Assessment of Muscle Fatigue during Isometric Contraction using Autonomic Nervous System Correlates

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Abstract

Muscle fatigue is a complex phenomenon that results in a reduction of the maximal voluntary force. Measuring muscle fatigue can be a challenging task that may involve the use of intramuscular electrodes (i.e., intramuscular electromyography (EMG)) or complex acquisition techniques. In this study, we propose an alternative non-invasive methodology for muscle fatigue detection relying on the analysis of two Autonomous Nervous System (ANS) correlates, i.e., the electrodermal activity (EDA) and heart rate variability (HRV) series. Based on standard surface EMG analysis, we divided 32 healthy subjects performing isometric biceps contraction into two groups: a fatigued group and a non-fatigued group.

EDA signals were analyzed using the recently proposed cvxEDA model in order to derive phasic and tonic components and extract effective features to study ANS dynamics. Furthermore, HRV series were processed to derive several features defined in the time and frequency domains able to estimate the cardiovascular autonomic regulation. A statistical comparison between the fatigued and the non-fatigued groups was performed for each ANS feature, and two EDA features, i.e., the tonic variability and the phasic response rate, showed significant differences. Moreover, a pattern recognition procedure was applied to the combined EDA-HRV feature-set to automatically discern between fatigued and non-fatigued subjects. The proposed SVM classifier, following a recursive feature elimination stage, exhibited a maximal balanced accuracy of 83.33%. Our results demonstrate that muscle fatigue could be identified in a non-invasive fashion through effective EDA and HRV processing.

Keywords: Muscle fatigue, Autonomic nervous system, Heart rate variability, Electrodermal activity, Pattern recognition

1. INTRODUCTION

Fatigue is a complex human condition that reflects several physiological and psychological states [1, 2]. Rather than being generated by a single cause [3], it depends on many central and peripheral phenomena [4] and can affect different sites of the neuromuscular system including the motor cortex, the moto-neuron, the neuromuscular transmission, the excitation-contraction coupling, or the contraction mechanisms [5]. According to the anatomical site (or sites), it is possible to distinguish among central fatigue, fatigue of the neuromuscular junction, or muscle fatigue [5].

Muscle fatigue is defined as a reduction in the ability of a muscle to generate force or power induced by a constant stimulation [6, 4]. Its quantitative measure is challenging, also because the exact moment of muscle exhaustion is unknown [4].

By monitoring the decrease of the maximal force or power produced over time, several indices of muscular fatigue have been proposed in the literature. Particularly, previous studies showed that muscle fatigue influences the surface electromyographic signal (sEMG) in a complex way [7], with a progressive decrease in the mean or median frequencies of the power spectral density function during isometric voluntary sustained contractions [8, 9, 10, 11, 12]. To this end, the analysis in the time-frequency domain of sEMG can be considered as a robust alternative to the gold standard approach of the intramuscular EMG [13, 14, 15]. To monitor fatigue during a physical exercise, especially in ecological scenarios, the wearability and comfort of the measuring system is extremely relevant. However, effective wireless wearable sEMG systems currently in the market are usually expensive and present several limitations regarding comfort, encumbrance, and intrusiveness.

Several studies showed that autonomic nervous system (ANS) dynamics is directly involved in voluntary muscle contractions [16, 17]. In detail, during such contractions both the sympathetic and parasympathetic branches of the ANS are activated through the brainstem activity [18, 16, 19], as well as, through chemo-, mechano- and baroreceptor afferents [20, 21]. The effects of ANS modulation during fatigue are evident when referring to cardiovascular dynamics, as heart rate, respiration rate, and blood pressure significantly changes during intense exercise [22, 23, 24, 23, 25]. Specifically, increased heart rate (HR), as well as oxygen uptake and EMG activity were observed during prolonged exercises [26, 27, 28, 29, 55, 31]. In this context, Heart rate variability (HRV) series, being modulated by sympathetic and parasympathetic interplay, was also investigated [32, 33, 34], with the hypothesis of a sympathetic withdrawal or a vagal tone restoration occurring after maximal exercise [35]. Due to methodological limitations, previous investigations observed significant changes in HRV spectra after
fatiguing training sessions [36, 37, 33, 38, 39], whereas few studies monitored heartbeat dynamics during the fatiguing task [40, 41, 42].

