

Uncertainty-aware Automated Assessment of the Arm Impedance with Upper-limb Exoskeletons

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2 ABSTRACT

3 Providing high degree of personalization to a specific need of each patient is invaluable to
4 improve the utility of robot-driven neurorehabilitation. For the desired customization of treatment
5 strategies, precise and reliable estimation of the patient's state becomes important, as it can
6 be used to continuously monitor the patient during training and to document the rehabilitation
7 progress. Wearable robotics have emerged as a valuable tool for this quantitative assessment as
8 the actuation and sensing are performed on the joint level. However, upper-limb exoskeletons
9 introduce various sources of uncertainty, which primarily result from the complex interaction
10 dynamics at the physical interface between the patient and the robotic device. These sources of
11 uncertainty must be considered to ensure the correctness of estimation results when performing
12 the clinical assessment of the patient state. In this work, we analyze these sources of uncertainty
13 and quantify their influence on the estimation of the human arm impedance. We argue that this
14 mitigates the risk of relying on overconfident estimates and promotes more precise computational
15 approaches in robot-based neurorehabilitation.

16 **Keywords:** reliable automated assessment, sensitivity analysis, human-exoskeleton interaction, uncertainty quantification, uncertainty-
17 aware simulation, neuromechanical state estimation

1 INTRODUCTION

18 Medical robotics have advanced greatly with application in many domains, such as robot-assisted
19 surgery (D'Etterre et al., 2021), service robots in healthcare (Holland et al., 2021) or rehabilitation
20 robotics (Laut et al., 2016). Particularly in the field of physical rehabilitation, an ever-increasing demand
21 for automation technology is observed. Stroke, for instance, is the second leading cause of death
22 worldwide (Feigin et al., 2014) with an increasing trend due to rising life expectancy in many parts
23 of the world (Boehme et al., 2017; Donkor, 2018). However, while stroke is a highly relevant cause
24 for motor impairment, many other neurological disorders, such as cerebral palsy, multiple sclerosis
25 or Parkinson's disease, require similar treatment strategies during rehabilitation to improve or retain
26 motor functions (Krebs et al., 2008). In particular, high-intensity (Ringleb et al., 2008) and repetition
27 training (Kwakkel et al., 1999) have been shown to produce promising recovery results. Due to these
28 requirements, effective rehabilitation is time- and labor-intensive, therefore, both patients and healthcare
29 professionals can benefit greatly from robot-assisted rehabilitation strategies.

30 In recent years exoskeletons, also referred to as wearable robotic devices (Lo and Xie, 2012), have
31 emerged as a powerful tool for rehabilitation. Since they are designed in a manner that the kinematic chain
32 aligns with the user, sensing and actuation can be performed at the joint level here. One of the main benefits
33 of rehabilitation robotics lies in their application during robot-aided patient assessment. Here, robotic
34 devices are used to monitor patients before, after, or during training, thereby tracking the recovery progress
35 and informing the treatment strategy. In the case of neurological disorders, there are multiple functional
36 impairments, e.g., arm hemiparesis, limited hand dexterity or over-rigid joints, that inhibit motor functions
37 of affected individuals (Carvalho-Pinto and Faria, 2016). Thus, the quantitative estimation of the dynamic
38 parameters underlying these effects using wearable robotic devices can greatly benefit neurorehabilitation.
39 Particularly relevant in the case of stroke is spasticity, a motor disorder described by hyperactivity in tonic
40 stretch reflexes (McLellan, 1981) which leads muscles to be overly resistive to elongations and thus reduced
41 mobility of the affected limb (Sommerfeld et al., 2004). In current clinical practice, spasticity assessment
42 scales, such as the Modified Ashworth Scale (MAS) are used to evaluate the muscle tone of patients. Here,
43 the clinician induces a passive motion by manually perturbing the target joint of the patient. Concurrently,
44 the muscle tone is assessed by tactually observing the movement resistance. Even though this method has
45 been proven to be useful in clinical practice (Gregson et al., 1999), there are shortcomings that could be
46 alleviated through robotic assessment. Specifically, the coarse and discrete nature of the scales limit the
47 level of precision. Additionally, the evaluation is subjective at its core, which can lead to possibly unreliable
48 and biased estimates that are not consistently reproducible (Blackburn et al., 2002; Raghavan, 2015).

49 Hence, the deployment of robot-aided assessment is expected to improve the objectivity and repeatability
50 of clinical evaluations (Lambercy et al., 2012). In particular, joint impedance is commonly used as a
51 concise measure for the patient state (Maggioni et al., 2016), since it describes the relationship between
52 joint motion and opposing torque, which is often abnormally increased (Chung et al., 2004). In recent
53 years, a multitude of these assessment approaches based on exoskeletons for upper-limb rehabilitation have
54 emerged. In Ren et al. (2013), an upper-limb exoskeleton quantitatively estimates the joint stiffness of the
55 shoulder, elbow and wrist joints. More recently, a decomposition of the coupled human arm dynamics is
56 proposed to allow the estimation of local and inter-joint stiffness effects following stroke (Zhang et al.,
57 2017). A more extensive impedance estimation is conducted in Wang et al. (2021), where an exoskeleton is
58 used to identify the inertia, viscosity and stiffness components of the elbow joint of patients' with spastic
59 arms using genetic algorithms. Despite the fact that the benefits of robot-aided assessment in comparison to
60 human-administered clinical scales have been demonstrated in studies (Bosecker et al., 2010), exoskeleton
61 applications suffer from the introduction of unintended interaction forces to the user (Jarrassé et al., 2010)
62 with adverse effects on the clinical evaluation. These interaction forces cannot be avoided completely due to
63 uncertainties in the complex physical human-exoskeleton interaction. In particular, sources of uncertainty
64 are known to arise due to kinematic incompatibilities, soft coupling and inaccuracies in the human dynamics
65 model (Pons, 2008). So far, the influence of these sources of uncertainty on the arm impedance estimation
66 has not been analyzed sufficiently, and a quantitative ranking of their impact is missing. However, since the
67 assessment is used to guide the therapy of patients, it is paramount to make these uncertainties explicit in
68 order to increase precision and ensure that clinicians are not misinformed by overconfident assessment
69 results. Therefore, it is important to investigate how uncertain the obtained impedance parameter estimates
70 are and how to effectively reduce uncertainty for exoskeleton-based automated assessment.

71 1.1 Related Work

72 The influence of uncertainties on the robot-aided impedance estimation can be quantified by mean of
73 a sensitivity analysis. These methods investigate how uncertainty in the output of a system, e.g., the
74 result of the automated assessment, is influenced by variations in the input of a system (Pianosi et al.,

2016), e.g., sources of uncertainty in the complex human-exoskeleton interaction. Thus, by analyzing these sensitivities and ascribing quantitative measures of importance to each source of uncertainty, the robustness of the automated assessment can be quantified (Thabane et al., 2013). Previously, it has been shown how sensitivity analysis methods are used to support efforts in uncertainty reduction (Hamm et al., 2006) and facilitate robust decision making under uncertainty (Nguyen and de Kok, 2007; Singh et al., 2014).

In general, sensitivity analysis can be approached in multiple ways, with three principle classes identified in (Christopher Frey and Patil, 2002): Analytical, statistical and graphical methods. Typically, analytical methods, such as (Kohberger et al., 1978; Ma et al., 2021), require access to a differential equation model of the system and perform analysis by monitoring the partial derivative over the uncertain parameters (Abraham et al., 2007). In Schiele (2008), an analytical 1 DoF model of the interaction forces induced by kinematic incompatibilities on the elbow joint is proposed. While the presented model was validated experimentally, remaining sources of uncertainty are not considered and it limits the utility of the model as interaction effects cannot be captured by it. Due to the complexity of the human-exoskeleton interaction dynamics, a closed-form description that captures all sources of uncertainty concurrently is not available, which makes analytical sensitivity analysis methods impractical. On the other hand, statistical and graphical approaches solely require access to input-output samples of the system (Christopher Frey and Patil, 2002). Here, samples are generated by evaluating the examined system for a factorial combination of all sources of uncertainty to obtain pertinent statistical information and gain rigorous insights, which is infeasible to do experimentally. Thus, simulations are often used instead (Iooss and Saltelli, 2017). However, to the best of the authors' knowledge, no human-exoskeleton simulation environment considers all of the key sources of uncertainty present during the complex, physical interaction. In (Agarwal et al., 2010), for instance, the authors analyzed challenges due to kinematic misalignments on the elbow joint to inform the simulation-based design of an arm exoskeleton. On the other hand, the effect of the human musculoskeletal model on lower-limb exoskeleton control during gait is investigated in (Khamar et al., 2019). Lastly, (Kühn et al., 2018) present an upper-limb simulation of the human, exoskeleton and their respective coupling where simplified 6 DoF springs are used to model soft-contacts. However, in order to fully understand the effect of uncertainty in exoskeleton-based impedance assessment, all sources of uncertainty and their interaction effects must be considered. Thus, a simulation platform which can systematically express the uncertain human-exoskeleton interaction is required in order to quantify the impact of sources of uncertainty on the estimated impedance parameter.

1.2 Contribution

In this work, we perform a sensitivity analysis that quantitatively investigates the influence of various sources of uncertainty on the exoskeleton-based arm impedance estimation. Through this process, a more precise understanding of the uncertainty composition and their prioritization is achieved, which facilitates effective measures to increase the performance of exoskeleton-based automated assessment and reduces the risk of relying on overconfident results. We propose a two-phase approach, where initially the negligible sources of uncertainty are identified, and then a ranking of the most influential factors is performed in the second phase. Due to the complexity of the human-exoskeleton interaction dynamics, we adopt a sampling-based sensitivity analysis which allows us to quantify the influence of each source of uncertainty independently as well as the interaction effects among them. In order to generate the samples required for the analysis, we develop a high-fidelity simulation environment of the human-exoskeleton system that includes the key sources of uncertainty, which are informed by the physical understanding of the system and identified in the literature.

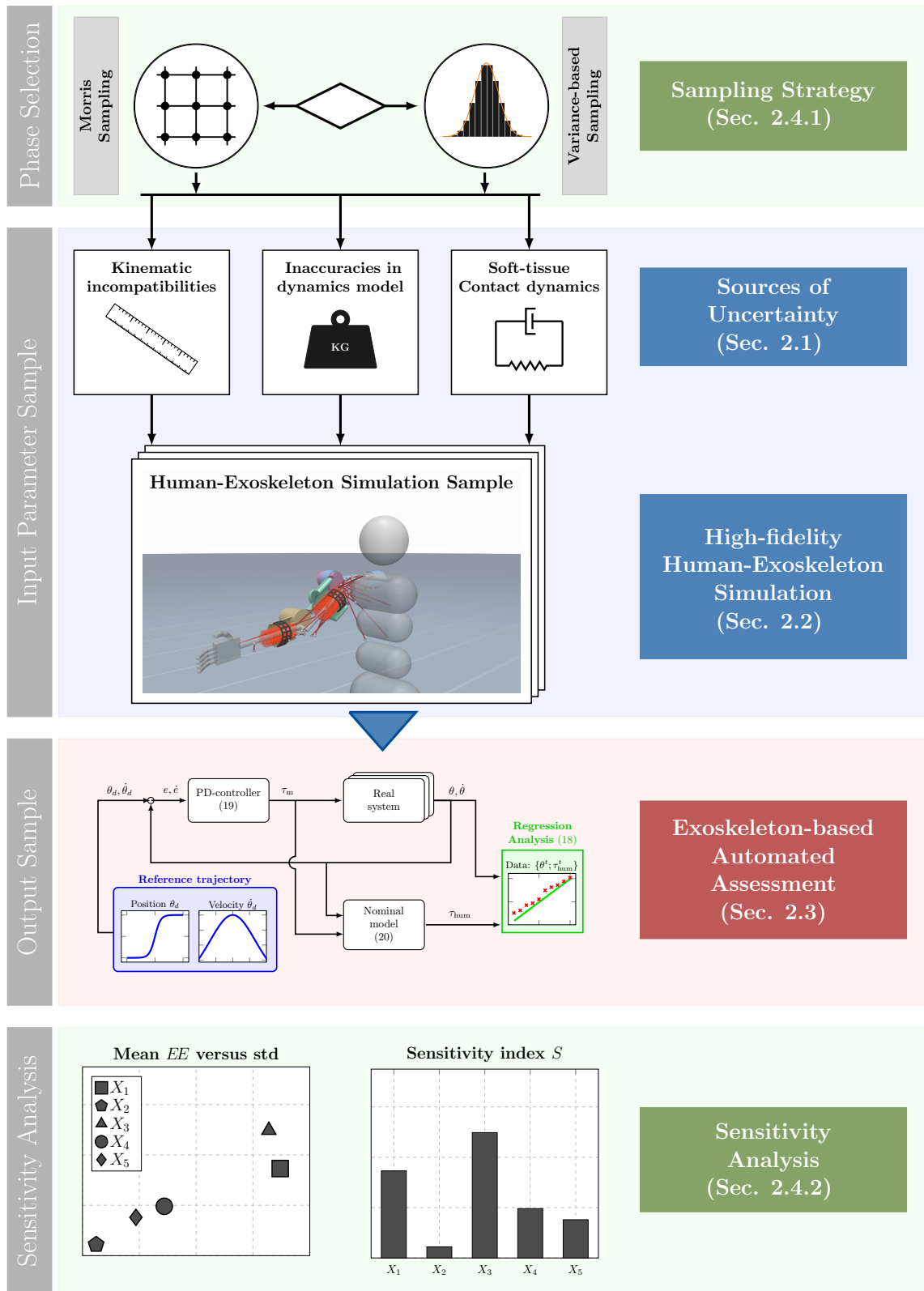


Figure 1. Depiction of the complete, proposed sensitivity analysis scheme. From top to bottom the blocks illustrate the different steps taken during the proposed scheme. First, during the *phase selection* the sampling strategy is determined. Subsequently, in the *input parameter sample* block input samples in the form of human-exoskeleton simulation instances are drawn. The *output sample* block illustrates the generation of output samples using the automated assessment process. Lastly, the input-output samples are used to obtain sensitivity measures which is visualized in the *sensitivity analysis*.

