-down correction for multiple comparisons.

303

- <sup>304</sup> Predictions of myoelectric responses were poor (RRV  $\approx 1.00$ ) across all muscles and models, except at the
- shortest prediction horizon ( $\Delta = 6.25\%$ ). At the shortest prediction horizon, both the LPV and NPV models'
- <sup>306</sup> predictions for the hamstrings, rectus femoris, and gastrocnemius were 10.7–15.0% more accurate than
- those of the PV model (p < 0.001; Fig. 5).

#### 16

309 Table II: Average (± 1SD) RRV values for kinematic and myoelectric predictions at a 12.5% prediction

310 horizon.

Output	PV	LPV	NPV
Ankle angle <sup>†‡</sup>	$0.50 \pm 0.14$	$0.30 \pm 0.14$	$0.33 \pm 0.14$
Knee angle <sup>†‡</sup>	$0.62 \pm 0.24$	$0.45 \pm 0.20$	$0.41 \pm 0.19$
Hip angle <sup>†‡</sup>	$0.77 \pm 0.15$	$0.44 \pm 0.13$	$0.51 \pm 0.12$
Tibialis anterior	$0.90 \pm 0.08$	$0.97 \pm 0.28$	$0.95 \pm 0.15$
Soleus	$0.86 \pm 0.21$	$1.02 \pm 0.66$	$1.04 \pm 0.64$
Gastrocnemius	$0.89 \pm 0.12$	$0.91 \pm 0.16$	$0.93 \pm 0.19$
Vastus medialis	$0.92 \pm 0.12$	$0.93 \pm 0.15$	$0.98 \pm 0.20$
Rectus femoris	$0.87 \pm 0.21$	$0.91 \pm 0.38$	$0.94 \pm 0.42$
Lateral hamstrings	$0.93 \pm 0.08$	$0.91 \pm 0.13$	$0.93 \pm 0.12$
Gluteus medius	$0.95 \pm 0.08$	$0.96 \pm 0.10$	$0.98 \pm 0.10$

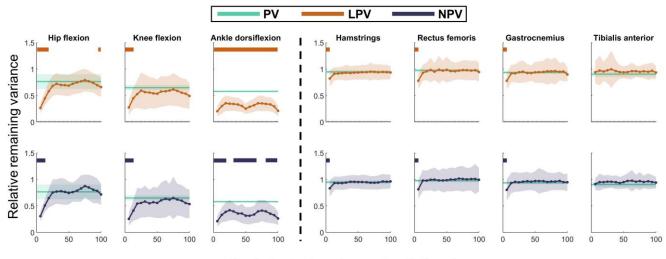
LPV = Linear phase-varying model; NPV = Nonlinear phase-varying model; PV = Phase-varying model

+ Significant difference in prediction accuracy between the PV and LPV models

**‡** Significant difference in prediction accuracy between the PV and NPV models

312

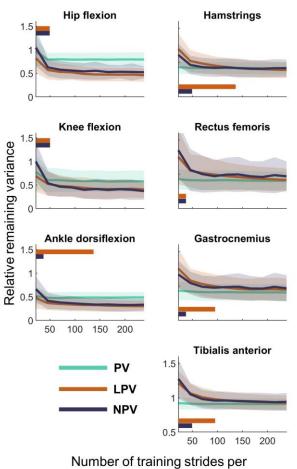
311



Prediction horizon (percent gait phase)