To our knowledge, other markers of ANS dynamics, such as electrodermal activity (EDA) [43, 44, 45, 46], has not been investigated in the frame of muscle fatigue induced by physical exercise. EDA is a widely used physiological signal modulated by attentional, affective, and motivational cognitive processes. EDA is directly controlled by the sympathetic branch of the ANS [47] and is used as an effective biomarker of emotional responsiveness, stress, and mental fatigue [48, 49, 50, 47].

Therefore, we hypothesize that combined HRV and EDA analyses can be profitably used to improve on the state of the art automatic recognition of the muscle fatigue condition from ANS. Contrary to the intramuscular EMG, both HRV and EDA are physiological series that can be monitored through non-invasive sensors (i.e., surface electrodes).

Accordingly, here we propose to assess muscle fatigue only using ANS measurements acquired from wearable devices, possibly opening novel applications for unobtrusive and ecological muscular fatigue monitoring. Specifically, we investigate whether parameters extracted from EDA and HRV during isometric exercise are able to statistically discriminate between a group of fatigued subjects and a control group (non-fatigue). Moreover, a pattern recognition system is proposed to automatically discern data from the two groups at a single-subject level.

2. METHODS

2.1. Subject Recruitment and Experimental protocol

Thirty-two young right-handed adults (mean age 29.25 ± 3.38 (SD) - 16 males) participated in the study. Before performing the experiment, all subjects signed an informed consent to take part in the study. All experimental procedures were approved by the local ethical committee. None of the participants had a history of neurological disorders, musculoskeletal injuries, or cardiovascular diseases, or assumed antidepressants that could have affected the ANS functioning and hence the experimental outcomes.

Subjects were required to perform two isometric force production tasks with their right arm: a maximum voluntary isometric contraction (MVC) and a submaximal fatiguing contraction. More in detail, the experimental paradigm consisted of three stages:

- each subject sat on a comfortable chair, with the elbow completely extended along the body, and a strap fixed on the wrist and connected to a load cell (i.e., dynamometer) [55]. To measure the MVC, the elbow flexion range of motion had to be performed completely, starting from a full extension to avoid/minimize possible compensation by the shoulders and trunk. We instructed subjects to increase the isometric force exerted by the right arm, from baseline to their maximum, and to maintain this maximum for 5s. All subjects were verbally encouraged to maximize their force production while their MVC was measured by the dynamometer [56, 3]. Subjects performed three MVC trials and the average of the three measures was used as the reference value to compute the target force for the submaximal fatiguing contraction. Subjects were given a rest period of 2 min after each MVC trial to minimize fatigue and ensure the correct exertion of the maximum force at each trial.