2 MATERIALS AND METHODS

118 In this section, the technical problem is formulated and the relevant material and methods are shown. An
 119 overview of the proposed uncertainty quantification procedure is shown in Figure 1. From top to bottom the
 120 colored blocks illustrate the *phase selection*, the process of obtaining *input parameter samples*, the process
 121 of obtaining *output samples* and the evaluation procedure using quantitative *sensitivity analysis* methods.
 122 First, during the *phase selection* the sampling strategy is determined, which is chosen in accordance to the
 123 objective of the respective sensitivity analysis method. Following this, the *input parameter samples* are
 124 generated. Here, the examined sources of uncertainty are sampled depending on the previously selected
 125 sampling strategy. Then, the input parameter samples are retrieved in the form of parameterized human-
 126 exoskeleton simulation instances, where the varied parameters are associated with different sources of
 127 uncertainty. Subsequently, the *output sample* block is applied. Here, the exoskeleton-based automated
 128 assessment is run for the sampled simulation parameterizations to obtain impedance parameter estimates for
 129 the human arm. Finally, the *sensitivity analysis* is performed. Depending on the sampling strategy chosen
 130 beforehand, different sensitivity analysis methods are deployed on the estimated impedance parameters
 131 to investigate the impact of the modelling uncertainties with respect to the observed estimation error. By
 132 deploying this sensitivity analysis scheme we are able to derive the most influential sources of uncertainty
 133 that influence the exoskeleton-based arm impedance estimation.

134 The remainder of the section is structured as follows: In Section 2.1, the dynamics governing the human-
 135 exoskeleton system are introduced and a qualitative account on uncertainties in the automated assessment
 136 is provided. Subsequently, a high-fidelity simulation of the human-exoskeleton interaction is presented
 137 in Section 2.2 with particular focus on including the key sources of uncertainty present in the system. In
 138 Section 2.3, the proposed assessment procedure is explained and technical details regarding the estimation
 139 process are provided. Finally, in Section 2.4, the deployed sampling strategies and sensitivity analysis
 140 methods are presented.

141 2.1 Uncertainty during Human-Exoskeleton Interaction

142 In order to perform the sensitivity analysis in an interpretable manner it is necessary to have an
 143 understanding of the investigated system. To this end, we first formulate the nominal human-exoskeleton
 144 interaction model. Subsequently, uncertainties are introduced to the nominal model. Finally, an automated
 145 spasticity assessment scheme is described and the impact of uncertainties on the assessment result is
 146 investigated.

147 2.1.1 Nominal Human-Exoskeleton Interaction Model

148 The instrumented assessment using an upper-limb exoskeleton is considered in this work. Therefore, we
 149 start by establishing the dynamics governing motion of the human arm. We model the dynamics using
 150 Euler-Lagrange equations (Featherstone, 2007) of the form,

$$M_h(\mathbf{q})\ddot{\mathbf{q}} + C_h(\mathbf{q}, \dot{\mathbf{q}})\dot{\mathbf{q}} + \mathbf{g}_h(\mathbf{q}) = \boldsymbol{\tau}_{\text{hum}} + \boldsymbol{\tau}_{\text{int,h}}. \quad (1)$$

151 Here, $\mathbf{q} \in \mathbb{R}^d$ is the d-dimensional state vector containing the joint configuration of the human arm, with
 152 $\dot{\mathbf{q}} \in \mathbb{R}^d$ describing the angular velocities and $\ddot{\mathbf{q}} \in \mathbb{R}^d$ describing the angular accelerations. On the left
 153 side of (1) the matrix $M_h: \mathbb{R}^d \rightarrow \mathbb{R}^{d \times d}$ denotes the human inertia matrix, $C_h: \mathbb{R}^d \times \mathbb{R}^d \rightarrow \mathbb{R}^{d \times d}$ the
 154 human Coriolis matrix and $\mathbf{g}_h: \mathbb{R}^d \rightarrow \mathbb{R}^d$ the human gravitational component. In addition to the human
 155 generated joint torques $\boldsymbol{\tau}_{\text{hum}}$, an interaction torque $\boldsymbol{\tau}_{\text{int,h}}$ acts on the human arm, due to the contact with
 156 the robotic system. In (1), $\boldsymbol{\tau}_{\text{hum}}$ represents the projected joint-level torques induced through variations of
 157 muscle lengths, muscle activation and the resulting tensions (Shin et al., 2009). Therefore, $\boldsymbol{\tau}_{\text{hum}}$ describes

158 the summed dynamics of internal origin and contains the relevant joint dynamics parameter necessary
 159 to quantify the patient's inner state. In the case of stroke, a viscoelastic model of the human-generated
 160 torque during passive mobilization tasks is proposed (McCrea et al., 2003). Thus, we can formulate the
 161 human-generated torque τ_{hum} as

$$\tau_{\text{hum}} = \mathbf{K}_h(\mathbf{q}, \dot{\mathbf{q}})\mathbf{q} + \mathbf{D}_h(\mathbf{q}, \dot{\mathbf{q}})\dot{\mathbf{q}}, \quad (2)$$

162 where $\mathbf{K}_h: \mathbb{R}^d \times \mathbb{R}^d \rightarrow \mathbb{R}^{d \times d}$ and $\mathbf{D}_h: \mathbb{R}^d \times \mathbb{R}^d \rightarrow \mathbb{R}^{d \times d}$ correspond to the joint stiffness and viscosity
 163 matrix, respectively. In McCrea et al. (2003) the validity of linear viscoelasticity parameters for the
 164 modelling of resistive torques in personas with chronic stroke is demonstrated. Therefore, it can additionally
 165 be assumed that the parameters are independent of the current configuration, which allows the application of
 166 standard regression methods. Thus, the instrumented assessment of the patient's state can be reformulated
 167 as a linear regression problem using the parametric model

$$\tau_{\text{hum}} = \mathbf{K}_h \mathbf{q} + \mathbf{D}_h \dot{\mathbf{q}}. \quad (3)$$

168 In order to estimate the impedance parameters \mathbf{K}_h and \mathbf{D}_h , it is first necessary to extract the human
 169 generated torque τ_{hum} in (1). This is not trivial in general, as the intrinsically generated human muscle
 170 torque cannot be measured directly. Hence, τ_{hum} has to be inferred using the available measurements and
 171 dynamics knowledge. For wearable robots deployed in clinical applications, measurements regarding joint
 172 positions and motor torques are typically available (e.g., Trigili et al., 2020). Unless additional expensive
 173 and possibly inconvenient force-torque sensors are mounted at the physical interface between human and
 174 exoskeleton (An and Hollerbach, 1987), the interaction torque $\tau_{\text{int,h}}$ is also unknown. To overcome this issue,
 175 knowledge regarding the dynamics model of the robotic system can be exploited to replace the unknown
 176 interaction torque $\tau_{\text{int,h}}$. Similar to the human, the exoskeleton is described by its rigid body dynamics

$$\mathbf{M}_e(\boldsymbol{\theta})\ddot{\boldsymbol{\theta}} + \mathbf{C}_e(\boldsymbol{\theta}, \dot{\boldsymbol{\theta}})\dot{\boldsymbol{\theta}} + \mathbf{g}_e(\boldsymbol{\theta}) = \tau_{\text{m}} - \tau_{\text{int,e}}, \quad (4)$$

177 where $\mathbf{M}_e: \mathbb{R}^n \rightarrow \mathbb{R}^{n \times n}$ is the inertia, $\mathbf{C}_e: \mathbb{R}^n \times \mathbb{R}^n \rightarrow \mathbb{R}^{n \times n}$ the Coriolis matrix and $\mathbf{g}_e: \mathbb{R}^n \rightarrow \mathbb{R}^n$ the
 178 gravitational component of the exoskeleton dynamics. The joint positions, velocities and accelerations of
 179 the robotic system are given by $\boldsymbol{\theta} \in \mathbb{R}^n$, $\dot{\boldsymbol{\theta}} \in \mathbb{R}^n$ and $\ddot{\boldsymbol{\theta}} \in \mathbb{R}^n$ respectively. In the following, we assume
 180 that the kinematic chain of human and exoskeleton align, thereby, resulting in $n = d$. Furthermore, the
 181 movement of the joints is driven by the motor torques τ_{m} and analogue to (1), an interaction torque $\tau_{\text{int,e}}$
 182 is exerted on the exoskeleton, which acts in the opposing direction in (4).

183 In the nominal model, three idealized assumptions are made: first, a perfect alignment of the human
 184 and exoskeleton kinematic chain is assumed. Second, no displacement of the attachments occurs during
 185 movement. Third, a completely rigid interface transmits forces between the human and exoskeleton. If these
 186 assumptions hold, both the human's and exoskeleton's joint kinematics match $\mathbf{q} = \boldsymbol{\theta}$ and the interaction
 187 torques can be written to

$$\tau_{\text{int,h}} = \tau_{\text{int,e}}. \quad (5)$$

For the sake of the derivation of the nominal model we hypothesize the dynamics of the robotic system and human to be known. Then, it is possible to derive the human generated torque τ_{hum} from (1), (4) and (5):

$$\tau_{\text{hum}} = \mathbf{M}_h(\boldsymbol{\theta})\ddot{\boldsymbol{\theta}} + \mathbf{C}_h(\boldsymbol{\theta}, \dot{\boldsymbol{\theta}})\dot{\boldsymbol{\theta}} + \mathbf{g}_h(\boldsymbol{\theta}) + \underbrace{\mathbf{M}_e(\boldsymbol{\theta})\ddot{\boldsymbol{\theta}} + \mathbf{C}_e(\boldsymbol{\theta}, \dot{\boldsymbol{\theta}})\dot{\boldsymbol{\theta}} + \mathbf{g}_e(\boldsymbol{\theta}) - \tau_{\text{m}}}_{\tau_{\text{int,h}}} \quad (6)$$

188 Since the motor torque τ_m and exoskeleton kinematics $\{\theta, \dot{\theta}, \ddot{\theta}\}$ are measurable and the dynamics are
 189 assumed to be known, the human torque τ_{hum} , as given in (6), is directly computable. Therefore, all the
 190 necessary input and output information are available to estimate the human joint viscoelasticity parameters
 191 K_h and D_h via linear regression using the parametric model (3):

$$y = X\omega, \tag{7}$$

192 where the labels y follows from the human torque computation according to (6), the input matrix X
 193 contains the human joint measurements under the assumption that $q = \theta$ and the viscoelasticity parameters
 194 of interest are described by ω . Thereby, performing the regression analysis for each joint yields

$$\underbrace{\begin{bmatrix} \tau_{\text{hum},i}^1 \\ \tau_{\text{hum},i}^2 \\ \vdots \\ \tau_{\text{hum},i}^T \end{bmatrix}}_y = \underbrace{\begin{bmatrix} q_i^1 & \dot{q}_i^1 \\ q_i^2 & \dot{q}_i^2 \\ \vdots & \vdots \\ q_i^T & \dot{q}_i^T \end{bmatrix}}_X \underbrace{\begin{bmatrix} k_{h,ii} \\ d_{h,ii} \end{bmatrix}}_\omega, \tag{8}$$

195 with $\{\tau_{\text{hum},i}^t\}_{t=1}^T$ denoting the computed human torques and $\{q_i^t, \dot{q}_i^t\}_{t=1}^T$ representing the kinematics
 196 measurements of the i -th joint at discrete time step t over the duration T of the assessment. Here, k_{ii} and d_{ii}
 197 are the i -th main diagonal entries of the joint stiffness and viscosity matrices, respectively. The parameter
 198 vector ω can be computed directly given access to inputs X and labels y as such:

$$\omega = (X^T X)^{-1} X^T y. \tag{9}$$

199 However, while the approach is mathematically convenient and can straight forwardly be implemented,
 200 it can result in large estimation errors, because it does not account for the uncertainties in the human-
 201 exoskeleton interaction dynamics.

202 2.1.2 Sources of Uncertainty

203 There are multiple factors that introduce uncertainties to the above described nominal model, which
 204 stem from variations in the biomechanics of individuals. In particular three key sources of uncertainty
 205 that adversely affect the physical interaction are identified in the literature (Pons, 2008): kinematic
 206 incompatibilities, soft contact dynamics and inaccuracies in the nominal dynamics model. In the following
 207 these sources of uncertainty and their impact on the nominal dynamics are presented in more detail.

208 Kinematic Incompatibilities

209 First, we consider kinematic incompatibilities between the exoskeleton and human, which are particularly
 210 prevalent in wearable robots with kinematic chains mirroring the human kinematics. These kinematic
 211 incompatibilities arise due to anatomical variations between users and variations within a user that occur
 212 during motion. Therefore, achieving a perfect alignment is infeasible (Jarrassé and Morel, 2012). Depending
 213 on the extent of the mismatch, it is considered a macro-misalignment or a micro-misalignment. Here,
 214 macro-misalignments are typically induced by offsets in the center of rotation (CoR) between the human
 215 and exoskeleton joints. These CoR offsets are the result of a multiple factors, such as an imprecise donning
 216 procedure or translations that occur in the instantaneous center of rotation of human joints for certain
 217 movements (Grant, 1973). In Figure 2A, the macro-misalignment due to CoR offsets is shown conceptually
 218 for a simplified two-link human-exoskeleton-system moving in the vertical plane. The top and bottom
 219 links represent the upper arm and forearm, respectively, emulating motion in flexion/extension direction.

220 Here, the CoR offsets are visualized by x_{off} and y_{off} using red arrows. While macro-misalignment can
 221 be reduced by performing careful donning and including redundant DoFs in the robotic kinematic chain,
 222 micro-misalignments still occur despite these mitigation strategies. This is for instance because the human
 223 kinematic chain is not comprised of idealized, circular joints. Therefore, misalignments cannot be removed
 224 completely in practice and must be explicitly considered for a robust automated assessment.

