

# Template matching and Matrix Profile for Signal Quality Assessment of Carotid and Femoral Laser Doppler Vibrometer Signals

Silvia Seoni<sup>1+</sup>, Simeon Beeckman<sup>2,3+</sup>, Yanlu Li<sup>4</sup>, Soren Aasmul<sup>5</sup>, Umberto Morbiducci<sup>6</sup>, Roel Baets<sup>4</sup>, Pierre Boutouyrie<sup>7</sup>, Filippo Molinari<sup>1</sup>, Nilesh Madhu<sup>3</sup> and Patrick Segers<sup>2\*</sup>

<sup>1</sup> *Polito<sup>BIO</sup> Med Lab, Biolab, Department of Electronics and Telecommunications Politecnico di Torino, Turin, Italy*

<sup>2</sup> *IBiTech-bioMMeda, Ghent University, Ghent, Belgium*

<sup>3</sup> *IDLab, Ghent University - imec, Ghent, Belgium*

<sup>4</sup> *Photonics Research Group, Ghent University-imec, Technologiepark-Zwijnaarde 126, 9052 Ghent, Belgium and Center for Nano- and Biophotonics, Ghent University, Technologiepark-Zwijnaarde 126, 9052 Ghent, Belgium*

<sup>5</sup> *Medtronic Bakken Research Center, Maastricht, The Netherlands*

<sup>6</sup> *Department of Mechanical and Aerospace Engineering, Politecnico di Torino, Turin, Italy*

<sup>7</sup> *INSERM U970, Université de Paris, Assistance Publique Hôpitaux de Paris, Paris, France*

<sup>+</sup> *Both authors contributed equally*

Correspondence\*:

Patrick Segers, Ghent University, IBiTech-bioMMeda, C. Heymanslaan 10, 9000 Ghent, Belgium  
patrick.segers@ugent.be

## 2 ABSTRACT

3 **Background:** Laser-Doppler Vibrometry (LDV) is a laser-based technique that allows measuring  
4 the motion of moving targets with high spatial and temporal resolution. To demonstrate its use  
5 for measurement of carotid-femoral pulse wave velocity, a prototype system was employed in a  
6 clinical feasibility study. Data were acquired for analysis without prior quality control. Real-time  
7 application, however, will require real-time assessment of signal quality. Here, we (1) use template  
8 matching and matrix profile for assessing the quality of these previously acquired signals; (2)  
9 analyse the nature and achievable quality of acquired signals at the carotid and femoral measuring  
10 site; (3) explore models for automated classification of signal quality.

11 **Methods:** LDV data were acquired in 100 subjects (50M/50F) and consisted of 4-5 sequences  
12 of 20-second recordings of skin displacement, differentiated twice to yield acceleration. Each  
13 recording consisted of data from 12 laser beams, yielding 410 carotid-femoral and 407 carotid-  
14 carotid recordings. Data quality was visually assessed on a 1-5 scale, and a subset of best quality  
15 data was used to construct an acceleration template for both measuring sites. The time-varying  
16 cross-correlation of the acceleration signals with the template was computed. A quality metric  
17 constructed on several features of this template matching was derived. Next, the matrix-profile  
18 technique was applied to identify recurring features in the measured time series and derived a  
19 similar quality metric. The statistical distribution of the metrics, and their correlates with basic  
20 clinical data were assessed. Lastly, logistic-regression-based classifiers were developed and their  
21 ability to automatically classify LDV-signal quality was assessed.

22 **Results:** Automated quality metrics correlated well with visual scores. Signal quality was  
23 negatively correlated with BMI for femoral recordings, but not for carotid recordings. Logistic  
24 regression models based on both methods yielded an accuracy of minimally 80% for our carotid  
25 and femoral recording data, reaching 87% for the femoral data.

26 **Conclusions:** Both template matching and matrix profile were found suitable methods for  
27 automated grading of LDV signal quality and were able to generate a quality metric that was  
28 on par with signal quality assessment of the expert. The classifiers, developed with both quality  
29 metrics, showed their potential for future real-time implementation.

30 **Keywords:** Laser Doppler Vibrometry (LDV), Matrix profile, Template matching, Logistic regression, Signal quality

## 1 INTRODUCTION

31 The aorta and large central arteries fulfil key physiological functions in the circulation, whereby their  
32 structure is apt to their function. They consist of complex composite soft tissues, concentrically organized in  
33 lamellar units, where sheets of elastin intertwine with layers of vascular smooth-muscle cells in a matrix of  
34 collagen and other proteins composing the extra-cellular matrix (Wolinsky and Glagov, 1967). This allows  
35 the aorta and large arteries to distend when the heart contracts and blood is ejected into the aorta and store  
36 elastic energy in the arterial wall, which is used during the relaxation phase of the heart to maintain blood  
37 pressure and drive the perfusion of organs and tissues. This function is also referred to as the “windkessel”  
38 or buffering function of the large arteries, and ensures that the pulsatile blood flow generated by the heart is  
39 transformed into a near steady flow when reaching the smaller arteries (Westerhof et al., 2009). It prevents  
40 excessive maximal (systolic) and too low minimal (diastolic) blood pressure. Arterial stiffening leads to  
41 a loss of this buffering function with detrimental effects on nearly all organ systems, and especially low  
42 resistance organs such as the brain, the kidneys and the heart itself (Chirinos et al., 2019). Arterial stiffening  
43 has received large attention over the past 3 decades, and there is a consensus that assessment of arterial  
44 stiffness is especially relevant in the assessment of an individual’s risk for cardiovascular disease and death  
45 (Vlachopoulos et al., 2010; Laurent et al., 2006).

46 Because of the distensible nature of arteries, cardiac contraction generates a wave (detectable as a change  
47 in pressure, flow or arterial diameter). This wave initially propagates from the heart to the periphery, but  
48 increases in complexity as it interacts on its way with the branching arterial tree and gets shaped because  
49 of wave reflection and transmission (Chirinos et al., 2019; Mitchell et al., 2004, 2011; O’Rourke and  
50 Kelly, 1993). The wave speed, or pulse wave velocity (PWV), is directly linked with the distensibility of  
51 the arteries (the stiffer the artery, the higher PWV) (Bramwell and Hill, 1922), and the current clinical  
52 standard method to measure arterial stiffness is by measuring the pulse wave velocity (Segers et al., 2020).  
53 In essence, the method is simple and straightforward: one detects the pulse at two locations a distance  $dx$   
54 apart, and from the time delay,  $dt$ , between the signals, one gets  $PWV = dx/dt$ . Despite the simplicity of the  
55 concept, there are still many hurdles in measuring PWV in practice, mainly related to the non-availability  
56 of sites to directly measure the pulse along the path of the aorta in a non-invasive way and without the need  
57 of clinical scanners (Segers et al., 2020). Accessible sites closest to the aorta are the neck (carotid artery)  
58 and groin (femoral artery) and carotid-femoral PWV is considered the best possible proxy for aortic PWV  
59 (Laurent et al., 2006).