- After the MVC measurement phase, we acquired EMG, EDA and ECG data as it follows: the EMG activity measured over the biceps brachii (short and long head), upper trapezius, deltoideus, triceps brachii, brachioradialis, flexor carpi radialis, and extensor digitorum communis was recorded using surface electrodes. Surface sensors were placed in single differential mode with two contact electrodes of 10.0 x 1.0 mm placed in parallel with an interelectrode distance of 10.0 mm. This is considered the optimum interelectrode spacing for reducing crosstalk in sEMG signals that contain baseline noise [57]. The two contact electrodes are assembled in a case of 41 x 20 x 5 mm. This kind of electrode is incorporated in the commercial equipment 16-bit A/D Bagnoli Delsys EMG Systems. Following the SENIAM guidelines, the sensors were placed along the longitudinal midline of each muscle (with the long side parallel to the muscle fibers) to minimize crosstalk [58]. To determine the precise location, origin, insertion, and function of the muscle under study as well as any nearby muscles that may produce undesirable signals, the placement procedure was preformed by a clinical expert in anatomy. In addition, to ensure that high-quality signals are recorded, once the sensors location was determined, subjects were asked to contract the muscle. Then, the sensor location was being slightly shifted until the detected signal was maximized. In this study, we focused only on the short-head biceps brachii (which assists with addition), since the experimental task (equivalent to a biceps curl) allowed to maximize biceps contraction. Concerning ECG and EDA sensors, standard disposable Ag/AgCl electrodes were used. ECG was recorded from a modified lead II configuration, which places the right electrode below the sternum and the left one on the left side of the torso below the ribcage. As suggested in [59], EDA electrodes were placed over the solar surface of the distal phalanges of the index and middle fingers of the left hand (non-dominant). A 24-bit A/D Biopac MP35 was used to acquire ECG and EDA signals. After the electrode montage, subjects were invited to comfortably sit with eyes open and their upper limbs along the body in a silent room for a 5 min resting state, while the EDA, ECG and sEMG were recorded with a sampling frequency of 2000 Hz.

- Finally, each subject performed a submaximal fatiguing task with the elbow flexed at 90 degree, the arm along the body (see Figure 1), and a strap fixed on the right wrist and connected to a load equal to 40% of their MVC. Then, subjects were asked to maintain it as long as they could.

To avoid confounding factors possibly affecting the recorded signals, we controlled for temperature and humidity level inside
the room. Throughout the experiment, subjects comfortably sat on a chair and were instructed to avoid speaking and strong changes in the respiration activity (such as breath hold or hyperventilation). Moreover, none of the participants were taking antidepressants or medications known to affect the ANS functioning or have had a history of neurological disorders, musculoskeletal injuries, or cardiovascular diseases. Before and after the experiment, each subject was asked to fill out psychometric tests to assess their stress level (State-Trait Anxiety Inventory test [51]), emotional state (Positive Affect Schedule [52]) and physical fatigue level through ad-hoc questions selected from the Multidimensional Fatigue Inventory and Fatigue Assessment Scale tests [53]. The two groups of subjects did not show any statistical significant differences between the scores of all psychometric tests.

2.2. sEMG Analysis

Surface EMG analysis was used as reference analysis to distinguish fatigued subjects from non-fatigued ones [5]. The EMG signals of the biceps and other monitored muscles were filtered with a Infinite Impulse Response (IIR) comb notch filter (50 Hz). Then a zero-phase Butterworth IIR band-pass filter was used to retain frequencies between 30 and 500 Hz. EMG spectral analysis was performed using the Discrete Fast Fourier Transform estimating the power spectra in a 1s moving Hamming window. In fact, it was proven that in case of isometric, constant force, fatiguating contractions, the EMG signal might be considered stationary for epoch durations of about 1–2 seconds [5]. There was no overlap between consecutive time windows, according to previous findings [60]. For each 1-s time window, we calculated the median frequency value to obtain the trend of the Median Frequency (MNF) along the time. Subjects who showed a decrease in MNF over time were considered as fatigued [5, 60].

2.3. EDA Processing using cvxEDA algorithm

EDA refers to changes in the electrical properties of the skin that are directly related to the eccrine sweat gland activity. The EDA is measured by monitoring skin conductance (SC) changes over specific body sites where the concentration of the eccrine glands is high. In this study the EDA was acquired on the finger phalanxes of the non-dominant hand [47]. Due to the direct control of the sweat glands activity performed by the sympathetic branch of the ANS (and in particular the sudomotor nerve), the EDA analysis is considered one of the best ways to monitor the sudomotor nerve activity (SMNA) and, consequently, ANS activity.