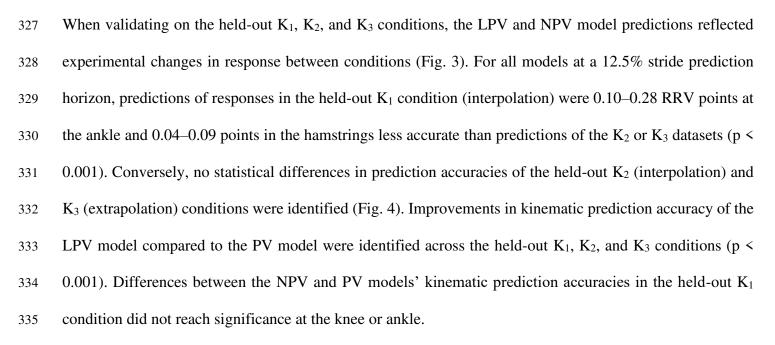
Fig. 5. Prediction accuracy decreased with increasing prediction horizon. The PV model's predictions (green) were constant across prediction horizons. Horizontal bars denote predictions that were significantly more accurate than the PV model. The LPV (top; orange) and NPV (bottom; purple) models' prediction accuracies approached nearly constant values for prediction horizons beyond 25.0% of a stride for kinematic responses and 6.25% of a stride for myoelectric responses. The LPV and NPV models' predictions of ankle kinematics remained more accurate than PV model predictions across almost all prediction horizons, while knee and hip kinematic predictions were similar to those of the PV model beyond 25.0% of a stride.

314	The LPV and NPV models' prediction accuracies improved with increasing training set size. As expected,
315	the PV model's prediction accuracy was nearly constant across training set sizes (p > 0.005; Fig. 6). For a
316	prediction horizon of $\Delta = 12.5\%$ of a stride, the LPV model's hip (RRV = 0.81) and knee (RRV = 0.78)
317	prediction accuracies were significantly worse than $RRV_{full}$ when using less than 50 strides of training data
318	per exoskeleton condition (p < $0.001$ ). Similarly, the NPV model's hip and knee prediction accuracies
319	approached $RRV_{full}$ with approximately 50 strides of training data per condition (p < 0.001). The LPV model
320	required more data – up to 150 strides per condition – for prediction accuracies to approach $RRV_{full}$ at the
321	ankle, gastrocnemius, and tibialis anterior (p < $0.001$ ), though predictions were only $0.02-0.05$ RRV points
322	greater than $RRV_{full}$ with 75 strides of training data per condition. The NPV model's myoelectric prediction
323	accuracies approached RRV <sub>full</sub> , in 25–75 strides of training data per condition ( $p < 0.001$ ; Fig. 6).
324	



exoskeleton condition

Fig. 6. Average ( $\pm 1$ SD; shaded region) prediction accuracy of kinematic (left) and myoelectric (right) outputs for the PV (green), LPV (orange), and NPV (purple) models over training set sizes ranging from 24 to 240 cycles (RRV<sub>full</sub>). Prediction accuracies were reported at a 12.5% stride prediction horizon. Orange (LPV) and purple (NPV) horizontal bars denote the training set sizes that yielded significantly worse predictions than those of the full training set. The PV model's prediction accuracies were not significantly different from RRV<sub>full</sub> at any training set size.



#### 336 VI. DISCUSSION

337 We evaluated the ability of subject-specific phase-varying models to predict kinematic and myoelectric 338 responses to ankle exoskeleton torques during treadmill walking. When predicting across three exoskeleton 339 torque conditions, both linear and nonlinear models predicted kinematic responses to exoskeletons without 340 knowledge of the specific user's physiological characteristics, supporting their potential utility as predictive tools for exoskeleton design and control. To our knowledge, this is the first study to predict kinematic and 341 342 myoelectric responses to ankle exoskeletons using phase-varying models. Consistent with Floquet Theory 343 and prior models of human locomotion, LPV models appear appropriate for predicting responses to exoskeleton torque over short prediction horizons, evidenced by its similar prediction accuracy to the more 344 345 complex NPV model and improved prediction accuracy over the less complex PV model [25-27, 29].