60 Several sensors can be used to detect the pulse in the neck and groin (Segers et al., 2020; Pereira  
61 et al., 2015), including applanation tonometry, ultrasound (pulsed Doppler recordings) or accelerometers.  
62 Motivated by the relatively high cost of equipment, the required level of expertise by the operator or  
63 contact-based nature of the measurement, we and others have explored the use of laser Doppler vibrometry  
64 to detect the motion of the skin atop the carotid and/or femoral arteries in response to the passage of the  
65 arterial pulse (De Melis et al., 2008; Scalise and Morbiducci, 2008; Campo and Dirckx, 2011; Kaplan et al.,  
66 2012; Morbiducci et al., 2007b). To eliminate motion drift and amplify the fast displacements associated  
67 with the arrival of the foot of the pulse (Morbiducci et al., 2007b), we have been using skin acceleration as  
68 the basic signal from which to derive time delays between the neck and groin for measuring carotid-femoral  
69 PWV.

70 The feasibility of the method has been shown using industrial-type LDV sensors (De Melis et al., 2008),  
71 and we have been working on the design and development of a multi-beam handheld device. The core  
72 of the device is a silicon photonics chip integrated in a micro-optical system which allows for flexible  
73 and compact multi-array designs (Li et al., 2013, 2020). A first prototype (consisting of 2 connected yet  
74 separable handheld pieces to measure in the neck and groin with each 6 laser beams) was developed  
75 within the context of the H2020-funded project CARDIS and included a clinical feasibility study whereby  
76 carotid-femoral PWV was assessed in 100 patients and compared with a reference method based on

77 applanation tonometry (Marais et al., 2019). Measurements were performed with a minimal visual feedback  
78 during the measurements and all the analyses were carried out in off-line modality.

79 A next generation version of the device is under development and will provide real-time measurement of  
80 carotid-femoral PWV. To do so, we need real-time assessment of the quality of incoming data to decide  
81 whether or not data records are of an acceptable quality for subsequent processing. This is, however, not a  
82 trivial assessment as there is little reference as to what makes LDV signal recordings appropriate for PWV  
83 estimation.

84 The aim of this study is therefore to identify a strategy to objectively and automatically assess the  
85 LDV-signal quality and set criteria for future use of this technology in arterial pulse detection. To do that  
86 we will use the existing CARDIS database of LDV recordings at the carotid and femoral measurement sites  
87 and subject them to two different strategies: the template matching and the matrix profile will be tested for  
88 (1) analysing the nature and achievable quality of the recorded signals, and (2) exploring models for an  
89 automated classification of LDV-signal quality.

## 2 MATERIALS AND METHODS

### 90 2.1 The CARDIS device

91 Technical details on the optics and overall design of the CARDIS device have been described in (Li  
92 et al., 2020). Briefly, the device consists of two handpieces (handpiece 1 contains the handgrip of the  
93 device, handpiece 2 is the add-on part of the device: we refer to Figure 1 for an illustration of the device  
94 and the positioning of the handpieces), each sending out 6 laser beams (wavelength 1550 nm), positioned  
95 along a line and 5 mm apart. The handpieces can be used separately for measurement of carotid-femoral  
96 PWV, or attached to measure signals on locations 25 to 50 mm apart, e.g. to locally measure pulse wave  
97 propagation along the carotid artery. A retro-reflective tape is attached to the skin at the measurement  
98 location to enhance reflection of the laser light, and the device is equipped with a spacer to ensure an  
99 appropriate optical focus distance and to stabilize measurements.

### 100 2.2 Study population and available database

101 The data used in this study were acquired with a clinical feasibility study in 100 patients, conducted at the  
102 Hôpital Européen Georges Pompidou (HEGP) in Paris, France, to assess the ability of the CARDIS device  
103 to measure signals in a configuration with simultaneous carotid-femoral or carotid-carotid recordings.  
104 Patients were in the age range 19-85 and presented with mild to stage 3 hypertension, controlled or not  
105 (Marais et al., 2019). For each subject, 4 to 5 datasets, each consisting of 20 second traces on 12 channels  
106 measured with the two handpieces, were acquired. In detail, the analysed database was made of 410  
107 datasets (4920 waveforms) from carotid-femoral recordings, and of 407 datasets (4884 waveforms) from  
108 carotid-carotid recordings. Raw IQ (In-phase and quadrature) LDV-data were acquired at a sampling  
109 frequency of 100 kHz, and LDV-displacement data were downsampled to 10 kHz upon demodulation.  
110 A low-pass filter with cut-off frequency of 30 Hz was applied to LDV displacement data, which were  
111 differentiated twice to yield acceleration. The same low-pass filtering strategy was applied after each  
112 differentiation operation.

### 113 2.3 Visual scoring of the data

114 A graphical interface displaying all the LDV acceleration signals derived from the six channel recordings  
115 per each handpiece was implemented in MATLAB environment (The MathWorks, Naticks MA, US). The  
116 acceleration signals were visually scored by an expert operator (Segers P.) on a 5-level grade scale taking  
117 values  $Q_{\text{vis}}$  according to table 1.

118 Note that the presence of brief artefacts in the 20 second acquired traces was not used as a criterion to  
119 score the signal quality. As such, signals qualified as excellent may still demonstrate a brief episode of poor  
120 data. Overall, the femoral data were of a markedly lower  $Q_{\text{vis}}$  'quality' than traces recorded at the carotid  
121 artery, which impacted the rating. Therefore, the  $Q_{\text{vis}}$  quality score 3 (borderline) was given to femoral  
122 traces that appeared to be of a much lesser quality than  $Q_{\text{vis}} = 3$  rated carotid traces. Such a borderline  
123 score was assigned when 5-10 beats were discernible in the signal. Representative carotid and femoral  
124 signals receiving the different scores are displayed in Figure 1.

## 125 2.4 Template Matching

126 Template matching technique is an effective approach for the automatic detection of a priori identified  
127 patterns in signal recordings (Won-Du and Chang-Hwan, 2014; Jiun-Hung et al., 2003) and images (Omachi  
128 and Omachi, 2007). A good-quality carotid LDV acceleration signal presents two sharp peaks for each  
129 heartbeat: the first peak corresponds to the systolic rapid upstroke of pressure and demarcates the foot of  
130 the arterial pulse; the second peak denotes the wave that is generated at the moment of closure of the aortic  
131 valve (the dicrotic notch). The LDV-femoral recording is devoid of clearly identifiable features related to  
132 the dicrotic notch because of the distance of the measurement site from the heart, whose final effect is  
133 filtering the recorded LDV pulses, in the femoral artery. An example of displacement, acceleration and  
134 ECG signals together are shown in Figure 2.