It is possible to decompose the EDA signals into two components, phasic and tonic, which differ for their time scales and their relationships with the external stimuli [61, 47]. The tonic component represents the conductance level (SCL) of the EDA signal, i.e., the slow-varying baseline level, whereas the phasic component refers to faster variations directly related to an exogenous stimulus. A single response to a stimulus is generally called skin conductance response (SCR).

The whole EDA signal results from the sum of these components, that provide relevant and non-redundant information about ANS dynamics. We extracted this information by decomposing the EDA signal through an previously validated approach: the cvxEDA model [61] (available online at: http://www.mathworks.com/matlabcentral/fileexchange/53326-cvxeda).

Here we briefly summarize the main methodological steps behind the cvxEDA model for EDA dynamics (details can be found in [61]). The model assumes EDA as the output of a linear time-invariant system to a sparse non-negative driver signal. CvxEDA describes the EDA signal \( y \) as a sum of three terms: a phasic component \( t \), a smooth tonic component \( s \), and an additive independent and identically distributed zero-average Gaussian noise term \( \epsilon \); 

\[
y = r + t + \epsilon.
\]

The tonic component is modeled by the sum of an offset term, a linear trend, and a smoothed cubic spline that are in accordance with physiological properties: 

\[
t = B t + C d.
\]

Where \( B \) is a matrix of cubic B-spline basis functions, \( t \) is the vector of spline coefficients, \( C \) is a \( N \times 2 \) matrix with \( C_{i,1} = 1 \) and \( C_{i,2} = i/N \) (with \( N \) the length of the EDA signal), and \( d \) is a \( 2 \times 1 \) vector containing the offset and slope coefficients for the linear trend.

The phasic component results from the convolution between a signal that represents the SMNA, \( p \), and an impulse response shaped like a biexponential Bateman function [63]:

\[
h(\tau) = (e^{-\frac{\tau}{\tau_1}} - e^{-\frac{\tau}{\tau_2}}) u(\tau),
\]

with \( \tau_1 = 2, \tau_0 = 0.7 \). This function results form a bi-compartmental pharmacokinetic model representing the diffusion of the sweat through the gland ducts, and can be represented, after its discrete-time approximation, in the matrix form of an autoregressive moving average model \( A^{-1} M \). Where: \( M \) is a tridiagonal matrix with elements \( M_{i,j} = M_{i,j-2} = 1, M_{i,j-1} = 2, 3 \leq i \leq N \); and \( A \) is a tridiagonal matrix with elements \( A_{i,j} = (\tau_1^{-1} \delta + 2)(\tau_0^{-1} \delta + 2)/(\tau_1^{-1} \delta^{2} - \tau_0^{-1} \delta^{2}), A_{i,j-1} = (2 \tau_1^{-1} \tau_0^{-1} \delta^2 - 8)/(\tau_1^{-1} \delta^2 - \tau_0^{-1} \delta^2), A_{i,j-2} = (\tau_1^{-1} \delta^2 - 2(\tau_1^{-1} \delta - 2)/(\tau_1^{-1} \delta^2 - \tau_0^{-1} \delta^2)), \) with sampling time interval \( \delta \).

Using an auxiliary variable \( q \) such that \( q = A^{-1} p \) and \( r = M q \), the final observation model results as:

\[
y = Mq + B t + C d + \epsilon.
\]

Given the EDA model (2), we aim at identifying the tonic component \( t \) and the maximum a posteriori (MAP) neural driver SMNA (\( p \), parametrized by \( [q, t, d] \), for the observed EDA signal \( y \). CvxEDA rewrites the MAP problem as a constrained minimization QP convex problem:
minimize $\frac{1}{2} \| M q + B \ell + C d - y \|_2^2 + \alpha \| A q \|_1 + \frac{\gamma}{2} \| f \|_2^2$  
subject to $A q \geq 0$.  