346

The small and variable responses to exoskeleton torque exhibited by the unimpaired adults in this work 347 highlight the challenge of altering kinematics with passive ankle exoskeletons. We found that even stiff 348 349 exoskeletons ( $K_3 = 5.08$  Nm/deg) only altered ankle kinematics on average by six degrees and integrated 350 muscle activity by 14%. These small changes may correspond to larger changes in joint powers or metabolic 351 demands and indicate that the present study is a rigorous test case [1, 2, 5, 24]. Despite small changes in gait, the LPV model's predictions explained more of the variance in kinematic responses to exoskeletons than the 352 PV model, regardless of whether predictions interpolated ( $K_1$  and  $K_2$ ) or extrapolated ( $K_3$ ) relative to the 353 354 training set. The LPV model's ability to predict kinematics within and slightly beyond the available training 355 data supports its potential utility for predicting responses to untested exoskeleton designs or control laws. 356 However, predictions of the held-out  $K_1$  condition highlight the importance of selecting experimental conditions that encode complex responses to torque. 357

358

Our hypothesis that the LPV model would predict kinematic and myoelectric responses more accurately than the PV model and as accurately as the NPV model was partially supported. The LPV model's kinematic and

myoelectric predictions were more accurate than those of the PV model only for prediction horizons less than 361 18.75% and 6.25% of a stride, respectively, but the LPV and NPV models exhibited similar prediction 362 accuracies across prediction horizons. The LPV and NPV models' similar predictions support research 363 364 demonstrating that nonlinear spring-loaded inverted pendula (SLIPs) have similar predictive accuracy to linear models of human movement [25]. Compared to a nonlinear SLIP, the NPV model's feedforward neural 365 network imposed fewer restrictions on model structure and enabling greater differences in prediction 366 accuracy compared to a linear model. Therefore, the similarity of LPV and NPV model predictions supports 367 368 the extension of Floquet Theory to gait with exoskeletons and indicates that, for rhythmic locomotion at a 369 constant speed over level ground, linear phase-varying models have sufficiently complex structure to predict 370 kinematic responses to exoskeletons [25-27, 29].

371

We observed comparable kinematic prediction accuracy to studies using physics-based and data-driven 372 373 models of locomotion. Maus et al. evaluated multiple models' abilities to predict center-of-mass height 374 during running and reported accuracies ranging from RRV  $\approx 0.15$  at a 15% prediction horizon to RRV  $\approx 0.85$ 375 beyond an 80% stride prediction horizon, for an exemplary participant [25]. Within a similar range of prediction horizons, the LPV model predicted kinematics across participants with average accuracies ranging 376 from (0.30 < RRV < 0.45) at a 12.5% prediction horizon and (0.34 < RRV < 0.77) at an 81.3% prediction 377 378 horizon. Similarly, Drnach et al. [43] used a hybrid linear model to predict response to functional electrical 379 stimulation, reporting median RRV values (transformed from a fitness score) ranging from approximately 380 0.11-1.04. However, the average unperturbed gait cycle was not subtracted from the data before computing 381 the fitness score in [43]. The average unperturbed cycle accounts for a substantial portion of the variance in 382 the perturbed signals, providing a less conservative prediction accuracy statistic than the RRV presented here. 383 For example, if the unperturbed cycle had not been subtracted from the data in the present study, the LPV 384 model's ankle predictions for one participant who exhibited large responses to torque would be RRV = 0.08385 rather than the more conservative 0.21 reported. Comparable prediction accuracies to prior work indicate that

21

386 phase-varying models are potentially useful predictive tools for locomotion with ankle exoskeletons and may 387 have similar predictive power to physics-based models of locomotion.