### 135 2.4.1 Constructing the templates

136 High-quality carotid and femoral LDV-acceleration traces were adopted for template construction. Traces  
137 with visual score value  $Q_{vis}$  of 4 and 5 were selected. To avoid subject-specific biasing in template  
138 construction, only one 20 s recording (from the acquired channel with highest  $Q_{vis}$ ) per subject was  
139 selected. Based on these selection criteria, 135 carotid LDV-acceleration traces from 20 different subjects  
140 and 40 femoral LDV-acceleration traces from 10 different subjects were identified as suitable for template  
141 construction in the CARDIS dataset. The selected carotid LDV-acceleration traces were from both  
142 handpieces.

143 The selected traces, characterized by the presence of sharp and pronounced peaks at the foot (and dicrotic  
144 notch for carotid recordings), were then segmented in epochs, each one corresponding to a single heartbeat.  
145 LDV-acceleration trace segmentation was carried out using ECG synchronous recordings (available for  
146 each subject in the CARDIS dataset, on which automatic R-peak detection was carried out, see Figure 3).  
147 Over each LDV trace single epochs were then defined within a time interval within the occurrence of two  
148 consecutive R peaks in the ECG trace (Figure 3A). By construction of the visual inspection classification,  
149 some of the identified single epochs might still not be of adequate quality for template construction,  
150 because of the presence of short-time artifacts/noise (Figure 3B). The lower quality single epochs in a  
151 LDV-acceleration trace were identified according to the following strategy: (1) for each LDV segmented  
152 trace a correlation matrix  $R_{ij}$  was built up, each element of the matrix being the Pearson-correlation  
153 coefficient between epochs  $i$  and  $j$ , used as a measure of their shape similarity; (2) a threshold value of the  
154 correlation coefficient was defined and single epochs with an average correlation coefficient with all the  
155 other epochs lower than the threshold was discarded, since they were not sufficiently similar in shape to the  
156 other epochs in the recorded trace (Figure 3C); (3) for each LDV-acceleration trace an ‘individual template’  
157 was built up by averaging only the identified highly correlated epochs (Figure 3D); (4) by adopting the same  
158 approach with the carotid and femoral LDV-acceleration traces, the final carotid and femoral ‘population  
159 templates’ were obtained (Figure 4).

160 Template construction is based upon the definition of a strategy to treat the issue of the different time  
161 length of single epochs (intra-individual RR variability) (Jensen-Urstad et al., 1997; Zhang, 2007) and of  
162 the individual templates as well. Hence, the time length of single epochs should be defined on the basis  
163 of what the template should represent. In the case under study, the carotid LDV-acceleration template  
164 longer than 350 ms will include by construction the foot of the wave (first peak) and the dicrotic notch  
165 (second peak). Here we speculate that a carotid LDV template incorporating the second peak may degrade  
166 in performance, as the distance between the two peaks is (intra-individually as well as inter-individually)  
167 variable. In figure 5, carotid and femoral LDV-acceleration templates constructed for different (predefined)  
168 time length are displayed. In detail, time lengths of 300, 400 and 500 ms were considered for the femoral  
169 LDV-acceleration template, and time lengths of 200, 400 and 600 ms for the carotid LDV-acceleration  
170 template. The impact of the time length in the LDV template performance when used for the automatic  
171 assessment of the quality of the CARDIS data was evaluated.

### 172 2.4.2 Template matching and beat selection

173 The matching between the templates and the LDV-acceleration traces in the CARDIS dataset was  
174 performed by applying a local moving-window function calculating the Pearson’s correlation coefficient  
175 between the LDV template and the 20 s-long acceleration trace at each time step, as displayed in Figure 6.

176 The locations of peaks in the time series resulting from this moving-window cross-correlation operation  
 177 identify the time instants where the sliding template is similar to a segment of the LDV-acceleration trace.

178 Setting a threshold for the value of the cross-correlation coefficient then demarcates the correspondence  
 179 level above which segments of the LDV-acceleration trace can be considered similar to the template. Based  
 180 on the set threshold value, single segments corresponding to single heartbeats in the LDV trace can be  
 181 considered of sufficient or not sufficient quality.

182 To further improve the identification of high-quality heartbeats in the LDV recorded traces, two further  
 183 selective criteria were added. Firstly, all the LDV-acceleration peaks in the recorded trace with an amplitude  
 184 lower than the 80% of the average peak amplitude were not considered. Then, if two successive peaks  
 185 were detected within a time window shorter than 500 ms, the second peak was discarded and only the  
 186 first one was considered. The latter criterion was adopted to avoid the dicrotic notch detection (second  
 187 peak), especially when the shorter carotid template was used. An explanatory example of peak detection,  
 188 presenting the LDV-acceleration trace, the moving-window cross-correlation function and detected peaks  
 189 is displayed in Figure 7.

#### 190 2.4.3 LDV traces classification based on template matching - finding threshold values

191 The performance of the template matching algorithm in classifying the quality of the CARDIS dataset was  
 192 evaluated by comparison with visual score classification, according to the following scheme: acceptable  
 193 heartbeat (label 1), corresponding to  $Q_{\text{vis}}$  values 4 or 5; not acceptable heartbeat (label 0) corresponding to  
 194  $Q_{\text{vis}}$  values 1 or 2. Signals with  $Q_{\text{vis}}$ -values of 3 are discarded in this analysis as these signals are difficult  
 195 to assign an absolute and correct classification (see discussion). The template matching-based classification,  
 196 as also mentioned before, depends upon the threshold value for the moving-window correlation function  
 197 and the number of detected heartbeats in the LDV-acceleration trace, which have to be appropriately set.

198 Here, we considered: true positive (TP) an acceptable LDV trace (based on  $Q_{\text{vis}}$  classified by the template  
 199 matching as acceptable; false negative (FN) an acceptable LDV trace classified by the template matching  
 200 as not acceptable; true negative (TN) an unacceptable LDV trace classified by the template matching as not  
 201 acceptable; false positive (FP) an unacceptable LDV trace classified by the template matching as acceptable.  
 202 On this basis, sensitivity and specificity values of the classifier are defined as:

$$\text{Sensitivity} = \frac{TP}{TP + FN} \quad (1)$$

203 and

$$\text{Specificity} = \frac{TN}{TN + FP} \quad (2)$$

204 Sensitivity and Specificity were then used to build up the Receiver Operating Characteristic (ROC) curves  
 205 and their area under the curve (AUC) was used to assess the performance of the classifier. Moving-window  
 206 cross-correlation coefficient threshold values and number of detected heartbeats yielding the highest AUC  
 207 were defined on the complete CARDIS dataset, and this for each one of the template lengths in time.

#### 208 2.4.4 LDV traces classification based on template matching - defining quality score and testing 209 on the CARDIS dataset

210 Once the best performing carotid and femoral templates time length and the associated moving-window  
 211 cross-correlation threshold values were identified, a quality score ( $Q_{\text{TM}}$ ) was estimated for each 20 s  
 212 LDV-trace recording, based on two main features.