(3)

Where $\alpha$ and $\gamma$ are parameters that depend on specific prior distributions (see [61]) and control the strength of the penalty (i.e., sparsity and smoothness) for the phasic and tonic components, respectively. CvxEDA finds the optimal $[q, \ell, d]$ and consequently the optimal tonic and phasic components that include the prior knowledge about the spiking sparse nature and nonnegativity of the neural signal representing the SMNA $(p)$ and the smoothness of the tonic SCL.

Table 1: List of features extracted from phasic and tonic EDA.

<table>
<thead>
<tr>
<th>Feature</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>SCRfreq</td>
<td>frequency of significant SCR wrw (i.e., exceeded the threshold of .5 uS [47, 62])</td>
</tr>
<tr>
<td>AUCphasic</td>
<td>Area under the curve of phasic component wrw ($\mu$S s)</td>
</tr>
<tr>
<td>MaxPeak</td>
<td>maximum amplitude of significant peaks of SMNA signal wrw ($\mu$S)</td>
</tr>
<tr>
<td>MeanAMP</td>
<td>Mean value of the SMNA signal wrw ($\mu$S)</td>
</tr>
<tr>
<td>STDphasic</td>
<td>Standard deviation of SMNA signal wrw ($\mu$S)</td>
</tr>
<tr>
<td>MeanTonic</td>
<td>Mean value of the tonic component wrw ($\mu$S)</td>
</tr>
<tr>
<td>EDAsym</td>
<td>Index of sympathetic nervous system activity ($\mu$S$^2$). PSD of the EDA signal wrw= within response window (i.e., 5 secs before the end of the task)</td>
</tr>
</tbody>
</table>

We then extracted several features from both the tonic and phasic signal as summarized in Table 1. In addition to the standard statistical features in the time domain, we also computed the power spectral density (PSD) of the EDA signal in the frequency band 0.045-0.25 Hz (i.e., EDAsym), which has been demonstrated to be a reliable parameter for the assessment of the sympathetic nervous system activity [63].

2.4. ECG processing

ECG signals were band-pass filtered (0.50-40 Hz) to reduce noise and motion artifacts. The QRS-waveforms and each relative R-peaks were detected using the Pan-Tompkins method [64]. All time series were visually inspected to ensure that no beat was discarded. After accounting for the missed R-wave beats, the R-R interval time series (RR) were computed. Moreover, the non-evenly sampled RR interval series were interpolated (4 Hz) by means of a cubic spline functions, i.e. the HRV time series. A set of features in the time and frequency domain was estimated from every RR and HRV series. More in detail, we computed the mean (meanRR) and the standard deviation (stdRR) of the RR intervals within a time window, the square root of the mean squared differences of successive RR intervals (RMSSD), the number of interval differences of successive RR intervals greater than 50 ms divided by the total number of RR intervals (pNN50).

In the frequency domain, we defined some features for the assessment of sympathetic and parasympathetic activities. More specifically, we estimated the power spectral density of HRV signals applying the Welch’s periodogram method with Blackman window and 50% of overlapped segments. The fast Fourier transform was computed for each windowed segment and the power spectra of the segments were averaged. We then calculated the power within the two main spectral bands: the Low Frequency band (LF, from 0.04Hz to 0.15Hz), and the High Frequency band (HF, from 0.15Hz to 0.4Hz). The HF components of HRV are known to be influenced by the parasympathetic system. In contrast, the low-frequency components (LF, 0.045-0.15 Hz) are influenced by both the sympathetic and parasympathetic nervous systems. Moreover, we calculated also the ratio between the LF and the HF power was computed in order to derive information about the sympatho-vagal balance (although this has not been fully accepted as an accurate measure of the ANS balance since the LF band also contains parasympathetic dynamics) [66].

In addition, also some geometrical indexes were extracted, such as the HRV triangular index (HRVtri), and the triangular interpolation of RR interval histogram (TINN). HRVtri was computed following the standard 1/128 seconds bin width, as recommended in [65].

All HRV features are listed with their relative brief description in Table 2.

Table 2: List of features extracted from HRV signals.