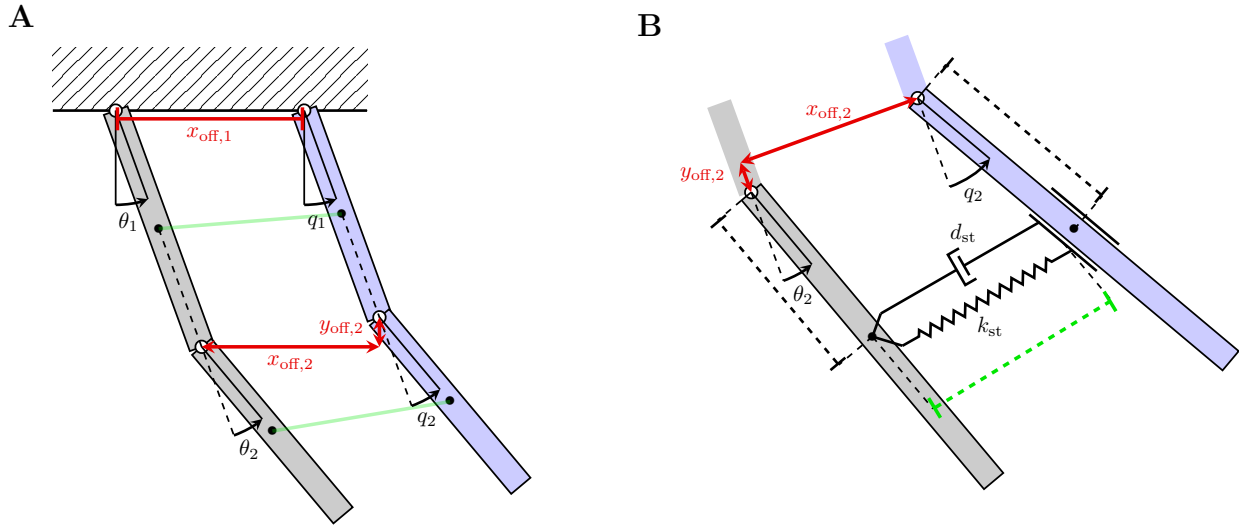


Figure 2. Two-link mechanical model of an interaction between a human (blue) and exoskeleton (grey) arm. Sub figure (A) illustrates kinematic incompatibilities and the resulting CoR offsets depicted with x_{off} and y_{off} . Sub figure (B) visualizes soft coupling between the human and exoskeleton link using a Voight-element.

225 The main consequence of these kinematic incompatibilities is induced displacements of the attachments
 226 between the exoskeleton and human limb during joint motion. Consequently, these displacements result in
 227 forces at the physical interface. The resulting impact on the nominal dynamics of the human-exoskeleton
 228 interaction can be observed at multiple points. First, the previously assumed joint alignment does not hold
 229 anymore, leading to a discrepancy in the joint angles, i.e., $\mathbf{q} \neq \boldsymbol{\theta}$ in general. Moreover, an offset and joint
 230 angle dependent displacement of the attachments along the axial direction occurs, which leads to a change
 231 in the interaction torque transmission (5):

$$\tilde{\boldsymbol{\tau}}_{\text{int,h}} = \mathbf{B}(\mathbf{x}_{\text{off}}, \mathbf{y}_{\text{off}}, \mathbf{q}, \boldsymbol{\theta}) \boldsymbol{\tau}_{\text{int,e}}, \quad (10)$$

232 where $\mathbf{B}: \mathbb{R}^{d \times d}$ is a d -dimensional diagonal matrix with the main diagonal entries describing the displaced
 233 attachment points. In (10), $\tilde{\boldsymbol{\tau}}_{\text{int,h}}$ represents the uncertain interaction torques which now depends on the
 234 CoR offsets denoted by \mathbf{x}_{off} and \mathbf{y}_{off} . Similarly, the induced displacement torques depend on the CoR
 235 offsets and joint angles deviations (Schiele, 2008). Therefore, we obtain following uncertain human torque
 236 under consideration of kinematic incompatibilities:

$$\tilde{\boldsymbol{\tau}}_{\text{hum}} = \mathbf{M}_h(\mathbf{q})\ddot{\mathbf{q}} + \mathbf{C}_h(\mathbf{q}, \dot{\mathbf{q}}) + \mathbf{g}_h(\mathbf{q}) + \tilde{\boldsymbol{\tau}}_{\text{int,h}}(\mathbf{x}_{\text{off}}, \mathbf{y}_{\text{off}}, \mathbf{q}, \boldsymbol{\theta}) + \tilde{\boldsymbol{\tau}}_d(\mathbf{x}_{\text{off}}, \mathbf{y}_{\text{off}}, \mathbf{q}, \boldsymbol{\theta}), \quad (11)$$

237 where $\tilde{\boldsymbol{\tau}}_d$ denotes the uncertain displacement torques. In addition to $\tilde{\boldsymbol{\tau}}_{\text{int,h}}$ and $\tilde{\boldsymbol{\tau}}_d$, uncertainty also arises in
 238 (11) due to the dependence on \mathbf{q} , since the human joint angle cannot be measured directly and cannot be
 239 inferred accurately from $\boldsymbol{\theta}$, since $\mathbf{q} = \boldsymbol{\theta}$ no longer holds. Note that, given completely rigid bodies, these

240 kinematic incompatibilities would theoretically make movements impossible and lead to extremely high
 241 interaction forces, due to the kinematic system being hyperstatic (Jarrassé and Morel, 2012). However, in
 242 practice deformation occurs at the physical interface, since the human limb is not rigid, which allows to
 243 retain mobility. The uncertainty that arises due to this plasticity is addressed in the following.

244 *Soft-tissue Contact Dynamics*

245 The second important aspect that introduces uncertainty to the physical human-exoskeleton interaction
 246 are morphological factors at the coupling between the robot and human. Specifically, the robotic system
 247 induces the desired movement by transmitting forces through the soft-tissue of the human limb at the
 248 attachment straps. Here, the considered soft-tissue primarily includes muscles, fat tissue and skin, but may
 249 also include smaller anatomical parts, such as ligaments, tendons or blood vessels. This is in contrast to the
 250 nominal dynamics model which assumes a rigid connection (11). Therefore, the dynamic properties of the
 251 human soft-tissue impact the description of the physical interaction.

252 Soft-tissue is most commonly modelled by elastic or viscoelastic components (Maurel, 1999). Viscoelastic
 253 dynamic behavior can for instance be represented by Voight-elements as illustrated in Figure 2B. Here, the
 254 soft coupling between the human and exoskeleton link is achieved via a Voight-element at the attachment.
 255 Hence, the displacement torques $\tilde{\tau}_d$ and the interaction torque $\tilde{\tau}_{\text{int,h}}$ become functions of the viscoelastic
 256 parameters, since all interaction forces are transmitted through soft contacts. It leads to

$$\tilde{\tau}_{\text{hum}} = \mathbf{M}_h(\mathbf{q})\ddot{\mathbf{q}} + \mathbf{C}_h(\mathbf{q}, \dot{\mathbf{q}}) + \mathbf{g}_h(\mathbf{q}) + \tilde{\tau}_{\text{int,h}}(\mathbf{x}_{\text{off}}, \mathbf{y}_{\text{off}}, \mathbf{q}, \boldsymbol{\theta}, \mathbf{K}_{\text{st}}, \mathbf{D}_{\text{st}}) + \tilde{\tau}_d(\mathbf{x}_{\text{off}}, \mathbf{y}_{\text{off}}, \mathbf{q}, \boldsymbol{\theta}, \mathbf{K}_{\text{st}}, \mathbf{D}_{\text{st}}), \quad (12)$$

257 where \mathbf{K}_{st} and \mathbf{D}_{st} denote the lumped viscoelastic properties of the coupling due to soft-tissue. In (Schiele,
 258 2008) a more detailed analysis of the displacement forces and their transmission through soft-tissue
 259 modelled as Voight-elements is presented. However, while linear, uniaxial models as shown in (12)
 260 are used for practicality, they describe the complex relationship between applied pressure and resulting
 261 deformation of the soft-tissue in a simplified manner. A more rigorous approach is to use discrete finite
 262 element to approximate the continuous medium and propagating the evolution of the deformation in
 263 simulations (Maurel et al., 2002). However, since this is an iterative procedure, it cannot straightforwardly
 264 be translated to an analytical model.

265 *Inaccuracies in the Human Dynamics Model*

266 Another source of uncertainty that needs to be considered are inaccuracies in the human dynamics
 267 model. This is due to significant variations in the biomechanics of each human. To mitigate this, precise
 268 measurements of geometrical and inertial properties of the anatomical links are necessary to compute
 269 the personalized model parameters required for the human rigid body dynamics (1). However, gathering
 270 the information needed to estimate the human model parameter can be expensive, cumbersome and
 271 time-intensive (Zajac et al., 2002). Therefore, in clinical practice most commonly standard tables of
 272 anthropometric parameters are used (de Leva, 1996) to infer model parameters by scaling the default
 273 dynamics model to the height and weight of a particular individual. However, since the approach only
 274 yields an approximate measure, uncertainties are introduced. Thus, the uncertain human torque $\tilde{\tau}_{\text{hum}}$ under
 275 additional consideration of the modelling inaccuracies is

$$\tilde{\tau}_{\text{hum}} = \tilde{\mathbf{M}}_h(\mathbf{q})\ddot{\mathbf{q}} + \tilde{\mathbf{C}}_h(\mathbf{q}, \dot{\mathbf{q}}) + \tilde{\mathbf{g}}_h(\mathbf{q}) + \tilde{\tau}_{\text{int,h}}(\mathbf{x}_{\text{off}}, \mathbf{y}_{\text{off}}, \mathbf{q}, \boldsymbol{\theta}, \mathbf{K}_{\text{st}}, \mathbf{D}_{\text{st}}) + \tilde{\tau}_d(\mathbf{x}_{\text{off}}, \mathbf{y}_{\text{off}}, \mathbf{q}, \boldsymbol{\theta}, \mathbf{K}_{\text{st}}, \mathbf{D}_{\text{st}}), \quad (13)$$

276 where $\tilde{\mathbf{M}}_h$, $\tilde{\mathbf{C}}_h$ and $\tilde{\mathbf{g}}_h$ denote the uncertain inertial, Coriolis and gravitational component of the human arm
 277 dynamics, which differ from the approximation obtained from the anthropometric tables. We summarize

278 the torque due to the uncertain passive dynamics of the human limb with

$$\tilde{\tau}_{\text{rbd,h}} = \tilde{M}_h(\mathbf{q})\ddot{\mathbf{q}} + \tilde{C}_h(\mathbf{q}, \dot{\mathbf{q}}) + \tilde{g}_h(\mathbf{q}). \quad (14)$$

279 Thereby, we can write (13) to a more compact form for improved readability

$$\tilde{\tau}_{\text{hum}} = \tilde{\tau}_{\text{rbd,h}} + \tilde{\tau}_{\text{int,h}} + \tilde{\tau}_{\text{d}}. \quad (15)$$

280 Here, $\tilde{\tau}_{\text{rbd,h}}$ denotes the uncertain rigid body dynamics of the human arm due to unknown parameters in
 281 \tilde{M}_h , \tilde{C}_h and \tilde{g}_h . Differently to the human limb, the model parameters governing the dynamics of the
 282 exoskeleton (4) can reasonably be assumed to be known or can be obtained accurately using classical
 283 identification procedures (Hollerbach et al., 2008). Note that in (15), both $\tilde{\tau}_{\text{int,h}}$ and $\tilde{\tau}_{\text{d}}$ are in principle
 284 torques that are induced by the interaction with the exoskeleton. However, they differ in the sense that $\tilde{\tau}_{\text{int,h}}$
 285 represents the desired loads that should be transmitted to the human limb, while $\tilde{\tau}_{\text{d}}$ are purely undesired
 286 torques due to kinematic incompatibilities. Since the human torque under consideration of uncertainties
 287 $\tilde{\tau}_{\text{hum}}$ (15) differs from the nominal human torque τ_{hum} (6) used in the regression analysis (8), errors are
 288 introduced to the estimated impedance parameters. In particular, deploying (6) for the computation of the
 289 human torque τ_{hum} implicitly allocates torques that are unaccounted for by the nominal dynamics model
 290 to be generated due to joint spasticity. Thus, solving the regression problem will not result in the true
 291 viscoelasticity parameter \mathbf{K}_h and \mathbf{D}_h . By directly comparing the nominal human torque τ_{hum} to the true,
 292 uncertain human torque $\tilde{\tau}_{\text{hum}}$, we obtain

$$\underbrace{\tau_{\text{hum}}}_{\mathbf{y}} = \underbrace{\tilde{\tau}_{\text{hum}}}_{\tilde{\mathbf{y}}} - \underbrace{\Delta\tau_{\text{rbd,h}} - \Delta\tau_{\text{int,e}} - \tilde{\tau}_{\text{d}}}_{\Delta\mathbf{y}}. \quad (16)$$

293 Here, $\Delta\tau_{\text{rbd,h}}$ denotes residual torques due to differences in the nominal human dynamics model $\tau_{\text{rbd,h}}$ and
 294 the unknown, true dynamics model $\tilde{\tau}_{\text{rbd,h}}$. Similarly, $\Delta\tau_{\text{int,e}}$ represents residual torques due to errors in the
 295 interaction torque modelling, while $\tilde{\tau}_{\text{d}}$ are the displacement torques due to kinematic incompatibilities.
 296 From (16) it can be seen that the labels \mathbf{y} deployed in (8) do not agree with the true output $\tilde{\mathbf{y}}$, i.e., the
 297 human torque $\tilde{\tau}_{\text{hum}}$ under consideration of uncertainties. The difference is summarized in (16) using $\Delta\mathbf{y}$.
 298 Moreover, the measurements for the desired input matrix \mathbf{X} according to (8) are not available, since
 299 kinematic incompatibilities result in a mismatch between the human joint angle \mathbf{q} and exoskeleton joint
 300 angle $\boldsymbol{\theta}$. Hence, it can be seen how the uncertainties qualitatively influence the outcome of the regression
 301 analysis and impact the automated assessment negatively. However, it remains unclear exactly how sensitive
 302 the assessment is with respect to the different sources of uncertainty, which we propose to quantify with a
 303 sampling-based sensitivity analysis in this work.

304 2.2 High-fidelity Human-Exoskeleton Simulation

305 In order to perform a sampling-based sensitivity analysis, a highly controlled environment is required.
 306 Obtaining the samples experimentally is infeasible, due to the missing ground-truth information and the
 307 large sample size that is required. Therefore, in this work we deploy a high-fidelity simulation environment
 308 of the human-exoskeleton system to generate samples. To this end, we develop a novel human-exoskeleton
 309 simulation which explicitly accounts for the complex contact dynamics present during physical interaction.
 310 Here, an optimization-based physics engine called MuJoCo (Todorov et al., 2012) is deployed which is
 311 widely used in the modelling of robotic and biomechanical systems in contact-rich environments (Acosta
 312 et al., 2022; Lowrey et al., 2016). In particular, three key features of the proposed simulation enable the

313 realistic emulation of the effects caused by sources of uncertainty and thereby facilitate the sampling-based
314 sensitivity analysis: A musculoskeletal model to simulate the human, the consideration of soft contact
315 dynamics at the attachments and a realistic load transmission via a mechanical interface. The proposed
316 human-exoskeleton simulation is shown in Figure 1 in the *input parameter sample* block. Here, the human
317 skeletal system is depicted in gray, while the muscular system is visualized with red lines. Furthermore, the
318 two red cylindrical shapes on the forearm and upper arm represent the simulated human soft-tissue. Also, it
319 can be seen that the physical interface is realized via cuffs and straps that wrap around the human upper and
320 forearm. The complete human-exoskeleton simulation environment is made publicly available¹. A brief
321 summary of the key components is presented below. Following this, a more detailed explanation of each of
322 the components of the simulation, their working principles and the performed validations is provided.