388

389 The convergence of LPV and NPV models' prediction accuracies to an approximately constant value at large prediction horizons (e.g. RRV<sub>LPV</sub>  $\approx 0.70$  for knee kinematics at  $\Delta > 25.0\%$  of a stride) may be useful when 390 selecting measurements for device design or control. The LPV and NPV models' kinematic prediction 391 accuracies decreased rapidly from 6.25% to 18.75% stride prediction horizons, before reaching an 392 393 approximately constant value. Ankle predictions remained better than those of the PV model across 394 prediction horizons. Higher prediction accuracy at the ankle was unsurprising due to large responses to exoskeletons and the ankle's direct piecewise-linear relationship to passive exoskeleton torque. Since we 395 396 trained on multiple exoskeleton conditions, the dynamics predicting future ankle kinematics are higherdimensional than the simple exoskeleton torque-ankle angle relationship, suggesting that accurate predictions 397 398 of ankle kinematics over large prediction horizons are likely for powered exoskeletons as well. Unlike the 399 ankle, hip and knee kinematics were indirectly impacted by exoskeleton torque and their RRV values 400 approached those of the PV model for prediction horizons above 18.75% of a stride. This result indicates that 401 stride-specific initial posture and exoskeleton torque were predictive of indirect exoskeleton impacts on 402 kinematics only for short prediction horizons. At large prediction horizons, measurements at an initial phase 403 did not, on average, improve predictions of future posture. However, some participants' hip and knee 404 kinematics were predicted up to 0.30 RRV points more accurately by the LPV and NPV models than the PV 405 model across prediction horizons, suggesting that the prediction horizon at which stride-specific 406 measurements no longer improve predicted responses to exoskeletons depends on the magnitude of the 407 individual's response. The LPV and NPV models' accurate predictions over short prediction horizons make 408 them primarily useful for exoskeleton control [10, 11]. For individuals that exhibit large responses to 409 exoskeletons, however, LPV model-based predictions over stance may inform passive exoskeleton parameter 410 selection. Guided adaptation and extended practice sessions [1, 17] or powered ankle exoskeletons [5, 6] may

411 elicit larger responses than those observed in this study and increase the maximum prediction horizons at 412 which measurements at an initial posture improve predicted responses to torque, potentially expanding the 413 settings in which model predictions are useful.

414

A major limitation of all three models was their inability to predict myoelectric responses. The LPV and NPV 415 models predicted myoelectric signals more accurately than the PV model only for the shortest prediction 416 417 horizon ( $\Delta = 6.25\%$ ). While exoskeleton torque and stiffness are known to impact average plantarflexor 418 activity, we found that the average unperturbed gait cycle accounted for only 30-60% of the variance in the K<sub>2</sub> data, compared to 60-95% in kinematic signals [1, 9, 12]. Consequently, poor prediction accuracy may 419 420 be partially attributed to small changes in muscle activity between the exoskeleton conditions. Alternatively, 421 kinematic and myoelectric input variables may fail to encode nonlinear musculotendon dynamics, which are impacted by ankle exoskeletons, between the initial and final phases [21, 40]. Studies predicting muscle 422 423 activity using physiologically-detailed models accounted for 60-99% of the variance in myoelectric signals, 424 though they evaluated predictions on unperturbed walking conditions only [44, 45]. Still, the difference in 425 prediction accuracy between the phase-varying models and physiologically-detailed models indicates that 426 encoding musculotendon dynamics in the input variables is likely needed to improve myoelectric predictions 427 for data-driven phase-varying models and represents an interesting area of future research.

428

Another limitation of subject-specific data-driven models, compared to physiologically-detailed models, is the amount of training data required to predict changes in gait with exoskeletons, which impacts models' utility in settings where minimizing data collection duration is critical to mitigating physical and logistical burdens on participants and families, such as in clinical gait laboratories. Improvements in prediction accuracy of the LPV and NPV models were small beyond 75–100 strides of training data per exoskeleton condition. The LPV model required more training data at the ankle, but a similar amount at the hip and knee to that used by Drnach et al., who trained a hybrid linear model using 45 seconds of data across two

experimental conditions [43]. For unimpaired, steady-state locomotion, data-driven linear models appear to require 75–125 strides of training data per condition, which supports their feasibility only in gait analysis settings with treadmills or long walkways [6, 7]. Additional dimensionality reduction, such as via sparse regression, may reduce the LPV model's complexity and demand for training data [25, 31, 46]. However, when only one training condition or a few strides are collected, as is standard in clinical gait analysis, phasevarying model predictions will be poor and physiologically-detailed or population-specific models may generate more accurate predictions [8, 19, 44, 45].