<table>
<thead>
<tr>
<th>Feature</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>meanRR</td>
<td>mean value of RR intervals [s]</td>
</tr>
<tr>
<td>stdRR</td>
<td>standard deviation of RR intervals [s]</td>
</tr>
<tr>
<td>pNN50</td>
<td>Number of successive RR interval pairs &gt;50 ms divided by the total number of RR intervals [%]</td>
</tr>
<tr>
<td>RMSSD</td>
<td>Square root of the mean squared differences between successive RR intervals [s]</td>
</tr>
<tr>
<td>LF</td>
<td>Absolute powers of LF band [$\mu$S$^2$]</td>
</tr>
<tr>
<td>LF/nu</td>
<td>LF/(LF+HF)</td>
</tr>
<tr>
<td>HF</td>
<td>Absolute powers of HF band [$\mu$S$^2$]</td>
</tr>
<tr>
<td>HF/nu</td>
<td>HF/(HF+HF)</td>
</tr>
<tr>
<td>LF/HF</td>
<td>Mean value of the tonic component wrw ($\mu$S)</td>
</tr>
<tr>
<td>TINN</td>
<td>The integral of the RR interval histogram divided by the height of the histogram</td>
</tr>
</tbody>
</table>

2.5. Statistical Analysis and Classification Procedure

In order to detect whether subjects were fatigued or not, according to a literature review (e.g. [5, 67] and references therein), we divided the isometric exercise session into two sub-sessions: the first 25% of trial duration; the last 25% of trial duration. The EMG-MNFs computed on the first and last quarter of the experiment were statistically compared, and a subject was classified as fatigued if a significant decrease were found (see two examples of a fatigued and a non-fatigued subjects in Figure 2). According to the results of this statistical analysis, we divided the subject-set into two groups: fatigued (F) subjects and non-fatigued (N-F) subjects.

For each subject, we computed the aforementioned features within the first and the last 25% of EDA signal recorded during
the isometric exercise, and considered the difference between these two values. Instead, given the frequency range of the HRV (<0.4Hz), we could not segment the HRV signal within the first and the last 25% of the session (time windows less than 30s). Therefore, in this case the features were calculated within the whole experimental session and the obtained value were subtracted by the one calculated during the resting phase.

2.5.1. Statistical Analysis

For each feature, we compared values between fatigued and non-fatigued conditions using the non-parametric Mann-Whitney U test under the null hypothesis that the medians of the two groups were equal. The choice of a non-parametric test was justified by the fact that most of the features were not normally distributed as demonstrated by the Shapiro-Wilk test ($p < 0.05$). All p-values were corrected following the Holm-Bonferroni’s method.

2.5.2. Classification Procedure

We applied a pattern recognition algorithm to distinguish between the two groups of subjects, only using features of ANS dynamics. The algorithm used a support vector machine classifier, which implemented a recursive feature elimination (SVM-RFE) procedure [68, 69, 70]. The algorithm was validated through a leave-one-subject-out procedure (LOSO) (i.e. leave-one-out cross validation), which is a nearly unbiased estimator of the out sample error [71, 50, 72].

In detail, within the LOSO scheme, considering $N$ subjects, iteratively we split the feature-set in a training set, comprising data from $(N \text{−} 1)$ participants, and a test set comprising data from the remaining subject. Moreover, for each training set, a recursive feature elimination (RFE) algorithm is applied. We specifically chose a recently developed nonlinear SVM-RFE that employs a radial basis function kernel and includes a correlation bias reduction strategy [68]. At each iteration of the RFE, the SVM model is trained. The feature with the smallest ranking criterion is removed since it has the least effect on classification [68]. The remaining features are kept for the SVM model used at the next iteration. This process is repeated until all the features have been removed. Then the features are sorted according to their removal order. The later a feature is removed, the more important it is. We computed the median rank for each feature over all folds. Results of the sub-feature-set that achieved the most accurate classification will be shown in the following section. The diagonal of the $2 \times 2$ matrix represents the percentage subjects correctly classified for each of the two classes (true positives and true negatives). The later a feature is removed, the more important it is. In the anti-diagonal, we showed the percentage of the fatigued subjects that were incorrectly marked as non-fatigued and vice versa.