323 **Human Musculoskeletal Model:** A musculoskeletal model is implemented for the shoulder and elbow.
324 Deploying a musculoskeletal model of the human arm here is necessary for two reasons. First, the
325 simulated muscular system is used to generate the human torque and emulate spastic behavior. Second,
326 the rigid skeletal system facilitates the introduction of variability in the human kinematics and dynamics.
327 Thereby, it is possible to sample over two of the three sources of uncertainty described in Section 2.1.2.

328 **Soft-tissue Simulation:** In the proposed simulation, soft-tissue is explicitly implemented by a composition
329 of multiple micro-elements, which together form an object with viscoelastic material properties. The
330 viscoelastic properties of the soft-tissue object can be varied, thereby allowing to sample over viscoelastic
331 properties of the soft-tissue.

332 **Physical Human-Exoskeleton Interface:** We simulate the mechanical interface explicitly by implementing
333 cuffs and straps, which enclose the human arm and facilitate a realistic load transmission. Thereby effects
334 that typically arise at the interface, such as attachment displacements, can be emulated.

335 2.2.1 Simulation of the Human Musculoskeletal System

336 A musculoskeletal model is used in the proposed simulation environment. Here, the rigid component of
337 the human arm has five DoFs, three on the shoulder joint and two at the elbow joint. For the shoulder, the
338 human simulation can rotate along the flexion-extension, abduction-adduction and internal-external axis.
339 Regarding the elbow, the simulation allows movement along the flexion-extension and pronation-supination
340 rotations. While a rigid wrist-hand model is also included in the simulation, in our envisioned interaction
341 scenario with the exoskeleton it is not pertinent. The inertial properties of the rigid skeletal system are
342 designed using statistical anthropometric data (Ramachandran et al., 2016) with a default reference person
343 of height 1.75m weighting 70kg. Thereby resulting in a nominal upper arm length of 36.37cm, a nominal
344 forearm length of 34.9cm, a nominal upper arm mass of 2.25kg and a nominal forearm mass of 1.31kg.
345 However, it is possible to adjust all of the parameters to account for variations in the target population.

346 In addition to the multi-link rigid body dynamics, the simulation accounts for the dynamics induced by
347 the muscular system. In MuJoCo, biological muscles are modelled by means of muscle-tendon systems
348 which induce dynamics dependent on origin and insertion sited and the forces generated by a muscle
349 actuator. Here, the generated muscle force F_m follows the dynamics

$$F_m(l, v, a) = -F_0 F_{lv}(l, v, a), \quad (17)$$

¹ Open-source code of the upper-limb human-exoskeleton simulation environment is available at: https://github.com/stesfazgi/rehyb_mujoco

350 where l is the scaled length of the muscle, v is the the scaled velocity and $a \in [0, 1]$ denotes the muscle
 351 activation level. Additionally, F_0 describes the peak active force and F_{lv} the force-length-velocity function,
 352 which are both fitted according to values derived from the experimental findings in (Holzbaur et al., 2005).
 353 The origin and insertion sites of the muscles are also implemented in accordance with anthropometric
 354 data (Ramachandran et al., 2016), thereby ensuring that the dynamics of the simulated musculoskeletal
 355 system follow the real-world dynamics closely.

356 Validation of the Human Musculoskeletal Model

357 In order to check the validity of the simulated human musculoskeletal model, a simulation experiment
 358 is performed. Specifically, it is examined whether the moments generated by the muscular system lie in
 359 similar ranges as those observed in real experiments. A common clinical procedure to assess the muscle
 360 strength is by means of the maximal isometric torque test (Amis et al., 1980; Garcia et al., 2016). Here,
 361 we use this procedure to adapt and validate the simulated elbow muscle contraction, which is a useful
 362 measure to quantify the neuromuscular properties of spastic muscles (Wang et al., 2019). In the proposed
 363 simulation, the dynamics of the elbow are governed by eight muscles. Specifically, four extensor muscles
 364 are considered, namely, the long, lateral and median triceps and the anconeus. Moreover, four flexor
 365 muscles are regarded, including the long and short biceps, the brachialis and the brachioradialis. The
 366 experimental procedure for the isometric torque test in flexion direction is as follows: First, the shoulder is
 367 flexed in the sagittal plane at 90 deg and mechanically locked in this configuration. While the shoulder is
 368 fixed in place, the elbow is flexed in discrete steps of 1 deg increments. At each of the discrete increments
 369 a maximum contraction of the elbow flexor muscles is applied, and the resulting torque is measured.

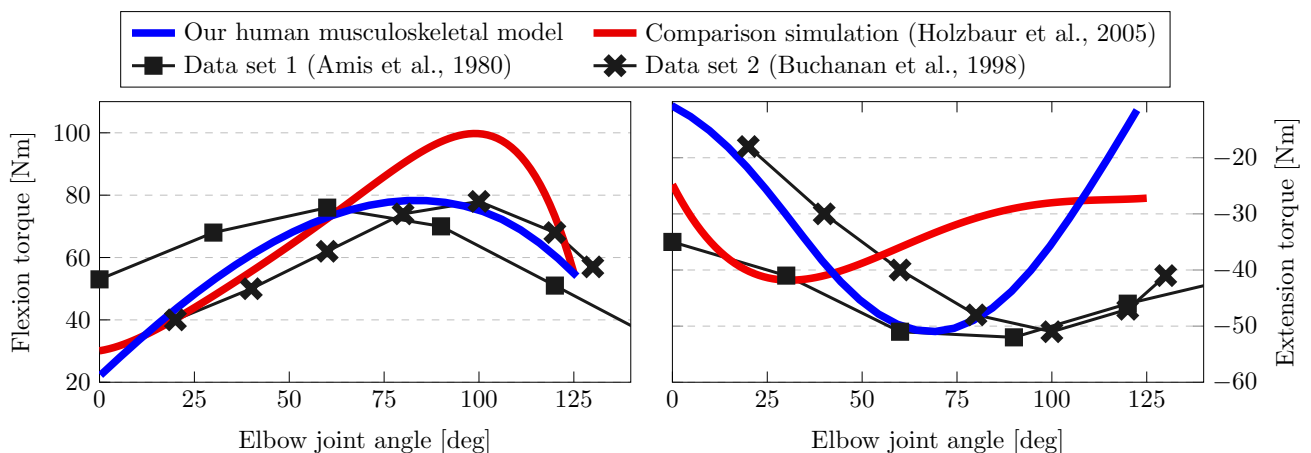


Figure 3. Results of the maximum isometric torque test. Here, the torque generated by the elbow flexors (left) and extensors (right) is shown over different elbow joint angle. Our human musculoskeletal model (blue) is shown to agree more closely with experimental data than the comparison simulation (red).

370 The results of performing the maximum isometric torque test in the simulation are shown in Figure 3.
 371 Here, the left-hand side shows the isometric flexion torque, while the right side depicts the extension torque.
 372 We compare our simulation results (blue) against related biomechanical models of the musculoskeletal
 373 system (Holzbaur et al., 2005) (red) and two experimental data sets (Amis et al., 1980; Buchanan et al.,
 374 1998). For the isometric flexion torque on the left, it is possible to see that our simulation results match the
 375 observed maximum torque of around 80 Nm closely, while the comparison simulation exhibits a higher
 376 peak at 100 Nm. Analogously, our simulation obtains a similar value for the peak extension torque as
 377 the experimental data set at -50 Nm, while the simulation in (Holzbaur et al., 2005) results in a lower

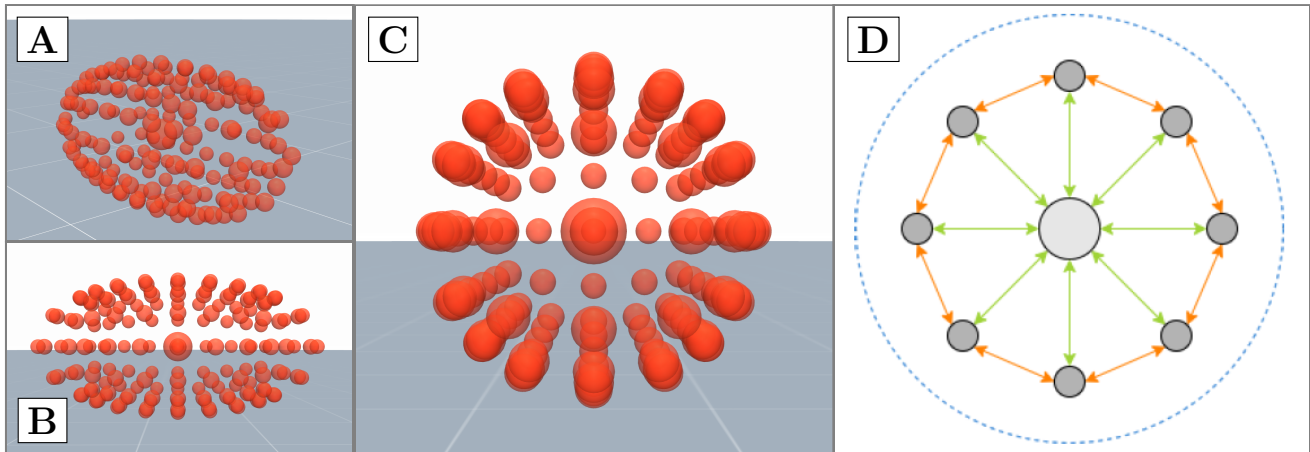


Figure 4. Sub figure (A), (B) and (C) depict a composite object with an ellipsoid shape from different viewing angles. (D) A cross-section of the composite object with the central element in light gray and external elements in dark gray. Three types of soft constraints hold the elements together: central-external constraints (green), external neighbors constraints (orange), and a global volume constraint (blue).

378 absolute value at at -41 Nm. With respect to the curve shape both data set 1 (Amis et al., 1980) and data
 379 set 2 (Buchanan et al., 1998) display different behaviors. This is to be expected due to variability in real
 380 experiments and between different subjects, however, the simulation results indicate that our model lies
 381 within this range. Particularly, when observing the joint angle at which the peak extension torque is reached
 382 for instance, it is clearly visible that our simulation agrees with the experimental data more closely.

383 2.2.2 Simulation of the Upper-limb Exoskeleton

384 In this work, the simulated robotic system is inspired by the specification detailed in Trigili et al. (2020),
 385 where an upper-limb exoskeleton with three actuated DoFs on the shoulder level and one actuated DoF
 386 for the elbow (flexion-extension) is presented. For the envisioned scenario, we consider all passive and
 387 regulatory DoFs to be fixed, therefore, the simulated upper-limb exoskeleton is a four-DoF open chain.
 388 Joint friction is implemented via viscous dampers and the inertial properties are designed to roughly match
 389 comparable robotic devices. While each joint is associated with an actuator in the simulation, we do not
 390 consider elastic actuators here. The actuating motors are also scaled in accordance with the maximum
 391 torques the real system can provide according to Trigili et al. (2020). Note that while the simulated
 392 exoskeleton is inspired by (Trigili et al., 2020), this represents an exemplary device and may be replaced
 393 by a different wearable robotic system of interest. The proposed method for the spasticity assessment and
 394 sensitivity analysis constitute a general methodology and are therefore not limited to this specific hardware
 395 and could be applied to other exoskeleton designs as well.

396 2.2.3 Physical Interface and Complex Contact Dynamics

397 In our simulation, the physical interface is composed of two contact areas which represent the exoskeleton
 398 attachments on the upper and lower arm of the human. On the human side, complexity of the contact
 399 dynamics is primarily caused by soft-tissues and their influence on the force transmission at the linkage
 400 between the human arm and exoskeleton. In order to replicate the behavior of human soft-tissue in the
 401 simulation, three-dimensional composite objects are used, where one central element is surrounded by
 402 multiple external elements. Here, the elements of the three-dimensional composite object are arranged
 403 such that the resulting geometry approximates the human limb shape and thus a simplification of the
 404 commonly used finite element method (Maurel et al., 2002) is achieved. Figure 4A, B and C depicts the
 405 composite object which takes an ellipsoid shape in the simulation environment, where the large sphere

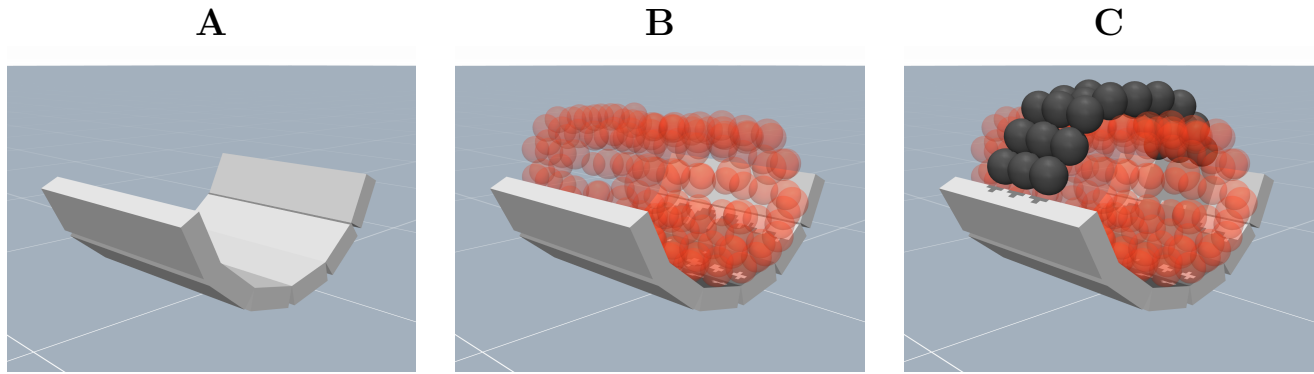


Figure 5. Depiction of the mechanical support at the physical interface in the simulation environment. (A) An illustration of the semi-cylindrical cuff composed of welded box primitives. (B) An illustration of the placement of the human limb within the cuff. (C) The implementation of straps using composite objects to fix the limb to the semi-cylinder.