213 The first feature ( $Q_1$ ) is the number of the detected acceleration peaks ( $n_{\text{peaks}}$ ), normalized with the  
 214 maximum expected number of peaks or heartbeats in the 20 s LDV-trace recording ( $\text{max}_{\text{peaks}}$ ). This value  
 215 was empirically set equal to 26 to ensure a maximal feature value of 1 in the investigated database:

$$Q_1 = \frac{n_{\text{peaks}}}{\max_{\text{peaks}}} \quad (3)$$

216 The second feature ( $Q_2$ ) is defined as the average time delay between the occurrence of maximum value  
 217 of each LDV-acceleration epoch in the recorded trace and the occurrence of the peak value on the template  
 218 ( $d_{\text{peak}_n}$ ), normalized to the template time length ( $N$ ):

$$Q_2 = \frac{\sum_{n=1}^{n_{\text{peaks}}} (1 - \frac{d_{\text{peak}_n}}{N})}{\max_{\text{peaks}}} \quad (4)$$

219 When the peaks in the template and in each LDV epoch are all perfectly aligned, and when all the peaks  
 220 in the LDV-trace are detected (i.e.,  $Q_1 = 1$ ), feature  $Q_2$  is equal to 1, indicating good quality of the LDV  
 221 trace recording. The final score based on template matching can be computed as the mean value of the  
 222 partial scores  $Q_1$  and  $Q_2$ :

$$Q_{\text{TM}} = \frac{1}{2}(Q_1 + Q_2) \quad (5)$$

223 By construction, the score  $Q_{\text{TM}}$  was set up so that the value is within the range  $[0, 1]$  (with  $Q_{\text{TM}} = 0$   
 224 representing the worst possible signal quality and  $Q_{\text{TM}} = 1$  indicating that the signal is of excellent quality).  
 225  $Q_{\text{TM}}$  was calculated for all the traces in the CARDIS database and compared to the corresponding assigned  
 226 visual score  $Q_{\text{vis}}$ , which is treated as the ground truth.

#### 227 2.4.5 A logistic regression model for signal classification based on template matching

228  $Q_{\text{TM}}$  Was a heuristically derived quality metric with equal weighting on the sub-components. Now we use  
 229 logistic regression models to find a better weighting of the contributions of  $Q_1$  and  $Q_2$ , and automatically  
 230 map this to a predicted quality of the signal. Logistic regression models are chosen since they can be  
 231 well applied to binary classification problems, and are typically used in medical research (Domínguez-  
 232 Almendros et al., 2011; Nick and Campbell, 2012; Austin and Steyerberg, 2012) when a two-class classifier  
 233 is required. These predictions were then compared to the ground truth labels (given by the visual scores).

234 Logistic regression models were trained and tested with the two template-matching derived scores (3)  
 235 and (4) as features, on both carotid and femoral LDV-acceleration traces. For this purpose, again the LDV  
 236 traces visually scored with  $Q_{\text{vis}}$  equal to 1 or 2 were labeled 0, and LDV traces visually scored with  $Q_{\text{vis}}$   
 237 equal to 4 or 5, were labeled 1. Again, signals with  $Q_{\text{vis}}$  score 3 were not included in the analysis.

238 The data available in the CARDIS database was split such that 80% was used for training the logistic  
 239 regression model and the remaining 20% used for testing purposes. The training-testing set partition was  
 240 randomly iterated 1000 times while storing the model accuracy every iteration, so that the overall accuracy  
 241 distribution of the logistic regression model approach could be assessed.

242 Of note, all features used to train logistic regression models were normalized via standardization. This  
 243 allowed the logistic regression-model coefficients to be interpreted as the corresponding feature weights,  
 244 granting information about which feature was most influential in labeling an LDV trace.

245 The accuracy distributions of logistic regression models trained on template-matching and the later  
 246 discussed matrix-profile derived features were evaluated.

## 247 2.5 Matrix profile

248 The matrix profile is a data structure that annotates a time series (Yeh et al., 2018; Zhu et al., 2020). It  
 249 allows for exact, simple and fast (Zhu et al., 2017b) similarity search or discord discovery and is among  
 250 the state-of-the-art techniques in the field of discrete time-series analysis (Zhu et al., 2017a; Madrid et al.,  
 251 2019). The matrix profile has been used in processing biological signals like EEG (Mueen et al., 2009),  
 252 ECG and gait cycles (Zhu et al., 2020). It was applied here to accurately identify recurring waveforms  
 253 in the LDV-acceleration data. Every such waveform is a subsequence of the original sequence or time

254 series. These subsequences, taken together, are collectively called a motif. We gauged the quality of an  
 255 LDV-measurement via several features determined by its best motif. The strength of the matrix profile  
 256 lies in the fact that it does not require a template or other input parameters except for the length  $m$  of the  
 257 desired motif subsequences. Analogous to the template matching analysis, waveforms were subsampled to  
 258 1 kHz. We set  $m$  to 200 ms, similar to the optimal length of the template described in previous sections.

### 259 2.5.1 Signal classification based on the matrix profile

260 A quality metric ( $Q_{MP}$ ) was constructed based on three features of the matrix profile-generated motif as  
 261 seen in (6). This metric was constructed so that its possible values lie between 0 and 1.

$$Q_{MP} = A_{MP} t_{d,MP} n_{MP} \quad (6)$$

262 The first feature used in calculating ( $Q_{MP}$ ) is the average relative maximum amplitude of a subsequence  
 263 in the motif ( $A_{MP}$ ) computed as in (7). The maximum amplitude of a subsequence  $A$  was compared with  
 264 the maximum amplitude of the reference subsequence  $A_{ref}$ . This reference is the first subsequence identified  
 265 by the matrix profile (the minimum of the matrix profile) and subsequently included in the motif. In good  
 266 quality measurements, most maximum amplitudes of subsequences in the motif were similar.

$$A_{MP} = \frac{1}{n_{mtf}} \sum_{n=1}^{n_{mtf}} \frac{A}{A_{ref}} \quad (7)$$

267 The second feature, the average relative time-instant of the subsequence peaks in the motif ( $t_{d,MP}$ ), is  
 268 computed as in (8). The time-instant of the subsequence peak was compared with that of the reference.  
 269 This value was then normalized over the length of the subsequence  $m$ . Ideally, all subsequences in the  
 270 motif represent the same heartbeat-related waveform with peaks at similar time instants. For poor quality  
 271 signals, these time instants tended to randomly vary over the length of the subsequence.

$$t_{d,MP} = \frac{1}{n_{mtf}} \sum_{n=1}^{n_{mtf}} \left(1 - \frac{d_{peak}}{m}\right) \quad (8)$$