443

Subject-specific data-driven phase-varying models of gait with exoskeletons have benefits and limitations 444 compared to predictive musculoskeletal models. While we investigated only a specific subset of phase-445 446 varying models, we showed that this class of model can predict kinematic responses to exoskeletons without detailed knowledge of the physiological and neuromuscular factors influencing responses to exoskeletons. 447 448 Conversely, uncertainty in the mechanisms driving complex responses to exoskeletons may limit 449 physiologically-detailed models' accuracy [13, 24]. While predictive musculoskeletal models may generate "what-if" predictions without experimental data, data may be needed to specify initial postures and tune 450 subject-specific parameters. Phase-varying models can similarly perform subject-specific "what-if" 451 predictions when application-specific training data are available. Unlike physiologically-detail models, this 452 453 and prior work exemplify phase-varying models' ability to take arbitrary measurements as inputs, enabling 454 their application using a range of experimental resources [25, 26, 31]. Extending data-driven predictions to "what-if" scenarios and improving predicted myoelectric responses to exoskeletons, combined with 455 456 analytical tools for phase-varying systems(e.g. [31]), may facilitate prediction and analysis of individualized 457 exoskeleton impacts on gait mechanics and motor control.

## 459 VII. CONCLUSION

To our knowledge, this is the first study to predict subject-specific responses to ankle exoskeletons using 460 461 phase-varying models. Without making assumptions about individual physiology or motor control, an LPV model predicted short-time kinematic responses to bilateral passive ankle exoskeletons, though predicting 462 myoelectric responses remains challenging. Results support the utility of LPV models for studying and 463 predicting response to exoskeleton torque. Improving data-driven models and experimental protocols to study 464 and predict myoelectric responses to exoskeletons represents an important direction for future research. 465 466 Modeling responses to exoskeletons or other assistive devices using a phase-varying perspective has the potential to inform exoskeleton design for a range of user groups. 467

## 469 VIII. AUTHOR CONTRIBUTIONS

470 471	MCR was involved in the conception and design of the study, carried out exoskeleton design and fabrication,
472	data collection, data preprocessing, statistical analyses and data interpretation, and drafted the manuscript.
473	BSB was involved in the conception and design of the study and carried out model development and coding,
474	contributed to the initial manuscript draft, and critically revised the manuscript. SAB critically revised the
475	manuscript and acquired funding for this work and was involved in the conception and design of the study,
476	model development, and data interpretation. KMS critically revised the manuscript, acquired funding for this
477	work, and was involved in the conception and design of the study and data interpretation.
478	
479	IX. DATA ACCESSIBILITY
480	All experimental datasets and code using in this study are freely available and can be accessed on
481	https://simtk.org/projects/ankleexopred.
482	
482 483	X. Ethics
	X. ETHICS This study was approved by the University of Washington Institutional Review Board #47744. All research
483	
483 484	This study was approved by the University of Washington Institutional Review Board #47744. All research
483 484 485	This study was approved by the University of Washington Institutional Review Board #47744. All research
483 484 485 486	This study was approved by the University of Washington Institutional Review Board #47744. All research participants provided informed consent prior to participating in the study, obtained by MCR.
483 484 485 486 487	This study was approved by the University of Washington Institutional Review Board #47744. All research participants provided informed consent prior to participating in the study, obtained by MCR. XI. FUNDING STATEMENT
483 484 485 486 487 488	This study was approved by the University of Washington Institutional Review Board #47744. All research participants provided informed consent prior to participating in the study, obtained by MCR. XI. FUNDING STATEMENT This material is based upon work supported by the U. S. Army Research Office under grant number W911NF-
483 484 485 486 487 488 489	This study was approved by the University of Washington Institutional Review Board #47744. All research participants provided informed consent prior to participating in the study, obtained by MCR. XI. FUNDING STATEMENT This material is based upon work supported by the U. S. Army Research Office under grant number W911NF- 16-1-0158 to SAB, the National Science Foundation under grant No. CBET-1452646 to KMS, the National
483 484 485 486 487 488 489 490	This study was approved by the University of Washington Institutional Review Board #47744. All research participants provided informed consent prior to participating in the study, obtained by MCR. XI. FUNDING STATEMENT This material is based upon work supported by the U. S. Army Research Office under grant number W911NF- 16-1-0158 to SAB, the National Science Foundation under grant No. CBET-1452646 to KMS, the National Science Foundation Graduate Research Fellowship Program under Grant No. DGE-1762114 to MCR, and