406 at the center of the ellipsoid visualizes the central element of the composite object, while the external
 407 elements are illustrated by the smaller spheres. The viscoelastic behavior of the resulting composite object
 408 is determined by several soft equality constraints on the relative distance between the different elements,
 409 which is illustrated in Figure 4D. Each soft equality constraint generates a force that can be approximately
 410 interpreted as a spring-damper link between two elements. Additionally, one constraint acting on all the
 411 elements is set to preserve the global volume of the composite object. The parameters of all constraints are
 412 fitted to approximate the viscoelastic behaviour of real human soft-tissue.

413 On the exoskeleton side, forces are generally transmitted to the human arm via the mechanical supports,
 414 e.g., cuffs and straps, which induce movement by pushing or pulling the limb (Pons, 2008). Therefore,
 415 we follow the same design principle in the simulation in order to render the contact dynamics in high
 416 fidelity. First, the arm supports are implemented using a hollow semi-cylinder shape. Since MuJoCo
 417 does not directly handle concave bodies, the desired shape is approximately realized by an arrangement
 418 of welded box primitives (Figure 5A). Second, the human arm is placed inside the support (Figure 5B).
 419 Third, the implementation of the arm straps is realized using composite objects which are arranged in a
 420 two-dimensional grid. By welding two opposing sides of the strap to the arm support, the human limb is
 421 fixed to the attachment as illustrated in Figure 5C.

422 *Validation of the Human-Exoskeleton Contact Dynamics*

In order to validate the geometric compliance of the simulated limb, the stress-strain relationship of the composite object is investigated in the form of a compression test. In the validation, a uniaxial tension is applied to a solid material and the relationship between compressing stress σ and axial strain ε is quantified (Pelleg, 2012). This property is called Young's modulus E and is computed as

$$E = \frac{\sigma}{\varepsilon} = \frac{F/A}{dl/l}, \quad (18)$$

423 where F is the applied force, A is the unit area and dl/l is the relative, normalized displacement of the
 424 composite body. It characterizes the compressive properties of a material, i.e. a higher Young's modulus
 425 E describes a stiffer material and a lower E indicates a softer material.

426 During the compression test, an incrementally increasing compressive stress is applied to the composite
 427 body via two rigid objects to opposing sides of the body. Subsequently, at each incremental step, the

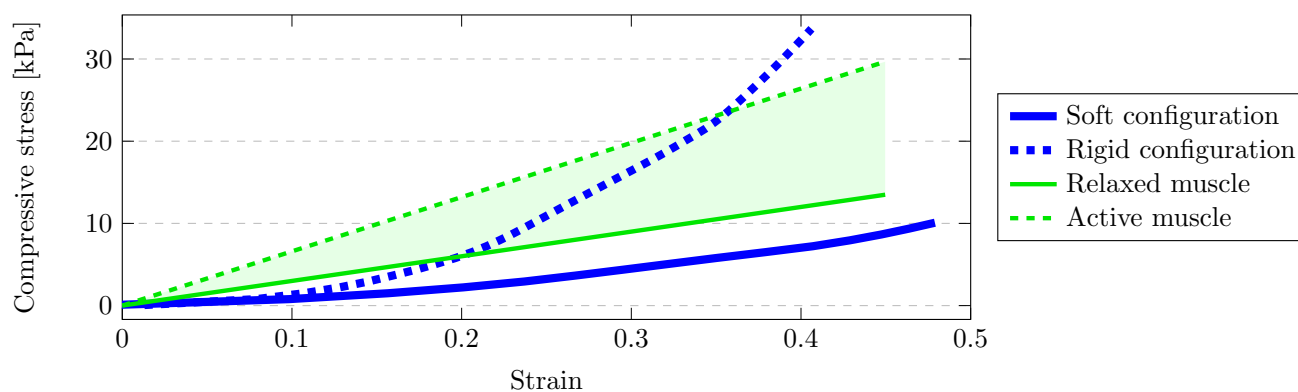


Figure 6. Result of the compression test. The green shaded area depicts the potential range of Young's moduli (Ogneva et al., 2010) determined by relaxed muscles (solid green) and active muscles (dashed green) from experimental data. The range of achievable Young's moduli in the simulation is bound by the soft configuration of the composite object (solid blue) and the rigid configuration (dashed blue).

428 Young's modulus was computed from the strain, i.e. the relative deformation, of the composite body. The
 429 results are compared with experimental data acquired from mammal muscular tissue (Ogneva et al., 2010)
 430 to verify the validity of the simulated soft-tissue. The results of this comparison are shown in Figure 6.
 431 Here, the green lines visualize the experimentally determined Young's moduli for relaxed (solid line) and
 432 contracted (dashed line) muscle fibers (Ogneva et al., 2010) and the green shaded area indicate the resulting
 433 range of potential Young's moduli. Analogously, the blue lines bound the range of achievable Young's
 434 moduli via the simulated composite object. The upper and lower bound are obtained by performing the
 435 above-described compression test for different parameterizations of the composite object. Given that the
 436 simulated, admissible values enclose the experimental data for higher strains, it is possible to approximate
 437 the elastic properties of muscle soft-tissue partially. Note however, that the Young's modulus provided
 438 from the experimental data (Ogneva et al., 2010) constitutes a linear fit and therefore does not exhibit
 439 the typical nonlinear stress–strain relationship which is normally characterized by a region of increasing
 440 modulus (Pons, 2008) as depicted by our simulation in Figure 6. Thus, the slight difference for lower
 441 strain levels can be explained due to approximation error caused by the linear fit in (Ogneva et al., 2010).
 442 Furthermore, the experimental data only considers muscle fibers and is therefore expected to vary from
 443 the considered soft-tissue, e.g., due to additional fat tissue at the attachments. The additional flexibility in
 444 the simulation environment to parameterize lower Young's moduli is thus favorable, since the expected
 445 variation generally leads to softer materials.

446 2.3 Exoskeleton-based Automated Assessment

447 With the nominal and uncertain dynamics model (Section 2.1) and a human-exoskeleton simulation
 448 that includes the key sources of uncertainty (Section 2.2) introduced, the required input samples for the
 449 sensitivity analysis can be generated. Here, the input samples are instantiations of the simulation with
 450 varying parameters for the different sources of uncertainty. Since we investigate how these uncertainties
 451 impact the results of an automated assessment, the output samples are in the form of estimated impedance
 452 parameter. The procedure by which these output samples are generated is explained in this section.

453 In order to perform the spasticity assessment in an automated manner, two components are necessary.
 454 First, a data generation procedure is required during which the robotic system interacts with the human
 455 arm to induce observations from which the impedance parameters can be inferred. Secondly, the captured
 456 data needs to be used to estimate the parameters. In this work, we propose a fully automated scheme for

457 the data generation and estimation that leverages model knowledge to produce the required labels \mathbf{y} . The
 458 complete scheme is illustrated with a block diagram in Figure 7. Here, the real system represents the true,
 459 uncertain human-exoskeleton system which is reproduced in the simulation environment. On the other
 460 hand, the nominal model block describes the idealized dynamics model that can be computed analytically.
 461 The reference trajectory $\theta_d, \dot{\theta}_d$ is depicted in the blue block and is used to observe the joint resistance along
 462 a predefined movement, similar to the passive mobilization that is typically performed by a clinician. It
 463 acts as an input to the PD-controller, which replicates the manual perturbation generated by the clinician
 464 using the exoskeleton.

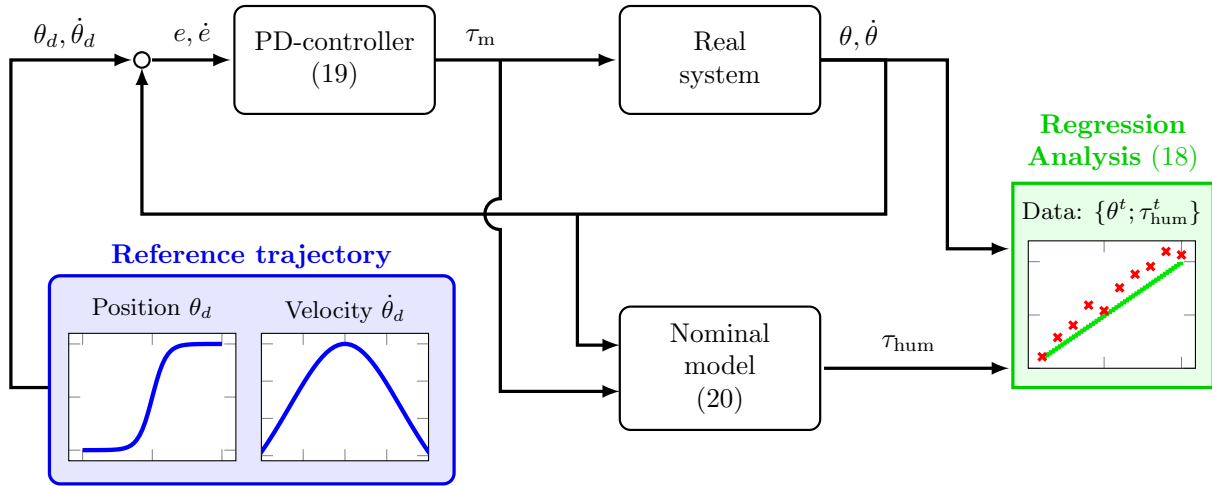


Figure 7. Block diagram of the data collection and estimation scheme for the automated assessment.

465 For the reference trajectory a sigmoid function is selected, since it is known to generate a minimum
 466 jerk profile on the joint level (Flash and Hogan, 1985), thus, leading to a natural and comfortable motion
 467 for the patient. With the reference trajectory being defined, the exoskeleton applies a torque on the
 468 human arm to emulate the manual perturbation performed by the clinician. This is achieved by using the
 469 feedback provided by the exoskeleton measurements $\theta, \dot{\theta}$ and feeding the current tracking error e, \dot{e} into a
 470 PD-controller to compute the required motor torque:

$$\tau_m(e, \dot{e}) = K_p e + K_d \dot{e}, \quad (19)$$

471 where $e = \theta_d - \theta$ and K_p, K_d are the feedback gains of the controller. By applying the motor torque (19),
 472 the human-exoskeleton system is moved and, given sufficiently high control gains, the desired trajectory
 473 θ_d is tracked. For the gains of the exoskeleton PD-controller $K_p = 50$ and $K_d = 15$ is set. In order to
 474 induce spastic behavior in the human simulation, a constant, co-contracting muscle activation of $a = 0.4$
 475 is simulated for the muscles associated with the examined joint. Thereby the human arm will produce
 476 a resisting torque opposing the exoskeleton during a change in joint position. The data that is generated
 477 during the passive mobilization is used for the regression analysis (8).

478 For the data generation according to the nominal model, perfect alignment between the human and
 479 exoskeleton kinematic chain is assumed. Thus, the measured angles $\theta, \dot{\theta}$ are assumed to match the human
 480 joint kinematics $\mathbf{q}, \dot{\mathbf{q}}$, thereby providing the nominal input variables \mathbf{X} for the linear regression (8).
 481 Furthermore, the output vector \mathbf{y} is required, which comprises measurements of the human internal torque
 482 τ_{hum} . Since τ_{hum} is not directly measurable, we exploit the nominal model in Figure 7 to overcome this

483 problem. Specifically, using the known motor torque (19) and the nominal dynamics model (6) we can
 484 compute the nominal human torque τ_{hum} to be

$$\tau_{\text{hum}}(\theta, \dot{\theta}, e, \dot{e}) = \underbrace{M_h(\theta)\ddot{\theta} + C_h(\theta, \dot{\theta}) + g_h(\theta)}_{\tau_{\text{rbd,h}}} + \underbrace{M_e(\theta)\ddot{\theta} + C_e(\theta, \dot{\theta}) + G_e(\theta) - \tau_m(e, \dot{e})}_{\tau_{\text{int,h}}}. \quad (20)$$

485 Here, The parameters of the nominal human model are chosen according to anthropometric
 486 data (Ramachandran et al., 2016) with a nominal reference person of height 1.75m weighting 70kg,
 487 which results in a nominal upper arm length of 33.37cm, a nominal forearm length of 31.9cm, a nominal
 488 upper arm mass of 2.25kg and a nominal forearm mass of 1.31kg. Thus, by measuring the trajectory of the
 489 exoskeleton joint kinematics $\theta, \dot{\theta}$ over time and computing the corresponding nominal human torques τ_{hum}
 490 according to (20), the regression analysis (8) can be performed for each joint independently.

$$\underbrace{\begin{bmatrix} \tau_{\text{hum},i}^1 \\ \tau_{\text{hum},i}^2 \\ \vdots \\ \tau_{\text{hum},i}^T \end{bmatrix}}_y = \underbrace{\begin{bmatrix} \theta_i^1 & \dot{\theta}_i^1 \\ \theta_i^2 & \dot{\theta}_i^2 \\ \vdots & \vdots \\ \theta_i^T & \dot{\theta}_i^T \end{bmatrix}}_{\hat{X}} \underbrace{\begin{bmatrix} k_{h,ii} \\ d_{h,ii} \end{bmatrix}}_{\omega}, \quad (21)$$

491 where, differently to (8), \hat{X} represent the inputs when the exoskeleton kinematic measurements $\theta, \dot{\theta}$ are
 492 used as a placeholder for the human joint kinematics q, \dot{q} . Note that deploying (20) for the computation of
 493 the human torques implicitly allocates torques that are unaccounted for by the nominal dynamics model to
 494 be generated due to spasticity in the patient’s joints. Intuitively, this is analogue to the principle applied
 495 during manual assessment, where the human limb is assumed to be passive and any encountered resistance
 496 is allocated to spasticity. However, as detailed in Section 2.1.2, different sources of uncertainty impact
 497 the human-exoskeleton interaction, which result in interaction torques that are not considered in (20).
 498 Thus, solving (21) will not result in the true viscoelasticity parameter K_h and D_h , due to the impact of
 499 uncertainties on the regression analysis.

500 **2.4 Sensitivity Analysis of Uncertainties**

501 The goal of this section is to quantify the impact of the uncertainties on the estimated impedance
 502 parameters during the exoskeleton-based automated assessment. To this end a sensitivity analysis is
 503 performed to examine how variations in the output of a numerical model or simulations can be ascribed
 504 to variations of its inputs. We consider uncertainties in the modelling of physical human-exoskeleton
 505 interaction as input factors to quantitatively assess their importance. Analogously, do the estimated
 506 viscoelasticity parameters K_h and D_h represent the output samples of the sensitivity analysis. Thereby,
 507 sensitivity is defined as the induced variability in the parameter estimates K_h and D_h due to variability
 508 in the uncertain inputs and is quantified by means of so-called sensitivity indices (Saltelli et al., 2004).
 509 Intuitively, these sensitivity indices represent importance measures, which are allocated to each input
 510 parameter of the simulation, i.e., each source of uncertainty (Pianosi et al., 2016). In this section, the
 511 methods used for the sampling-based sensitivity analysis procedure are presented. First, the input sample
 512 generation is described in Section 2.4.1. Following this, Section 2.4.2 details the deployed methods for the
 513 computation of the sensitivity indices.