272 Lastly, the third feature ( $n_{MP}$ ) was calculated as the expected amount  $n_{exp}$  versus the effective amount  
 273  $n_{mtf}$  of subsequences in the motif, shown in (9).  $n_{exp}$  was estimated based on a discrete-Fourier-transform  
 274 analysis of the entire signal recording. More specifically, the peak corresponding to the heartbeat during  
 275 the measurement was identified as the most prominent peak in the signal spectrum, in the range 0.5 -  
 276 1.5 Hz. The effective amount of subsequences in the motif  $n_{mtf}$  was based on how many heartbeats the  
 277 matrix-profile technique was able to pick up.

$$n_{MP} = \frac{n_{mtf}}{n_{exp}} \quad (9)$$

278 Before a subsequence is included in the motif, three criteria decide the inclusion: (1) If a subsequence  
 279 maximum amplitude was lower than 0.8 times the reference maximum amplitude it was excluded from the  
 280 motif. (2) If the time instant of the peak deviated 30 ms or more from that of the reference, the subsequence  
 281 was also removed from the motif. (3) If two subsequences were closer than 0.8 times the expected time  
 282 delay between two subsequent heartbeats, the one with the lower matrix-profile value (higher similarity  
 283 to the reference) of the two was preserved, the other was removed. The applied thresholds levels were  
 284 determined empirically from excellent and poor quality signals.

285 Figure 8 shows an example of a signal being scored by first finding the motif so that as many heartbeats  
 286 as possible are present within it, then calculating the features of that motif. Both the relative amplitude and  
 287 time-instant of subsequence peak features of one subsequence in the motif are indicated on the figure.

288 The auto-generated matrix-profile based quality metric was computed for all carotid-carotid and femoral-  
 289 carotid datasets and results were compared to the visual scores.

## 290 2.5.2 A Logistic regression model based on the matrix profile

291 Similar to template matching, we also designed logistic regression models using the previously discussed  
292 matrix-profile derived features. These models allowed for more freedom in weighting the features to come  
293 to a better classification result. Models were trained and tested on the three features mentioned above.  
294 Signals were labeled and available data was split into training and testing sets analogous as in the previously  
295 discussed template-matching case.

## 296 2.6 Relation between signal quality and physiological variables

297 Lastly, we investigated the existence of possible associations of quality of the LDV-acceleration traces  
298 with age, body mass index (BMI) and systolic blood pressure. The statistical analysis was performed  
299 using  $Q_{MP}$  as quality score variable. In detail, the existence of a linear correlation was tested using the  
300 Pearson-correlation coefficient on both CC and CF datasets, with data analysed per handpiece. For all  
301 analyses, significance was assumed for  $p < 0.05$ .

# 3 RESULTS

## 302 3.1 Visual scoring

### 303 3.1.1 Carotid-carotid measurements

304 By visual inspection, about 12% of all LDV-acceleration traces were qualified as bad and close to 30%  
305 as poor (Figure 9, panel B). Which implies that about the 42% of the recorded LDV traces was evaluated  
306 to not be of sufficient quality for further analysis. About 22% of all recordings was scored from good  
307 to excellent, and are deemed suitable for further analysis. About 37% of the traces was visually scored  
308 borderline, i.e. these traces might be of sufficient quality for further analysis with advanced processing.  
309 The number of LVD traces scored with  $Q_{vis}$  4 or 5, and recorded using handpiece 2 was higher than the  
310 number using handpiece 1. For handpiece 1, channel 1 scored almost systematically very low; the best  
311 channels were channels 3 and 4. For the second handpiece, the best channels were channels 2 and 3.

### 312 3.1.2 Carotid-femoral measurements

313 The bottom row of Figure 9 illustrates that, concerning femoral LDV-acceleration traces (handpiece  
314 1), 20% of all recordings was qualified as bad, and another 32% as poor, meaning that over 50% of all  
315 recordings is not usable for analysis. About 15% of the measured signals get a score good to excellent,  
316 deemed immediately suitable for analysis. Best channels are channels 3 and 4 with 21.9% (beam 3) and  
317 19.5% (beam 4) of the recordings good to excellent. For handpiece 2 (carotid recordings), about 20%  
318 gets a score good to excellent. This is less than what was obtained for handpiece 2 for the carotid-carotid  
319 recordings, where close to 25% of all recordings were rated good to excellent. On the other hand, less  
320 signals received grade 1 and 2. Best channels are channels 4 (24.9%) and 5 (24.1% of the recordings  
321 scoring good to excellent).

## 322 3.2 Template matching

### 323 3.2.1 Carotid-carotid (CC) measurements

324 From the analysis carried out on the complete CC dataset, it emerged that using the carotid template of  
325 200 ms length guarantees the best performance in terms of specificity, setting the cross-correlation threshold  
326 to 0.74 and the minimum number of detected heartbeats per trace to 15 (AUC = 0.89, sensitivity 74%,  
327 specificity 89%; template of 400 ms length: AUC = 0.89, sensitivity 81%, specificity of 83%; template of  
328 600 ms length: AUC = 0.92, sensitivity 87%, specificity 86%). For each template length, the corresponding  
329 confusion matrix is presented in Table 2. The adoption of specificity for the evaluation of the performance  
330 of the template matching strategy was dictated by the need of maximizing the removal of LDV traces with  
331 inadequate quality. More in detail, it emerged that in general the template matching performed excellently  
332 in correctly classifying visual scores 1 and 5, while accuracy decreased for visual scores 2 and 4 (Table 2).  
333 Interestingly, using the shorter template length of 200 ms led to score 42% of the LDV acceleration traces  
334 visually scored 3 (borderline) as acceptable data.



335 The level of agreement obtained between  $Q_{TM}$  and  $Q_{vis}$  on the CC recordings dataset, with template  
336 matching adopting a 200 ms template length, is presented in Figure 10. This suggests that the median of  
337 the  $Q_{TM}$  values, computed on traces that have a  $Q_{vis} = 3$ , could be adopted as a threshold value for the  
338 automatic quality checking of an LDV trace (i.e. in the case under study, traces with a  $Q_{TM} > 0.5$  could be  
339 considered of adequate quality; note that, manually setting these thresholds is not required for the logistic  
340 regression models since this is implicitly learnt in the training).

341 The accuracy distributions of logistic regression models trained on quality scores derived from template-  
342 matching are displayed in figure 11( 11A- 11B). On average, the accuracy on traces acquired using  
343 handpiece 2 is higher than handpiece1 ( $85\pm 1.6\%$  and  $80\pm 1.70\%$ , respectively; the results are summarized  
344 in Table 4).

### 345 3.2.2 Carotid-femoral (CF) measurements

346 From the analysis carried out on the complete CF dataset, it emerged that using the carotid template of  
347 500 ms length guarantees the best performance in terms of specificity, setting the cross-correlation threshold  
348 to 0.56 and the minimum number of detected heartbeats per trace to 10 (AUC = 0.89, sensitivity 77%,  
349 specificity 92%; template of 400 ms length: AUC = 0.88, sensitivity 76%, specificity of 91%; template of  
350 300 ms length: AUC = 0.87, sensitivity 72%, specificity 92%). The confusion matrices are shown in Table 3  
351 for each template.