- 493 XII. REFERENCES
- 494 [1] Collins, S.H., Wiggin, M.B. & Sawicki, G.S. 2015 Reducing the energy cost of human walking using an
  495 unpowered exoskeleton. *Nature* 522, 212-215.
- 496 [2] Kerkum, Y.L., Buizer, A.I., van den Noort, J.C., Becher, J.G., Harlaar, J. & Brehm, M.-A. 2015 The
- 497 effects of varying ankle foot orthosis stiffness on gait in children with spastic cerebral palsy who walk with
- 498 excessive knee flexion. *PloS one* **10**, e0142878.
- 499 [3] Owen, E. 2010 The importance of being earnest about shank and thigh kinematics especially when using
- ankle-foot orthoses. *Prosthetics and orthotics international* **34**, 254-269.
- 501 [4] Lerner, Z.F., Harvey, T.A. & Lawson, J.L. 2019 A Battery-Powered Ankle Exoskeleton Improves Gait
- 502 Mechanics in a Feasibility Study of Individuals with Cerebral Palsy. *Annals of biomedical engineering* **47**,
- 503 1345-1356.
- 504 [5] Mooney, L.M. & Herr, H.M. 2016 Biomechanical walking mechanisms underlying the metabolic 505 reduction caused by an autonomous exoskeleton. *Journal of neuroengineering and rehabilitation* **13**, 4.
- 506 [6] Zhang, J., Fiers, P., Witte, K.A., Jackson, R.W., Poggensee, K.L., Atkeson, C.G. & Collins, S.H. 2017
- 507 Human-in-the-loop optimization of exoskeleton assistance during walking. *Science* **356**, 1280-1284.
- 508 [7] Ding, Y., Kim, M., Kuindersma, S. & Walsh, C.J. 2018 Human-in-the-loop optimization of hip assistance
- 509 with a soft exosuit during walking. *Science Robotics* **3**, eaar5438.
- 510 [8] Ries, A.J., Novacheck, T.F. & Schwartz, M.H. 2014 A data driven model for optimal orthosis selection
- 511 in children with cerebral palsy. *Gait & posture* **40**, 539-544.
- [9] Steele, K.M., Jackson, R.W., Shuman, B.R. & Collins, S.H. 2017 Muscle recruitment and coordination
  with an ankle exoskeleton. *Journal of biomechanics* 59, 50-58.
- 514 [10] Åström, K.J. & Murray, R.M. 2010 Feedback systems: an introduction for scientists and engineers,
- 515 Princeton university press.