514 **2.4.1 Sampling Sources of Uncertainty**

515 For the input sample generation, we draw samples over different parameterization of the human-
 516 exoskeleton simulation. Here, each sampled simulation instance represents a distinct patient with the

517 individual variations present in the population. Six biomechanical parameters are chosen as input factors,
 518 where each parameter is associated with a different source of uncertainty. An overview of the parameters,
 519 their respective uncertainties and the value ranges is depicted in Table 1. Here, kinematic incompatibilities
 520 are produced by varying the length of the human limb. In particular, changes in the upper arm length lead to
 521 macro-misalignments and a resultant CoR offset, since the exoskeleton link length remains unchanged. In
 522 contrast, varying the human forearm length induces micro-misalignments. The second source of uncertainty
 523 investigated during the sensitivity analysis are inaccuracies in the dynamics model. By perturbing the
 524 mass of the upper and forearm, errors in the nominal model are evoked, as the gravitational component
 525 and inertia of the human limb are dependent on the mass. Lastly, uncertainties due to soft-tissue contact
 526 dynamics are considered by sampling over different elasticities of the human upper arm and forearm at the
 527 attachments. The value ranges of the samples shown in Table 1 are derived from statistical information
 528 provided by anthropometric data (Ramachandran et al., 2016). Here, a fixed viscosity of 100Ns/m is chosen
 529 for the micro-elements comprising the soft-tissue to avoid numerical instabilities.

Table 1. Sources of uncertainty and associated simulation parameters for the input sample generation.

Uncertainty	Simulation Parameter	Value range
Kinematic incompatibilities	Length upper arm	27.28cm - 37.78cm
Kinematic incompatibilities	Length forearm	28.27cm - 34.55cm
Inaccuracies in dynamics model	Mass upper arm	0.3kg - 3.41kg
Inaccuracies in dynamics model	Mass forearm	0.1kg - 1.82kg
Soft contact dynamics	Elasticity upper arm	100.5N/m - 974.43N/m
Soft contact dynamics	Elasticity forearm	100.5N/m - 974.43N/m

530 In addition to defining the input variability space, i.e., the value ranges shown in Table 1, further design
 531 choices regarding the sampling strategy have to be made. In general two classes of sampling concepts can
 532 be differentiated, One-At-a-Time (OAT) and All-At-a-Time (AAT) methods (Pianosi et al., 2016). While in
 533 OAT methods variations are induced by perturbing one input parameter only and keeping all other fixed,
 534 AAT methods induce output variations by varying all input parameters concurrently. The main advantage
 535 of OAT in comparison to AAT sampling is the reduced computational load due to fewer samples being
 536 required. However, because of the concurrent sampling in AAT methods, the joint influence of input factors
 537 due to interaction between the parameters can be analyzed, thereby, providing more insights (Pianosi et al.,
 538 2016). Depending on the deployed method to estimate the importance measures, both approaches can be
 539 beneficial. Therefore, the following section presents sensitivity analysis methods with distinct sampling
 540 strategies for different investigation purposes.

541 2.4.2 Sensitivity Analysis Methods

542 Depending on the setting and purpose of the sensitivity analysis, different methods are appropriate. In
 543 Saltelli et al. (2008) two main purposes are introduced. First, the goal of ranking the most relevant input
 544 factors which is called *factor prioritization*. Second, identifying input factors with negligible impact which
 545 is called *factor fixing*. Beyond these two main settings, other purposes are introduced as well. However,
 546 given that the proposed sensitivity analysis is supposed to inform the decision making process in clinical
 547 practice and lead to more robust spasticity assessment, our quantitative analysis is mainly focused on
 548 factor prioritization and factor fixing, since these information lead to a practical guide to performing more

549 robust automated assessment. Additional information may also be derived by qualitative sensitivity analysis
 550 methods, e.g., using scatter plots (Beven, 1993; Kleijnen and Helton, 1999).

551 Furthermore, potential interactions between the investigated sources of uncertainty should also be
 552 considered. Since these interactions may emerge for various parameters and it is a-priori unknown how
 553 the interactions behave with respect to the magnitude of the parameters, we ideally want to perform
 554 a dense sampling over the input variability space. To this end global sensitivity analysis methods are
 555 preferred, which investigate variations over the complete range of admissible inputs. Global sensitivity
 556 analysis methods have previously been shown to facilitate tasks such as supporting efforts in uncertainty
 557 reduction (Hamm et al., 2006) and facilitating robust decision making (Nguyen and de Kok, 2007; Singh
 558 et al., 2014).

559 *Elementary Effects Method*

560 Given these requirements, there are multiple viable sensitivity analysis methods. First, Morris
 561 method (Morris, 1991), also referred to as *elementary effects test*, is an efficient and suitable approach to
 562 perform factor prioritization and fixing. Here, a perturbation-based design is deployed, where the whole
 563 input space is explored by applying perturbations to each input factor separately and computing global
 564 sensitivity measures from the probed samples. This is done by computing so-called elementary effects *EE*
 565 for each input factor x_i

$$EE_i = \frac{f(x_1, \dots, x_{i-1}, x_i + \Delta_i, x_{i+1}, \dots, x_K) - f(x_1, \dots, x_K)}{\Delta_i}, \tag{22}$$

where $\mathbf{x} = (x_1, x_2, \dots, x_K)$ represents a set of input parameters, $f(\mathbf{x})$ denotes the function that maps inputs to model responses, K is the total amount of examined input parameters and Δ_i is the perturbation applied to the i -th input parameter. In order to achieve a global measure of sensitivity, the input space is sampled with r trajectories, each consisting of $K + 1$ sampling points, where each point differs in just one input factor by a fixed amount Δ (Morris, 1991). Thereby, each trajectory allows for the computation of one *EE* per input factor and the sensitivity measures for each parameter can be computed as such:

$$\mu_i = \frac{1}{r} \sum_{j=1}^r EE_i^j \tag{23}$$

$$= \frac{1}{r} \sum_{j=1}^r \frac{f(x_1^j, \dots, x_i^j + \Delta_i^j, \dots, x_K^j) - f(x_1^j, \dots, x_K^j)}{\Delta_i^j}$$

$$\sigma_i^2 = \frac{1}{r - 1} \sum_{j=1}^r (EE_i^j - \mu_i)^2, \tag{24}$$

566 where Δ_i^j represents the perturbation of the i -th input parameter x_i^j in trajectory j and EE_i^j denotes the
 567 computed elementary effect associated with parameter x_i along trajectory j . Here, the mean μ and standard
 568 deviation σ of the elementary effects *EE* are proposed as sensitivity measures (Saltelli et al., 2008). In
 569 particular, μ (23) represents how much the input parameter affects the output, while σ (24) is a measure
 570 for the induced effects due to interaction with other inputs, i.e., how much EE_i varies when changes in
 571 the remaining $i - 1$ parameters occur. Specifically, a small σ_i implies that the effect of parameter x_i on
 572 the output, which is shown by μ_i , is independent of the other parameters. Therefore, Morris method is
 573 particularly well suited for factor fixing, since a simultaneous consideration of both μ and σ allows the

574 identification of negligible input factors, which have both little interaction with the other inputs (small σ)
 575 and do not influence the output strongly (small μ). Moreover, applying this approach requires relatively few
 576 samples, which further increases its utility for factor fixing in cases where model evaluations are expensive.
 577 However, since it is a perturbation-based OAT method, it may lead to erroneous results if the target system
 578 exhibits high-frequencies in its response to variations in the input (Pianosi et al., 2016).

579 *Variance-based Sensitivity Analysis*

580 An alternative approach that facilitates the analysis of output sensitivity with respect to each input
 581 factor over their complete value range are variance-based sensitivity analysis methods, also referred to
 582 as Sobol method (Sobol, 1993). Here, modelling uncertainty is specifically considered by regarding the
 583 input parameters as stochastic variables with a defined probability distribution. Thereby, a conceptual link
 584 between sensitivity and uncertainty is exploited and sensitivity is analyzed by investigating how uncertainty
 585 in the input propagates to the output variables. Subsequently, the relative contribution of each input is
 586 decomposed and used as a measure of sensitivity. To this end variance is used as a measure to quantify
 587 uncertainty. The so-called *first-order effect* S_i , which is a measure for the individual contributions of inputs
 588 to the output variance, is computed as

$$S_i = \frac{\mathbb{V}(z) - \mathbb{E}_{x_i}[\mathbb{V}_{x_{-i}}(z | x_i)]}{\mathbb{V}(z)}, \quad (25)$$

589 where $z = f(\mathbf{x})$ is the output variable, \mathbb{E} denotes the expectation and \mathbb{V} the variance. Here, $\mathbb{V}_{x_{-i}}(z | x_i)$
 590 expresses the conditional variance of the output z over x_{-i} , i.e., all inputs except x_i , given that x_i is
 591 fixed. Analogously, $\mathbb{E}_{x_i}(z | x_i)$ denotes the conditional expected value. Therefore, the second term in (25)
 592 expresses the expected variance in the output given that the i -th input x_i is fixed. A small value for this
 593 expectation, and consequently a high value for S_i , implies that a significant reduction in output variance
 594 can be achieved by fixing x_i (Saltelli et al., 2008). Thus, the first-order index S_i is a measure for the direct
 595 contribution of an input to the output variance, which in turn functions as a place-holder for sensitivity.

596 On the other hand, the *total-order index* S_{Ti} indicates the total effect of an input x_i on the output variance
 597 including interactions with other input factors (Homma and Saltelli, 1996) and is defined as

$$S_{Ti} = \frac{\mathbb{E}_{x_{-i}}[\mathbb{V}_{x_i}(z | x_i)]}{\mathbb{V}(z)}. \quad (26)$$

598 Moreover, variance-based methods allow for the computation of further, higher-order indices, such as
 599 second-order or third-order ones. Thereby, by computing all $2^K - 1$ orders, variance-based sensitivity
 600 measures can theoretically capture the sensitivities present in the system completely. However, since this
 601 is computationally infeasible in practice, a good approximation can be achieved by computing only the
 602 first-order and total-order terms (Saltelli et al., 2004).

603 Thus, variance-based methods are well equipped to analyze sensitivities in a principled manner by both
 604 quantifying the importance of individual inputs and groups of inputs. Moreover, an uncertainty-aware
 605 modelling paradigm is supported and, by sampling the input space using probability distributions, the
 606 full range of input variations can be investigated. However, due to their sampling-intensive nature, it
 607 is impractical to deploy them directly when model evaluations are expensive. Therefore, we propose
 608 to use both the elementary effect test and variance-based sensitivity analysis in conjunction. Thereby,
 609 non-influential input parameters are detected by the efficient elementary effect method and can be discarded
 610 prior to performing a more extensive analysis using variance-based methods.

3 RESULTS

611 In this section we present the findings of performing the proposed two-phase sensitivity analysis scheme.
 612 First, in Section 3.1 the elementary effect test is deployed to screen parameters that do not effect the
 613 automated assessment outcome significantly and can therefore be fixed for subsequent investigations.
 614 Second, the variance-based sensitivity analysis is performed on the remaining input parameter in Section 3.2
 615 to determine the relative importance of the different model uncertainties. Lastly, a qualitative analysis of
 616 the obtained samples is conducted in Section 3.3 to provide further insights. For clarity of presentation the
 617 automated assessment is limited to the estimation of the elbow joint stiffness. The presented sensitivity
 618 analysis is implemented in Matlab using the SAFE toolbox (Pianosi et al., 2015), while the simulation
 619 model is implemented in Python using the MuJoCo physics engine (Todorov et al., 2012).

620 3.1 Factor Fixing using Elementary Effects

621 In order to identify non-influential parameters, we deploy the elementary effect method as described in
 622 Section 2.4.2. To this end, input parameter samples are drawn for which the human-exoskeleton simulation
 623 is instantiated and subsequently the automated assessment is run for each model instance to generate the
 624 respective output samples. Here, we use a radial design for sampling the input parameter hyperspace, since
 625 it has been shown to achieve superior performance for computing elementary effects (Campolongo et al.,
 626 2011). A total of $r = 150$ trajectories is generated for $k = 6$ input parameters, which are listed in Table 1,
 627 resulting in 1050 sampling points. For the generation of the random sampling vectors required in the radial
 628 design, the well-established Latin hypercube approach (McKay et al., 1979; Helton and Davis, 2003) is
 629 used. Moreover, a uniform distribution of the input parameter space is assumed.

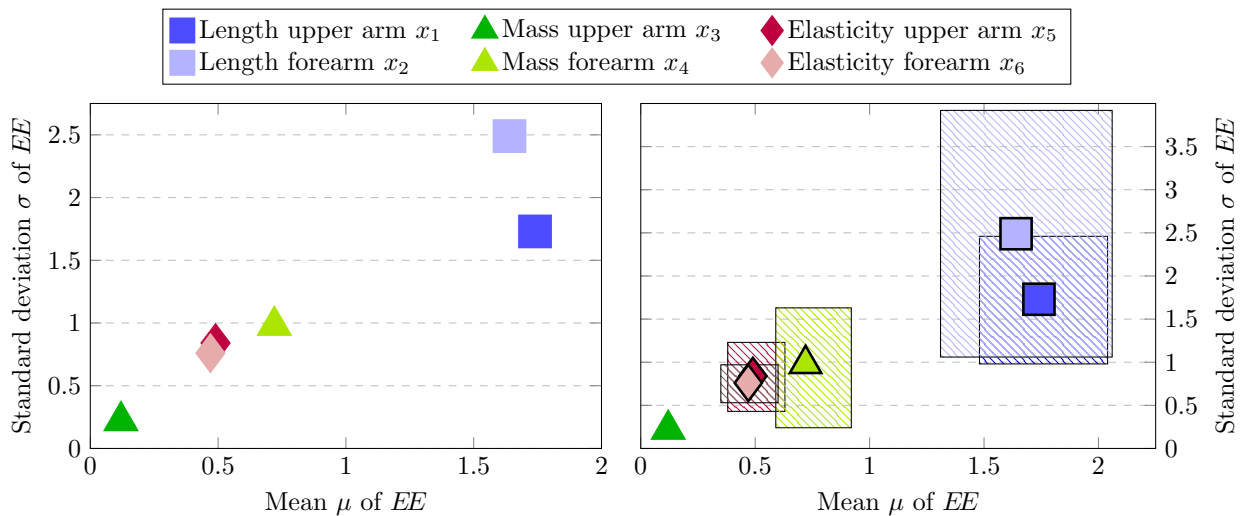


Figure 8. Estimated mean μ versus standard deviation σ of the elementary effects EE (left) and approximated 95% confidence bounds via bootstrapping (right). Here, each input factor is represented by one marker and the confidence bounds are represented by the patterned area associated with each marker.