352 As for the carotid traces, the performance of the template matching algorithm was based on the specificity  
353 values, in order to remove the bad quality signals. More in detail, the template matching strategy shows  
354 excellent performance for the classification of visual scores 1 and 5 (accuracy of 96% and 93% respectively),  
355 while the accuracy decreases for class 2 and class 4 (91% and 72% respectively). In the femoral case, the  
356 method classified a majority of LDV traces with a visual score of 3 (borderline) as inadequate. Indeed,  
357 considering the template of 500 ms, the template matching method classified 65% of score 3 as inadequate  
358 signals and the other 35% (borderline) as adequate.

359 The level of agreement obtained between  $Q_{TM}$  and  $Q_{vis}$  on the CF recordings dataset, using the 500 ms  
360 template length, is shown in Figure 10C. The results indicate that from the median  $Q_{TM}$  values scored  $Q_{vis}$   
361 = 3, a threshold value could be adopted for the automatic quality checking of the LDV trace (i.e. in the  
362 case under study, traces with a  $Q_{TM} > 0.23$  could be considered of adequate quality; again, this threshold  
363 is not required when working with the logistic regression models.)

364 The accuracy distributions of logistic regression models trained on quality scores derived from template-  
365 matching are displayed in figure 11( 11E and 11F). On average, the accuracy on traces acquired using  
366 handpiece 1 is higher than handpiece 2 ( $87\pm 1.3\%$  and  $81\pm 1.9\%$ , respectively; the results are summarized  
367 in Table 4).

## 368 3.3 Matrix profile

369 On good quality data, i.e., those visually scored at 4 or 5, the matrix profile technique was able to include  
370 nearly all heartbeats in the motif. On poor quality data the matrix profile was unable to identify most  
371 heartbeats because of noise or artifacts in the measurement. On some measurements that contain pure noise,  
372 the matrix profile picked up random noisy waveforms that were less prevalent and differed much compared  
373 to the desired foot-of-the-wave waveform.

### 374 3.3.1 Quality metric results

375 The signals measured at the carotid measuring site were given a matrix profile-derived quality score that is  
376 compared with their visual scores in figure 10B. A positive, linear relation between the two scoring methods  
377 is observed for the carotid-carotid database. The same information is shown for the femoral measuring site  
378 in figure 10D. The difference between poor and good quality signals is apparent. Signals with visual score  
379 1,2 or 3 have significantly lower  $Q_{MP}$  than those with visual score 4 or 5.

### 380 3.3.2 Logistic regression models performance

381 Figure 11C,D,H show the accuracy distributions of the repeated logistic regression model-training  
382 experiment for signals measured in the neck with the different handpieces. All accuracy averages are  
383 above 80% with 82% ( $\pm 1.64\%$ ) and 88% ( $\pm 1.53\%$ ) for carotid-carotid recordings with handpiece 1 and 2,

384 respectively. For carotid-femoral recordings, carotid data recorded with handpiece 2 yielded an accuracy of  
385  $85\% \pm 1.71\%$ . The distributions were assumed to be normally-distributed after a Shapiro-Wilk test and  
386 thus the Gauss-curves are drawn onto the subfigures of figure 11.

387 The same data for the femoral data (measured with handpiece 1 during carotid-femoral recordings), is  
388 shown in figure 11G. An average accuracy of  $86\%$  with a standard deviation of  $1.43\%$  is observed. All  
389 accuracy statistics of the different measurement situations are summarized in table 4.

### 390 3.4 Signal quality vs. physiological variables

391 The results from the correlation analysis between  $Q_{MP}$  and age, BMI and systolic blood pressure  
392 are shown in figure 12 for the femoral data (carotid-femoral recording, handpiece 1; CF hp1) and the  
393 carotid-carotid recordings with handpiece 2 (CC hp2) showing the strongest trends. Significant negative  
394 correlations were found between age and  $Q_{MP}$  for CF hp1 ( $r=-0.253$ ,  $P<0.05$ ) and CC hp2 ( $r=-0.365$ ,  
395  $P<0.001$ ). The correlation with BMI (figure 12B) was significant only for the femoral recording ( $r=-0.304$ ,  
396  $P<0.01$ ) while the correlation with systolic blood pressure was significant only for CC hp2 ( $r=-0.206$ ,  
397  $P<0.05$ )(figure 12C). In a multivariate regression model including both age and systolic blood pressure,  
398 the correlation between carotid signal quality and systolic blood pressure was no longer significant (due to  
399 the correlation between age and systolic blood pressure). In contrast, in a multivariate model of femoral  
400 signal quality, both age and BMI remained significantly correlated with signal quality. The same relations  
401 are found when repeating the analysis with  $Q_{vis}$  or  $Q_{MP}$  (data not shown).

## 4 DISCUSSION

402 The potential of Laser-Doppler Vibrometry (LDV) for non-contact measurement of physiological  
403 (cardiovascular) signals has been reported since about 2000 in explorative studies (Pinotti et al., 1998;  
404 Kaplan et al., 2012; Morbiducci et al., 2007a; Rohrbaugh et al., 2013) making use of bulky industry-time  
405 devices, and the technique has been suggested for measurement of carotid-femoral PWV by (De Melis  
406 et al., 2008). An important technological breakthrough to enable LDV-based measurements in a clinical  
407 setting is the use of silicon photonics to miniaturize and integrate the optical components onto chips (Li  
408 et al., 2013) that are easily built in into hand-held devices as the CARDIS prototype used in this study. That  
409 prototype was used in a clinical feasibility study where measurements were performed on the carotid and  
410 femoral artery and we previously reported on the agreement of LDV-based carotid-femoral PWV with a  
411 reference method (Marais et al., 2019). In that paper, data was processed off-line and algorithms for foot  
412 detection relied on the ECG and gating was applied on carotid and femoral tracings to ensure identification  
413 of the correct characteristic points on the waveforms. Further developments aim for ECG-independent  
414 measurements and will require a more stringent quality assessment in real-time application to ensure that  
415 data is captured from which transit times can be derived. Unlike the CARDIS device, future versions of the  
416 device will provide real-time feedback on signal quality and valid measurements will only be accepted after  
417 a minimal number of data samples have been retrieved from signals passing predefined quality criteria. In  
418 this paper, we explored two possible strategies for such quality assessment, template matching and matrix  
419 profile, and benchmarked them using visual scoring as reference.