- 516 [11] Koller, J.R., Jacobs, D.A., Ferris, D.P. & Remy, C.D. 2015 Learning to walk with an adaptive gain
- 517 proportional myoelectric controller for a robotic ankle exoskeleton. *Journal of neuroengineering and* 518 *rehabilitation* **12**, 97.
- [12] Jackson, R.W. & Collins, S.H. 2015 An experimental comparison of the relative benefits of work and
  torque assistance in ankle exoskeletons. *Journal of Applied Physiology* 119, 541-557.
- 521 [13] Ries, A.J., Novacheck, T.F. & Schwartz, M.H. 2015 The efficacy of ankle-foot orthoses on improving
- the gait of children with diplegic cerebral palsy: a multiple outcome analysis. *PM&R* 7, 922-929.
- 523 [14] Bregman, D., Van der Krogt, M., De Groot, V., Harlaar, J., Wisse, M. & Collins, S. 2011 The effect of
- ankle foot orthosis stiffness on the energy cost of walking: a simulation study. *Clinical Biomechanics* 26,
  955-961.
- 526 [15] Rosenberg, M. & Steele, K.M. 2017 Simulated impacts of ankle foot orthoses on muscle demand and
- recruitment in typically-developing children and children with cerebral palsy and crouch gait. *PloS one* 12,
  e0180219.
- 529 [16] Uchida, T.K., Seth, A., Pouya, S., Dembia, C.L., Hicks, J.L. & Delp, S.L. 2016 Simulating Ideal
- Assistive Devices to Reduce the Metabolic Cost of Running. *PloS one* **11**, e0163417.
- <sup>531</sup> [17] Selinger, J.C., Wong, J.D., Simha, S.N. & Donelan, J.M. 2019 How humans initiate energy optimization
- and converge on their optimal gaits. *Journal of Experimental Biology* **222**, jeb198234.
- 533 [18] Delp, S.L., Anderson, F.C., Arnold, A.S., Loan, P., Habib, A., John, C.T., Guendelman, E. & Thelen,
- 534 D.G. 2007 OpenSim: open-source software to create and analyze dynamic simulations of movement. *IEEE*
- transactions on biomedical engineering **54**, 1940-1950.
- [19] Pitto, L., Kainz, H., Falisse, A., Wesseling, M., Van Rossom, S., Hoang, H., Papageorgiou, E.,
  Hallemans, A., Desloovere, K. & Molenaers, G. 2019 SimCP: a simulation platform to predict gait
- 538 performance following orthopedic intervention in children with Cerebral Palsy. *Frontiers in neurorobotics*
- 539 **13**, 54.

- [20] Handsfield, G.G., Meyer, C.H., Abel, M.F. & Blemker, S.S. 2016 Heterogeneity of muscle sizes in the
- 541 lower limbs of children with cerebral palsy. *Muscle & nerve* **53**, 933-945.
- 542 [21] Sawicki, G.S. & Khan, N.S. 2016 A Simple Model to Estimate Plantarflexor Muscle–Tendon Mechanics
- 543 and Energetics During Walking With Elastic Ankle Exoskeletons. IEEE Transactions on Biomedical
- 544 *Engineering* **63**, 914-923.
- 545 [22] Steele, K.M., Rozumalski, A. & Schwartz, M.H. 2015 Muscle synergies and complexity of
- neuromuscular control during gait in cerebral palsy. *Developmental Medicine & Child Neurology* **57**, 1176-
- 547 1182.
- 548 [23] Chisholm, A.E. & Perry, S.D. 2012 Ankle-foot orthotic management in neuromuscular disorders:
- recommendations for future research. *Disability and Rehabilitation: Assistive Technology* **7**, 437-449.
- 550 [24] Jackson, R.W., Dembia, C.L., Delp, S.L. & Collins, S.H. 2017 Muscle-tendon mechanics explain
- unexpected effects of exoskeleton assistance on metabolic rate during walking. *Journal of Experimental Biology* 220, 2082-2095.
- [25] Maus, H.-M., Revzen, S., Guckenheimer, J., Ludwig, C., Reger, J. & Seyfarth, A. 2015 Constructing
   predictive models of human running. *Journal of The Royal Society Interface* 12, 20140899.
- [26] Wang, Y. & Srinivasan, M. System identification and stability analyses of steady human locomotion. *Foot* **300**, 600.
- [27] Hartman, P. 2002 *Ordinary differential equations*. 2nd ed. ed. Philadelphia, Philadelphia : Society for
  Industrial and Applied Mathematics.
- [28] Burden, S.A., Revzen, S. & Sastry, S.S. 2015 Model reduction near periodic orbits of hybrid dynamical
  systems. *IEEE Transactions on Automatic Control* 60, 2626-2639.
- 561 [29] Ankaralı, M.M., Sefati, S., Madhav, M.S., Long, A., Bastian, A.J. & Cowan, N.J. 2015 Walking
- dynamics are symmetric (enough). *Journal of the Royal Society Interface* **12**, 20150209.
- [30] Revzen, S. & Guckenheimer, J.M. 2008 Estimating the phase of synchronized oscillators. *Physical Review E* 78, 051907.