630 The results of the elementary effect test are depicted in Figure 8. On the left-hand side, it is clearly visible
 631 that the estimated sensitivity measures indicate the mass of the upper arm x_3 as the least influential input
 632 parameter. The low value estimated for both the mean and standard deviation implies that the input factor
 633 has both little direct impact on the estimated joint stiffness during the automated assessment procedure
 634 and moreover does not interact strongly with the remaining parameters. This makes sense intuitively since
 635 the mass of the upper arm is not expected to influence the estimated torque on the elbow level. However,

636 due to the design of the passive mobilization experiment in Section 2.3, it is first necessary to drive the
637 human arm into the desired initial configuration to start the procedure. Thereby, different upper arm mass
638 parameterization could potentially influence the precise starting state, which in turn can lead to slight
639 changes in the estimated stiffness. However, from the results of the elementary effect test it is apparent
640 that these disturbances do not impair the assessment process. Differently, the length of the upper and
641 forearm exhibit the highest sensitivity both with respect to the mean and standard deviations. Therefore,
642 the elementary effect method identifies the parameters associated with uncertainties due to kinematic
643 incompatibilities as the most dominant ones. Lastly, the remaining parameters regarding the soft-tissue
644 contact dynamics and the mass of the forearm are estimated to have a comparable sensitivity measure with
645 the mass having a slightly bigger impact in both μ and σ .

646 Sampling-based sensitivity analysis methods inherently approximate the true sensitivity indices given
647 the observed samples. Therefore, especially when working with small to medium sample sizes, it is
648 pertinent to validate the robustness of the obtained results. In order to investigate this, an additional
649 robustness analysis can be performed (Pianosi et al., 2016), which assesses whether similar sensitivity
650 measures would have been obtained with different input samples. This can be achieved in a sample-efficient
651 manner by approximately computing the confidence bounds of the estimated similarity measures using
652 bootstrapping (Efron and Tibshirani, 1993). Note that while bootstrapping is an efficient technique, the
653 obtained confidence intervals do not constitute theoretically guaranteed bounds in general and can result
654 in overly optimistic estimates when applied to Morris method (Romano and Shaikh, 2012; Yang, 2011).
655 However, applying the method still allows to retrieve valuable insights regarding the estimated sensitivity
656 indices. The results of the robustness analysis are displayed in Figure 8 on the right. Here, a total of 300 μ
657 and σ values are computed for each input factor, where each value is generated by drawing 150 samples
658 with replacement from the original 1050 sampling points. Notably, the confidence bounds for the upper
659 arm mass x_3 are very small, thereby, indicating that the mass of the upper arm can confidently be regarded
660 as a non-influential input factor that can be fixed for subsequent analysis. Differently, the upper arm length
661 x_1 and forearm length x_2 , which are identified as the most important ones by the elementary effect test,
662 are associated with large confidence intervals. In particular the forearm length x_2 features the highest
663 uncertainty in the estimated sensitivity measures. Therefore, the results are not conclusive to make reliable
664 statements beyond the screening of the upper arm mass and the deployment of further sensitivity analysis
665 methods is required.

666 3.2 Factor Prioritization using Variance-based Sensitivity Analysis

667 Following the elementary effect test in the previous evaluation, we perform an additional variance-based
668 sensitivity analysis to obtain a more rigorous understanding of the uncertain sensitivity patterns present in
669 the human-exoskeleton system. To this end we exploit the findings of the prior section to fix the upper arm
670 mass x_3 , as it is identified as a non-influential factor, which leads to a reduction of the computational load
671 of the proposed variance-based analysis. For the input sample generation of the remaining parameter we
672 use the two-phase sample procedure proposed for the variance-based approximation of the first-order and
673 total-order indices (Saltelli et al., 2010). In the first phase, a total of $2N$ random samples is generated, which
674 are referred to as base samples. Subsequently, KN additional input samples are produced by resampling
675 vectors of the base samples. Thereby, this method requires $N(K + 2)$ model evaluation for the estimation
676 of the first-order and total-order effects and is computationally more efficient than a naive approach (Saltelli
677 et al., 2010). Here, we set $N = 3000$ and investigate $K = 5$ input factors leading to a total of 21000
678 simulation runs. The random base samples are again obtained using the Latin hypercube method assuming
679 a uniform distribution over the input parameters.

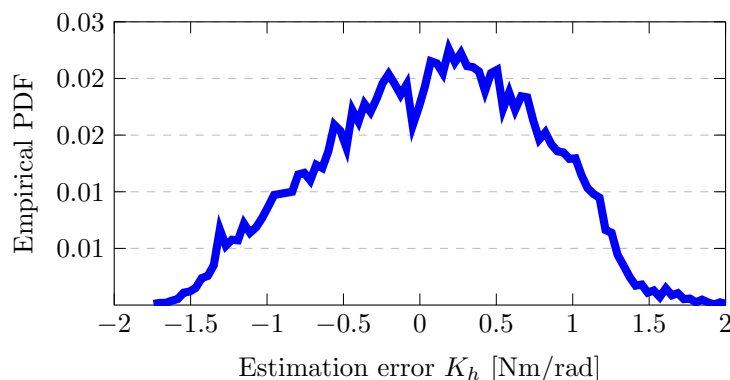


Figure 9. Visualization of the output distribution, i.e., error in the impedance parameter estimation, due to the sampled input parameters. Here, the empirical probability distribution function (PDF) is shown.

680 The resulting output distributions is shown in Figure 9 with the empirical probability distribution function
 681 (PDF), which is approximated from the output samples. Here, the output distribution, i.e., the estimation
 682 error in K_h , resembles a Normal distribution with a mean estimation error slightly larger than 0 Nm/rad.
 683 Thereby, it can be seen how the sampling of uncertainties in the input variability space induces an
 684 output distribution and impacts the assessment results. Note that an implicit assumption in variance-based
 685 sensitivity analysis is that variance is an appropriate measure to capture uncertainty (Pianosi et al., 2016).
 686 Since the empirical PDF in Figure 9 resembles a Normal distribution and is neither multi-model nor
 687 highly-skewed, this assumption holds true, thus strengthening the viability of deploying the approach here.

688 Figure 10 depicts the resulting first-order indices S_i on the left and total-order indices S_{Ti} on the
 689 right. Additionally, the 90% confidence interval are shown by the error bars, which are computed using
 690 bootstrapping. From the first-order effects it is clearly visible that the factors x_1 , x_2 and x_3 are the most
 691 influential ones, with the length of the forearm x_2 having the highest impact. Moreover, the results indicate
 692 that the softness of the upper and forearm x_5 and x_6 are negligible, since their respective total-order indices
 693 are close to zero. Note that a total-order index of value zero constitutes a necessary and sufficient condition
 694 for an input factor to be non-influential (Pianosi et al., 2016). The negative signs for the first-order indices of
 695 x_5 and x_6 can be attributed to numerical errors, which are known to occur for input factors with negligible
 696 sensitivity indices when using the deployed sampling method (Saltelli et al., 2008). Moreover, the sum of
 697 the first-order effects computes to 0.78, while the sum of the total-order effects is 1.13. Since both sums are
 698 not equal to 1, it can be concluded that there are interaction effects present among the input factors in the
 699 system. Additionally, it can be seen in Figure 10 that the total-order indices of each factor are greater than
 700 the respective first-order indices. Thus, it can be inferred that all of the studied input parameter participate
 701 in the interactions.

702 Finally, we perform a convergence analysis to affirm the reliability of the obtained results. Since the
 703 sensitivity indices are approximated from samples, a convergence analysis assesses whether the evaluated
 704 sample size is sufficiently large to make a statement regarding the importance of the input factors. This
 705 can be done efficiently by recomputing the results from increasing sets of sub-samples of the original data
 706 set and analyzing the convergence of the observed indices (Nossent et al., 2011; Pianosi et al., 2016). The
 707 results of the performed convergence analysis are shown in Figure 11. Here it can be seen that both the first
 708 and total-order indices converge quickly when increasing the size of the sub-samples with few changes in
 709 the indices after sub-samples of half the size of the original set. This indicates that a sufficiently large input
 710 sample size is chosen in the evaluation. Since the error bars in Figure 10 are also small when compared

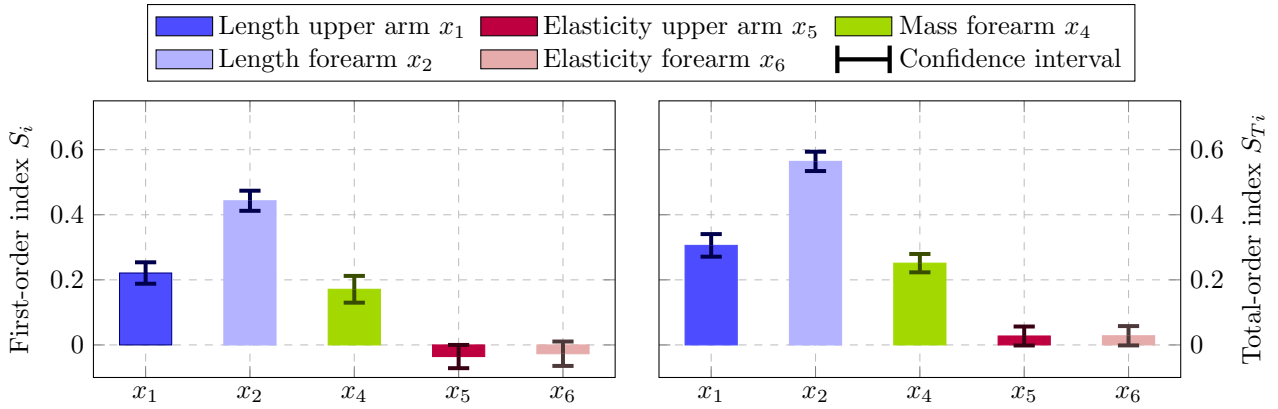


Figure 10. Estimated first-order indices S_i (left) and total-order indices S_{T_i} (right) with 90% confidence intervals using the variance-based sensitivity analysis. The left figure shows the most influential factor is x_2 followed by x_1 and x_4 . The total-order effects on the right identify both x_5 and x_6 to have no impact, since $S_{T_i} = 0$ constitutes a necessary and sufficient condition.

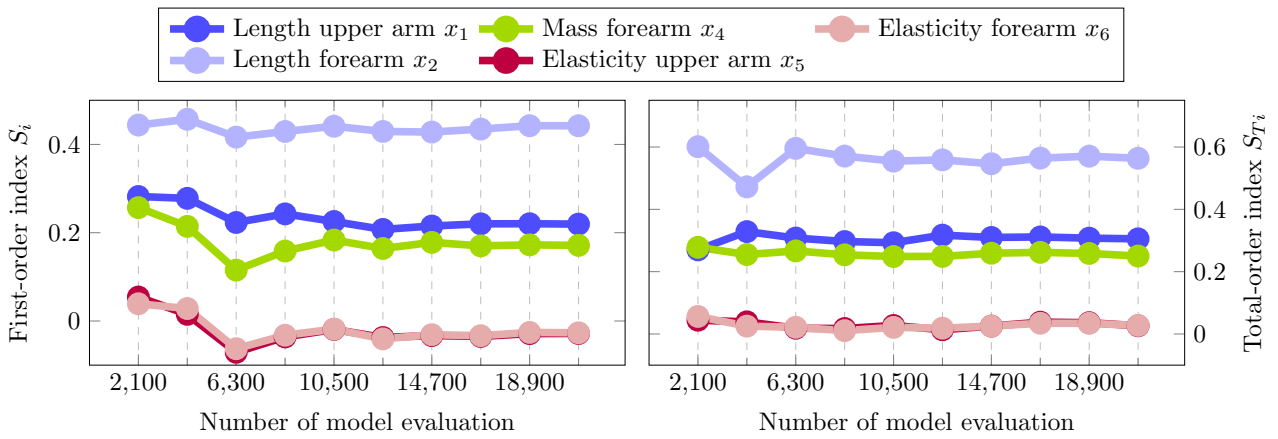


Figure 11. Convergence plot illustrating the estimated sensitivity indices using an increasing amount of sub-samples. Both the first-order and total-order indices converge quickly, which implies that a sufficient sample size is chosen for the variance-based sensitivity analysis.

711 to the estimated indices, the obtained results can be deemed robust. Therefore, we can conclude that the
 712 length of the forearm is the most influential source of uncertainty, with the upper arm length and the mass
 713 of the forearm following as the next most important factors.

714 3.3 Qualitative Sensitivity Analysis

715 In previous sections, we have analyzed the impact of uncertainties on the human-exoskeleton interaction
 716 from a quantitative manner, which is a particularly suitable approach when screening for influential and
 717 non-influential factors and when ranking those. By applying the elementary effect test and variance-
 718 based sensitivity analysis in Section 3.1 and Section 3.2, input parameters associated with kinematic
 719 incompatibilities and erroneous dynamics model are identified as the most relevant uncertainties. However,
 720 little information regarding their functional influence on the system is retrieved and, while interaction
 721 between the inputs is indicated, their exact nature remains unclear. Therefore, we perform an additional
 722 qualitative sensitivity analysis to gain further insights into the most influential sources of uncertainty.