420 The visual grading was done by what we considered an expert observer, but is inherently subjective.  
421 The graphical user interface that was developed showed all data within one single window for reasons of  
422 efficiency, but inevitably leads to a weighed appreciation where data from different channels do get, to  
423 some extent, a degree of relative scoring. This mainly applies to the scores good (4)-excellent (5) where  
424 recordings of certain channels could have likely received different rankings if they had been individually  
425 assessed without the knowledge of the signal on the other channels. This remark may also pertain to the  
426 grade borderline. As future use of the device will target acquiring the best possible signals in a given subject,  
427 we particularly focused on signals graded 4 or 5. Figure 9 provides a visual overview of observed quality  
428 across the complete database. Each handpiece of the device is equipped with 6 channels in line, spanning  
429 2.5 cm with the aim to have minimally one channel that detects a strong signal. It is clear that channel 1  
430 on handpiece 1 systematically yields very low scores, which was attributed to a hardware problem with  
431 inadequate alignment of the optical components during device assembly. For carotid measurements, the  
432 middle channels 3 and 4 yielded the the highest quality signals (as expected), but this shifted to channels 2  
433 and 3 for handpiece 2. Also, the overall signal quality was slightly higher for handpiece 2. We speculate that  
434 the use of the spacer underneath handpiece 1 may contribute to the difference in signal quality between both

435 handpieces. These data can be compared to the data from handpiece 2 during carotid-femoral measurement,  
436 where handpiece 2 is now equipped with a spacer (see Figure 9 for the measuring configurations). The  
437 mean signal quality is now in the same range as it was for handpiece 1 on the local carotid measurements.  
438 An extra factor, however, is the fact that carotid-femoral measurements are technically more demanding,  
439 requiring the simultaneous acquisition of signals at 2 distinct locations. The same conclusions can be drawn  
440 on the basis of the automatically calculated scores  $Q_{TM}$  and  $Q_{MP}$ .

441 In essence, one very good to excellent channel recording on each of the handpieces should guarantee a  
442 reliable transit time estimation from one handpiece to the other. This was achieved in 27% of the local  
443 carotid datasets and in 13% of the carotid-femoral datasets mainly due to the suboptimal femoral recording  
444 that is more challenging due to fact that the operators has to manipulate two sensors on two distinct  
445 locations as well as the deeper positioning of the femoral artery leading to weaker signals. That does not  
446 imply that the remaining datasets cannot be processed (especially when the ECG is available; see (Marais  
447 et al., 2019)) or that LDV would not be suitable as measuring technique; we just speculate that these results  
448 can be drastically improved with real-time feedback on the signal quality upon measurement.

449 The main objective of this paper was to explore different methods for an automated signal quality  
450 assessment, where we first explored template matching. The template should minimally contain the foot  
451 fingerprint of the wave, apparent on both the carotid and femoral measuring locations. That pattern turned  
452 out to be fairly robust across the tested population. Even though the amplitude of acceleration signals was  
453 lower at the femoral measuring site, the pattern of the foot is quite similar on both measuring locations. A  
454 practical choice that has to be made is on the length of the template. For carotid signals, it may be relevant  
455 to extend the template such that it also encompasses the dicrotic notch. We preferred the shorter template  
456 of 200 ms (which does not extend beyond the dicrotic notch) as the time delay between the wave's foot and  
457 the dicrotic notch is not constant but varies in between subjects and also within one subject from cycle to  
458 cycle due to physiological variations in blood pressure and heart rate. The shorter template was found to  
459 result in a somewhat higher specificity in correctly classifying poor signals, but overall, the performance of  
460 the carotid templates with different lengths was not very different, as can be observed from the confusion  
461 matrix (Table 2). On the other hand, for the femoral artery, we preferred the longest template of 500 ms  
462 which should detect epochs characterized by one prominent peak, the foot of the wave, followed by a long  
463 tail of low amplitude signals.

464 We then determined optimal thresholds levels for the magnitude of the cross-correlation and the number  
465 of detected beats using ROC analysis, whereby we maximized the classification performance of a binary  
466 classifier on the basis of  $Q_{TM}$ . In this exploratory study, that analysis was done on the complete database  
467 and further optimizations should be done on the used features and repeating the analysis with a separate  
468 training and testing data set. Using the resulting thresholds, the agreement between  $Q_{TM}$  and  $Q_{vis}$  was  
469 overall satisfactory. The logistic regression model analyses learned that a template matching approach is a  
470 valuable option to automatically classify signal quality as acceptable or not acceptable with an accuracy of  
471 over 80%.

472 As a second method, we considered the matrix profile as a technique to identify recurring patterns in the  
473 LDV-measurements in an automatic manner (Zimmerman et al., 2019), with very few control parameters.  
474 The potential advantage of a matrix profile approach over template matching is that no prior knowledge is  
475 required on the shape of the signal feature that one is looking for. Also, using the matrix profile allows  
476 the generation of a 'user-dependent' template in situ. Signal quality was quantifiable using features of  
477 the motifs found by the matrix profile and combined into the quality metric  $Q_{TM}$ , which showed a good  
478 agreement with the ground truth of visual scores as can be observed from Figure 10.

479 As for the template matching approach, the average accuracy of a logistic regression model trained and  
480 tested on features derived from motifs provided by the matrix profile technique is in all cases higher than  
481 80%. Overall, only relatively small differences are observed between the two techniques. Both techniques  
482 perform similarly well which suggests that both, or a combination of the two, can be used for classifying  
483 new, future data into 'bad, unusable' or 'good, usable'. This allows us to state that a logistic regression  
484 model suffices, along with the signal features and techniques that are considered, to accurately assess  
485 incoming data in future real-time applications.

486 In our logistic model training, we purposely discarded datasets visually labelled 'borderline' (score 3) as  
487 these data were simply hard to classify visually in an unequivocal way. That difficulty is relatively well  
488 reflected in the values of the quantitative metrics for these signals (figure 10) and the performance of the

489 classifiers as quantified by the confusion matrix (table 2 and 3). Especially for the carotid artery, automated  
490 classification leads to a close to fifty-fifty percent labeling of data as acceptable or not acceptable. For the  
491 femoral recordings, there is a larger tendency to classify signals with visual score 3 as not acceptable. This  
492 is in line with our own perception that femoral data may have received higher scores than carotid data of a  
493 similar quality, and underlines the need for objective tools to score signal quality.

494 Interestingly, the quality score, exemplified by  $Q_{MP}$ , correlates negatively with age and especially  
495 with BMI when signals are measured in the groin on the femoral artery. This observation supports the  
496 operator impression that measuring good quality LDV-signals on more obese subjects is consistently more  
497 challenging. The deeper the positioning of the artery and the more surrounding tissue, the stronger the  
498 signal attenuation. Such relation with BMI was absent for neck recordings. Also, skin inelasticity or  
499 thickness is expected to play a role on the transmission of intra-arterial vibrations and likely contributed to  
500 the observed negative correlation between and signal quality at the carotid and femoral locations in the  
501 study populations. The negative correlations between signal quality and age for carotid-carotid recordings  
502 with handpiece 2 were less strong, and were not found for the other carotid recordings (carotid-carotid  
503 handpiece 1 or carotid-femoral handpiece 2 recordings). A possible explanation may be the use of the  
504 spacer for these latter measurements, which may mechanically interfere with the transmission of the  
505 vibrations from within the artery to the skin and exert an effect on the recordings. Overall, this effect is  
506 considered minor, but it may nonetheless be a factor contributing to observed differences in the recordings.