- 565 [31] Revzen, S. & Guckenheimer, J.M. 2011 Finding the dimension of slow dynamics in a rhythmic system.
- *Journal of The Royal Society Interface* **9**, 957-971.
- [32] Kadaba, M.P., Ramakrishnan, H. & Wootten, M. 1990 Measurement of lower extremity kinematics
  during level walking. *Journal of orthopaedic research* 8, 383-392.
- 569 [33] Hermens, H.J., Freriks, B., Merletti, R., Stegeman, D., Blok, J., Rau, G., Disselhorst-Klug, C. & Hägg,
- 570 G. 1999 European recommendations for surface electromyography. *Roessingh research and development* **8**,
- 571 13-54.
- 572 [34] Rajagopal, A., Dembia, C.L., DeMers, M.S., Delp, D.D., Hicks, J.L. & Delp, S.L. 2016 Full-body
- 573 musculoskeletal model for muscle-driven simulation of human gait. *IEEE transactions on biomedical*
- 574 *engineering* **63**, 2068-2079.
- 575 [35] Villarreal, D.J., Poonawala, H.A. & Gregg, R.D. 2016 A robust parameterization of human gait patterns
- across phase-shifting perturbations. *IEEE Transactions on Neural Systems and Rehabilitation Engineering*25, 265-278.
- 578 [36] Revzen, S. 2020 BIRDS Lab mathmisc. (GitHub).
- [37] Glorot, X. & Bengio, Y. 2010 Understanding the difficulty of training deep feedforward neural networks.
- 580 In Proceedings of the thirteenth international conference on artificial intelligence and statistics (pp. 249-
- 581 256.
- 582 [38] LeCun, Y., Bengio, Y. & Hinton, G. 2015 Deep learning. *nature* **521**, 436-444.
- 583 [39] Zajac, F.E., Neptune, R.R. & Kautz, S.A. 2002 Biomechanics and muscle coordination of human
- walking: Part I: Introduction to concepts, power transfer, dynamics and simulations. *Gait & posture* 16, 215-
- 585 232.
- 586 [40] Zajac, F.E. 1989 Muscle and tendon: properties, models, scaling, and application to biomechanics and
- 587 motor control. *Critical reviews in biomedical engineering* **17**, 359-411.
- 588 [41] Carter, G.C. 1987 Coherence and time delay estimation. *Proceedings of the IEEE* **75**, 236-255.
- 589 [42] Glantz, S. 2012 Primer of Biostatistics, 7th edn, pp. 65–67. (New York: McGraw-Hill.

- 590 [43] Drnach, L., Allen, J.L., Essa, I. & Ting, L.H. 2019 A Data-Driven Predictive Model of Individual-
- 591 Specific Effects of FES on Human Gait Dynamics. In 2019 International Conference on Robotics and
- 592 Automation (ICRA) (pp. 5090-5096, IEEE.
- 593 [44] Geyer, H., Seyfarth, A. & Blickhan, R. 2006 Compliant leg behaviour explains basic dynamics of
- walking and running. *Proceedings of the Royal Society B: Biological Sciences* **273**, 2861-2867.
- 595 [45] Martelli, S., Calvetti, D., Somersalo, E. & Viceconti, M. 2015 Stochastic modelling of muscle
- recruitment during activity. *Interface focus* **5**, 20140094.
- 597 [46] Brunton, S.L., Proctor, J.L. & Kutz, J.N. 2016 Discovering governing equations from data by sparse
- identification of nonlinear dynamical systems. Proceedings of the National Academy of Sciences 113, 3932-
- 599 3937.