723 Figure 12 visualizes the relationship between input and output samples for x_1 , x_2 and x_4 . Each black
 724 dot in the scatter plot indicates an input-output sample pair, while the larger red dots depict the average

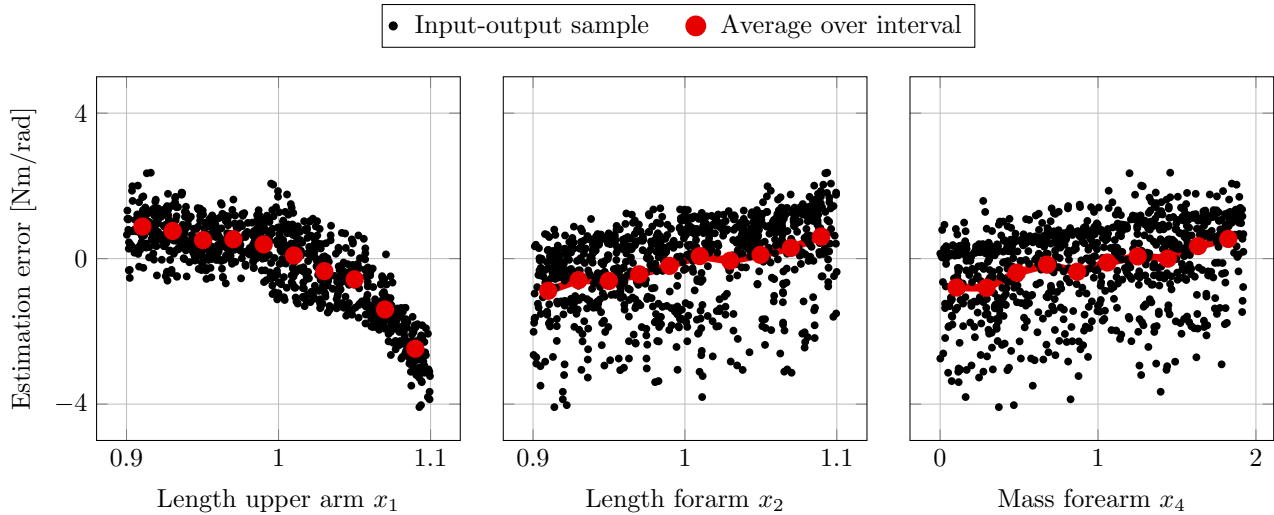


Figure 12. Scatter plot visualizing the output samples against input samples for variations of the upper arm length x_1 (left), variations of the forearm length x_2 (middle) and variations of the forearm mass x_4 (right). The red dots illustrate the mean output for equidistant bins along each input.

725 output values over an interval range of the respective input. Here, equidistant intervals that split the input
 726 value ranges into 10 bins are used, which result in a width of 0.02 for x_1 and x_2 , and 0.17 for x_4 . For the
 727 evaluation, a total of 1500 input samples are generated assuming a uniform distribution for each parameter.
 728 Note that here the x_2 sample range is slightly larger compared to the previous evaluation, since the sampling
 729 strategy of the qualitative sensitivity analysis is more robust to erroneous model responses, which can occur
 730 due to simulation failures caused by unreasonable input parameter combinations. In Figure 12 it is clearly
 731 visible that variation in the length of the upper arm x_1 induce a nonlinear change in the output, while both
 732 forearm length changes x_2 and forearm mass changes x_4 have a linear influence. The linear relationship
 733 in x_2 and x_4 is consistent with the physical intuition for the examined system, since the gravitational
 734 component of the human arm dynamics in (1) is a linear function in the link length and the mass. Thus,
 735 it is indicated that the forearm length x_2 has to be considered as a source of uncertainty with respect
 736 to both kinematic incompatibilities and modelling errors, which leads to a better understanding of the
 737 high sensitivity ranking of x_2 in the variance-based analysis. Differently, the output exhibits a nonlinear
 738 behavior in x_1 with a continuous decrease in the slope for larger upper arm lengths. Thereby, it can be
 739 derived that beyond a certain threshold the misalignment in the center of rotations due to variations in x_1 ,
 740 lead to extreme errors in the output value any may cause catastrophic failures. Thus, despite the relative
 741 lower prioritization in Section 3.2, the upper arm length remains a significant uncertainty and it needs to be
 742 ensured that the mismatch to its nominal values is below certain runaway boundary conditions.

743 Finally, we visualize the interaction between the input parameters using colored scatter plots in Figure 13,
 744 where one input factor is depicted x-axis against another one on the y-axis with the marker color indicating
 745 the output value. Here, the emergence of patterns provides an indication for the interaction between two
 746 factors. From Figure 13 on the far right it can be seen that little interaction is taking place between upper
 747 arm length x_1 and forearm mass x_4 , since the output values do not change significantly with concurrent
 748 changes in the input parameters. However, it can be detected that the upper arm length x_1 is dominant
 749 for very large values, since the markers along the maximal y-axis values are all colored in red. On the
 750 other hand, a slight interaction between the forearm length x_2 and mass x_4 can be inferred from the middle
 751 plot, where the estimation error appears to grow strongly, if both input parameters are increased jointly.

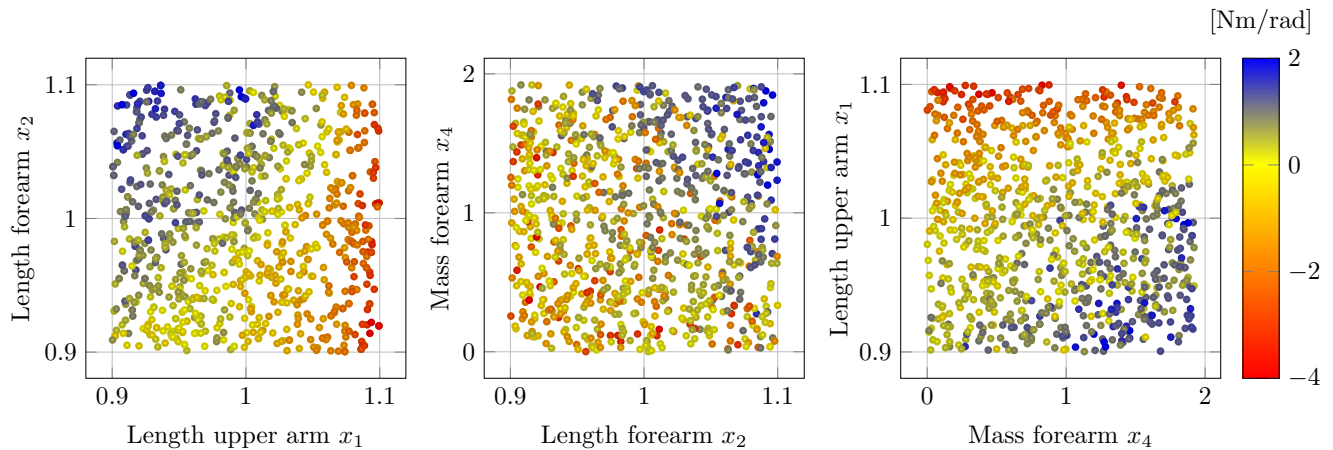


Figure 13. Colored scatter plot depicting samples of the i -th input parameter on the x-axis against the j -th on the y-axis, where the marker color indicates the respective estimation error. By observing emerging pattern in the plot, conclusions regarding the interaction of two input factors can be derived.

752 Intuitively, this can be ascribed to the fact that an increase in the forearm length also shifts the center of
 753 mass of the link, which in turn increases the influence of the forearm mass. Lastly, in Figure 13 on the left
 754 it can clearly be seen that for very high values of x_1 the upper arm length dominates the output, which is
 755 indicated by the red marker coloring along maximal x-axis values.

4 DISCUSSION

756 The present study performed a quantitative sensitivity analysis of the major sources of uncertainty present
 757 in an upper-limb human-exoskeleton system, and their impacts on the arm impedance parameter estimation
 758 was investigated. The performed analysis indicates kinematic incompatibilities and errors in the nominal
 759 dynamics model as the most influential sources of uncertainty. Specifically, variations in the assumed
 760 forearm length belong to both classes of uncertainty and appear to be the most significant factor according
 761 to the results in Figure 10. However, given a wider input variability space, the influence due to variations
 762 in the upper arm length dominates, as shown in the qualitative analysis in Figure 12 and Figure 13. Here,
 763 the results indicate that for slight kinematic misalignments within a 5% range of the nominal upper arm
 764 length, the resulting estimation error only grows approximately linearly. However, when the upper arm
 765 misalignment increases beyond the approximately linear range, the nonlinear functional behavior results in
 766 a blow up of the estimation error. While qualitative sensitivity analysis approaches are more ambiguous, this
 767 finding makes sense intuitively, as the upper arm length is associated with offsets in the center of rotation,
 768 which is typically considered a significant source of uncertainty (Jarrassé and Morel, 2012; Schiele, 2008).
 769 In addition to the above-described link lengths, the mass of the forearm is the third-most relevant source
 770 of uncertainty according to both the elementary effect test and the variance-based sensitivity analysis. Here,
 771 the forearm mass has implications regarding the nominal dynamics model, since it is relevant for both the
 772 gravitational and inertial properties of the human arm. In contrast, the contact dynamics due to soft-tissue
 773 at the attachment are the least relevant as the results in Figure 10 indicate them to be non-influential.

774 Given the results, it can be seen that uncertainty has a significant effect on the exoskeleton-based
 775 arm impedance estimation. In order to help reduce overconfidence in assessment results, the estimation
 776 procedure may benefit from employing uncertainty-aware regression techniques, e.g. Gaussian Processes,
 777 which model uncertainty explicitly, and thus make it transparent for the clinician (Rasmussen and Williams,
 778 2005). Besides modelling the uncertainty, practical steps can be taken to increase the precision of the

779 assessment by exploiting insights provided by our sensitivity analysis. In particular, reducing the effect
780 of kinematic incompatibilities should be prioritized here. More specifically, a close alignment of the center
781 of rotations has to be ensured. Inclusion of passive DoFs on the shoulder as well as the elbow level can
782 mitigate the influence of kinematic incompatibilities (e.g., Vitiello et al., 2013). Additionally, special care
783 should be taken during the donning procedure to ensure an ideal alignment before and during the usage.
784 Second, our sensitivity analysis shows that errors in the nominal dynamics model, due to inaccuracies in
785 the modelling of gravitational and inertial properties of the human arm, adversely affect the impedance
786 estimation result. Therefore, measures should be taken to reduce these effects. This can be achieved
787 by performing more extensive identification procedures for the human arm model instead of relying on
788 standardized models derived from anthropometric data. The benefits of deploying more personalized models
789 has been demonstrated recently in rehabilitation scenarios (Just et al., 2020). While modelling inaccuracies
790 are expected to be less prevalent for the robotic system, they may also adversely affect the assessment. For
791 example in scenarios where unknown and nonlinear friction components influence the robot joints (Chang
792 et al., 2009), the device dynamics may differ from the original identification. Therefore, ensuring the
793 accuracy of the robot model also needs to be considered in practice when performing automated assessment.

794 The simulation environment proposed in the presented study emulates realistic load transmissions between
795 the human and exoskeleton via a mechanical interface composed of supporting cuffs and straps. In addition,
796 we facilitate soft contacts by augmenting the human musculoskeletal model by simulated soft-tissue at
797 the attachment areas. To the best of the authors' knowledge, it is the first upper-limb human-exoskeleton
798 simulation that acknowledges the contact dynamics at the mechanical interface between human and robot
799 by implementing both the interface and the human soft-tissue explicitly. Therefore we believe that the
800 developed high-fidelity simulation platform lends itself well for exploitation in diverse use cases and is
801 particularly suitable to investigate safety and ergonomics in control development. The consideration of
802 ergonomics in physical human-robot interaction is a field that has recently gained growing attention and is
803 considered crucial for driving advances in human-robot collaboration (Gualtieri et al., 2021; Sunesson et al.,
804 2023). Having an explicit implementation of the physical interface is particularly relevant here, in order to
805 accurately represent loads arising at the human limb during interaction with an exoskeleton. Moreover, our
806 proposed simulation platform also provides utility in assisting simulation-based hardware development of
807 wearable robotics, as the consideration of safety and ergonomics is desirable here (Agarwal et al., 2010).

808 While the present study quantitatively analyzed how uncertainties in the human-exoskeleton interaction
809 impact the arm impedance estimation, some simplifying assumptions were made. First, an idealized,
810 fully known robotic system is assumed. Despite the fact that inertial and gravitational components can
811 reasonably be derived for the exoskeleton, commonly, unknown friction dynamics remain. However, we do
812 not expect this to be a significant issue, since a multitude of friction compensation strategies exist (Huang
813 et al., 2019), which can straight-forwardly be applied in the considered scenario. Another assumption
814 was made with respect to the simulation of spastic behavior of the human arm. In particular, we did not
815 consider joint synergies or phase-dependent descriptions of spasticity. Since in this work the focus lied on
816 isolating the influence of uncertainties on the mechanical interaction and consequently on the assessment,
817 the consideration of a more complex spasticity model would provide limited additional benefit to the
818 objective of the study. Still the presented human musculoskeletal simulation allows for the inclusion of
819 different spasticity behaviors in principle. Thus, despite these limitations, the presented results enable us
820 to derive the most relevant sources of uncertainty that impact the physical human-exoskeleton interaction,
821 and thereby help increase the precision of exoskeleton-based arm impedance estimation.

5 CONCLUSION

822 We conclude that this work presents a novel framework to analyze the influence of sources of uncertainty
823 in the human-exoskeleton interaction and their impact on the exoskeleton-based impedance estimation.
824 Due to an increasing demand for robot-based neurorehabilitation and assessment, we argue that the
825 explicit consideration and quantification of uncertainties is paramount, as this allows for more robust and
826 trustworthy estimates. To this end, a human-exoskeleton simulation environment is developed to facilitate
827 the use of sampling-based sensitivity analysis methods. The performed sensitivity analysis indicates that
828 uncertainties significantly impact the impedance estimation, and are primarily caused due to kinematic
829 incompatibilities and inaccuracies in the nominal rigid body dynamics model of the human arm. Therefore,
830 the findings of the study may also be used to increase the precision of exoskeleton-based automated
831 assessment, i.e. by extending model calibrations of the human arm, more careful donning procedures or
832 by deploying uncertainty-aware regression techniques. In the future, we plan to exploit this framework to
833 develop approaches for uncertainty reduction during exoskeleton-based impedance estimation, in order to
834 reduce the estimation uncertainty below pre-defined tolerances. Thus, providing a constructive approach
835 for improving exoskeleton-based automated assessment.

CONFLICT OF INTEREST STATEMENT

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

AUTHOR CONTRIBUTIONS

838 ST and RS implemented the simulation and validated the components with experimental data. ST performed
839 the sensitivity analysis. ST and RS wrote the first draft of the manuscript. SE and SH provided the critical
840 revisions. All authors contributed to the conceptualization and methodology, read and approved the
841 manuscript.

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DATA AVAILABILITY STATEMENT

844 The code for this work can be found online https://github.com/stesfazgi/rehyb_mujoco

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