3. RESULTS

The statistical analysis on the EMG-MNF time series of the biceps demonstrated that 12 subjects experienced fatigue of this muscle. In fact, for these subjects a significantly decreasing trend on EMG-MNF was shown between the first and last 25% of the exercise duration. The remaining 20 subjects did not show fatigue for the biceps brachii and were labeled as non-fatigued. Same evaluation procedures, although limited to the other muscles considered in this study, showed non-fatigued conditions. In addition, a visual inspection analysis was performed to confirm a proper separation of the two groups according to the MNF trend. Figure 2 shows two MNF trends over time from a fatigued subject (lower) compared to a non-fatigued subject (upper). A decreasing MDF trend, indicating fatigue condition, is reported in red, while in blue it is shown a MDF trend related to a subject who, based on our analysis, did not experience the fatigue condition.

![Figure 2: Example of MDF trends. In red it is reported a decreasing MDF trend related to a subject belonging to the fatigue group (F). Differently, in blue it is reported an example of a subject who did not experience the fatigue condition (N-F).](image)

We then compared EDA and HRV features between the two groups. Concerning EDA, results are depicted in Fig. 3. Most of these features did not show significant differences. However, the standard deviation of the tonic component and the frequency of the phasic responses showed significant differences between groups.

![Figure 3: Comparison of EDA features between fatigued (F) and non-fatigued (N-F) groups.](image)

No group-wise statistical differences were found in HRV features (see Fig. 4).

Results for the classification procedure gathered at each iteration of the RFE procedure are depicted in Figure 5. Moreover, the confusion matrix related to the best feature combination is shown in Table 3. The SVM-RFE showed that the most informative feature-subset comprised features from both the EDA and HRV series, specifically using markers of parasympathetic activity (HFnu, HF, RMSSD) and electrodermal sympathetic activity (AUCphasic, SCRfreq, MaxPeak). Of note, such a pattern recognition system was able to classify the fatigued and non-fatigued subjects with a balanced accuracy of 83.33%.
We investigated ANS correlates of muscle fatigue through combined HRV and EDA analyses. To this end, 32 healthy subjects (16 males) were monitored during a prolonged isometric contraction. The proposed experimental task naturally induced the activation of multiple muscles of the upper limb. However, these conditions mostly led to a major involvement of the biceps brachii. By limiting our study to the monitoring of superficial electromyographic activity of the main upper limb muscles, we labeled participants as fatigued and non-fatigued according to the sEMG MNF of the biceps brachii (short and long head), upper trapezius, deltoideus, triceps brachii, brachioradialis, flexor carpi radialis, and extensor digitorum communis. To our knowledge, this is the first study proposing an automatic muscle fatigue recognition at a single-subject level based on electrodermal and cardiovascular dynamics. Indeed, our results demonstrate that it is possible to recognize muscle fatigue condition of biceps using a proper combination of EDA phasic features and linear/spectral quantifiers of cardiovascular variability.

From a methodological viewpoint, we employed a recently proposed model of EDA in order to efficiently decompose the signal into its phasic and tonic components. This processing represents an essential step to effectively derive psycho-physiological markers [61, 74]. In addition, markers of sympathovagal control were derived from HRV analysis performed...
in the time and frequency domains. In this way, we collected features from two of the most widely used ANS correlates, carrying complementary information.

As a preliminary, exploratory step, we performed a statistical analysis to compare ANS features between the first and last quarter of the fatiguing exercise. Features from cardiovascular variability did not show significant changes. Of note, the inability to calculate the HRV features within a smaller time window, during the exercise, may have limited our possibility to detect possible changes of the sympathovagal dynamics. On the other hand, EDA measurements revealed a significant decrease of the sudomotor nerve spike frequency, as well as an increase of the tonic variability in the case of fatigue. Previous studies report that the vagal activity is predominant during the first part of an isometric exercise, whereas the sympathetic effect prevails close to the completion of the exercise [56]. As EDA neural control is exclusively sympathetic [47, 74], we can assume that the sympathetic activity, when muscle fatigue occurs, is reflected in a more variable tonic signal (slow variations) together with a decrease of faster phasic variations.

We have also proposed a pattern recognition approach able to automatically recognize muscle fatigue at a single-subject level using only ANS correlates. In addition we demonstrated that a proper combination of HRV and EDA features significantly improves balanced classification accuracy. The automatic muscle fatigue classification was performed using nu-SVM, which was validated by means of a LOSO cross-validation. Results showed a satisfactory balanced accuracy of 83.33% when a specific feature combination was taken into account (see Table 3). This accuracy results from the combination of a specificity of 91.67%, and a sensitivity of 75.00%. Our results demonstrate that the discrimination of ANS states associated with muscle fatigue can be supported by a multivariate machine learning analysis implemented at a single-subject level, rather than using group-wise statistics. Of note, the accuracy of the non-fatigued class, i.e., 75.00%, was fairly lower than the fatigue class. This could be explained by the conservative technique used to label the subjects according to a statistical analysis on sEMG spectral features. More in detail, four out of the five subjects who were misclassified in the non-fatigued group (i.e., the 25% of false positives) showed a decrease in the MNF trend from the first to the last quarter of the task. However, paired statistical analyses on such trends did not show significant differences, possibly due to the high inter-subject variability. At a speculation level, this may suggest possible muscle fatigue also in subjects belonging to the non-fatigued group. In particular, misclassified subjects may be the most fatigued ones within the non-fatigued group. Furthermore, a rigorous SVM-RFE procedure revealed that the most informative features are strongly correlated with the parasympathetic branch of cardiovascular neural control and the sympathetic activation of the electrodermal processes. Of note, the implemented SVM-RFE uses a radial basis function kernel and reduces bias due to highly correlated features. Our results also suggest that the muscle fatigue process is modulated by both the ANS branches. In this sense, EDA, which is controlled only by the sympathetic nervous system, and the HF spectral information of the HRV signal, which is strongly associated with the parasympathetic activity, have been confirmed to be an ideal way to monitor the ANS dynamics.

To summarize, this work proposes a non-invasive methodology for muscular fatigue detection. However, we are aware the non-invasiveness and simplicity of our approach comes with limitations. First, sEMGs cannot consider the effect of deeper muscles that may contribute to the task and got fatigued, and sometimes it could possibly lead to controversial outcomes, e.g., the compensation of MNF decrease in submaximal task due to the recruitment of new motor units. These issues could be addressed relying on invasive procedures or through EMG array-based acquisition and processing techniques, which can in principle identify the discharge of multi-channel action potentials by individual muscle units [73]. This would certainly improve the reliability of the results but, on the other side, it would be in contrast with the minimalistic approach that drove our work, under a sensing and/or computational point of view. A comparison between these different approaches is interesting, and it will deserve future analyses to identify the right trade-off between effectiveness and usability/simplicity.

These results pave the path towards unobtrusive and wearable devices to assess fatigue in working environments or during rehabilitative procedures in an ecological fashion. Indeed, the devices that are able to monitor, for example, HRV series are already very popular in the everyday life (e.g., smartphones and smartwatches). In addition, ANS measurements could be also used to infer the emotional state of the subjects [74, 75, 46], which can play a crucial role in the rehabilitation procedures [76].

Future endeavors will be directed to study EMG dynamics from several muscles locations, and to implement our approach into a portable device, which integrates, e.g., into a smartwatch, HRV and EDA sensors (i.e., Empatica wearable systems [77]).

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