507 The CARDIS prototype has a laser wavelength of 1550 nanometer which is insufficiently reflected  
508 by the skin. We therefore attached retroreflective patches to the skin at the measurement locations to  
509 enhance reflection. The next-generation prototype aims for measurements without the retroreflective patch  
510 to facilitate practical use. A wavelength of 1300 nanometer, for which there is a relative peak in skin  
511 reflectance (Rockwell and Goldman, 1974), will be used but the impact of skin pigmentation or sweating  
512 on data quality will have to be investigated.

513 In this study, signal quality was assessed off-line on 20 second recordings. Future developments will focus  
514 on real-time assessment of data quality as data is being captured and where the considered techniques will  
515 be used for epoch detection and subsequent quality quantification. Although a template-matching approach  
516 has the benefit that prior knowledge can be used to assess incoming data from the start, we assume that  
517 both techniques provide similarly useful features and that both are suitable for real-time implementation. It  
518 may be an option to hybridize the two techniques to come to a stronger, even more robust algorithm when  
519 implementing them into the device.

## 5 CONCLUSION

520 In conclusion, template matching and matrix profiling are methods suitable for the automated assessment  
521 of the signal quality of acceleration data measured from the skin in the neck and groin using laser Doppler  
522 velocimetry. Both methods allow to identify epochs in a data stream, and provide quantifiable features that  
523 can be combined into a quality score, or be used as input for logistic regression models for an automated  
524 classification of signals as acceptable or not acceptable. Models based on both methods yielded an accuracy  
525 of minimally 80% in our CARDIS database of carotid and femoral recordings, reaching as high as 87% for  
526 the femoral data.

## REFERENCES

- 527 Austin, P. C. and Steyerberg, E. W. (2012). Interpreting the concordance statistic of a logistic regression  
528 model: Relation to the variance and odds ratio of a continuous explanatory variable. *BMC Medical*  
529 *Research Methodology* 12, 1–8. doi:10.1186/1471-2288-12-82
- 530 Bramwell, J. C. and Hill, A. V. (1922). The velocity of pulse wave in man. *Proceedings of the Royal*  
531 *Society of London. Series B, Containing Papers of a Biological Character* 93, 298–306
- 532 Campo, A. and Dirckx, J. (2011). Dual-beam laser doppler vibrometer for measurement of pulse wave  
533 velocity in elastic vessels. In *22nd Congress of the International Commission for Optics: Light for the*  
534 *Development of the World, Puebla, Mexico* (International Society for Optics and Photonics), vol. 8011,  
535 80118Y
- 536 Chirinos, J. A., Segers, P., Hughes, T., and Townsend, R. (2019). Large-artery stiffness in health and  
537 disease: Jacc state-of-the-art review. *Journal of the American College of Cardiology* 74, 1237–1263

- 538 De Melis, M., Morbiducci, U., Scalise, L., Tomasini, E. P., Delbeke, D., Baets, R., et al. (2008). A  
539 noncontact approach for the evaluation of large artery stiffness: a preliminary study. *American journal*  
540 *of hypertension* 21, 1280–1283
- 541 Domínguez-Almendros, S., Benítez-Parejo, N., and Gonzalez-Ramirez, A. R. (2011). Logistic regression  
542 models. *Allergologia et Immunopathologia* 39, 295–305. doi:10.1016/j.aller.2011.05.002
- 543 Jensen-Urstad, K., Storck, N., Bouvier, F., Ericson, M., and Lindblad, L. E. (1997). Heart rate variability  
544 in healthy subjects is related to age and gender, 235–241
- 545 Jiun-Hung, C., Chu-Song, C., and Yong-Sheng, C. (2003). Fast Algorithm for Robust Template Matching  
546 With M-Estimators. *IEEE TRANSACTIONS ON SIGNAL PROCESSING* 51
- 547 Kaplan, A. D., OrSullivan, J. A., Sirevaag, E. J., Lai, P.-H., and Rohrbaugh, J. W. (2012). Hidden state  
548 models for noncontact measurements of the carotid pulse using a laser doppler vibrometer. *IEEE*  
549 *Transactions on Biomedical Engineering* 59, 744–753. doi:10.1109/tbme.2011.2179297
- 550 Laurent, S., Cockcroft, J., Van Bortel, L., Boutouyrie, P., Giannattasio, C., Hayoz, D., et al. (2006). Expert  
551 consensus document on arterial stiffness: methodological issues and clinical applications. *European*  
552 *heart journal* 27, 2588–2605
- 553 Li, Y., Marais, L., Khettab, H., Quan, Z., Aasmul, S., Leinders, R., et al. (2020). Silicon photonics-based  
554 laser doppler vibrometer array for carotid-femoral pulse wave velocity (pwv) measurement. *Biomedical*  
555 *Optics Express* 11, 3913–3926
- 556 Li, Y., Segers, P., Dirckx, J., and Baets, R. (2013). On-chip laser doppler vibrometer for arterial pulse wave  
557 velocity measurement. *Biomedical optics express* 4, 1229–1235
- 558 Madrid, F., Imani, S., Mercer, R., Zimmerman, Z., Shakibay, N., and Keogh, E. (2019). Matrix profile XX:  
559 Finding and visualizing time series motifs of all lengths using the matrix profile. *Proceedings - 10th*  
560 *IEEE International Conference on Big Knowledge, ICBK 2019*, 175–182doi:10.1109/ICBK.2019.00031
- 561 Marais, L., Khettab, H., Li, Y., Segers, P., Baets, R., Reesink, K., et al. (2019). Measurement of aortic  
562 stiffness by laser doppler vibrometry: The cardis study. *Journal of Hypertension* 37, e88
- 563 Mitchell, G. F., Parise, H., Benjamin, E. J., Larson, M. G., Keyes, M. J., Vita, J. A., et al. (2004). Changes  
564 in arterial stiffness and wave reflection with advancing age in healthy men and women: The Framingham  
565 Heart Study. *Hypertension* 43, 1239–1245. doi:10.1161/01.HYP.0000128420.01881.aa
- 566 Mitchell, G. F., van Buchem, M. A., Sigurdsson, S., Gotal, J. D., Jonsdottir, M. K., Kjartansson, Ó., et al.  
567 (2011). engArterial stiffness, pressure and flow pulsatility and brain structure and function: the Age,  
568 Gene/Environment Susceptibility–Reykjavik study. *Brain : a journal of neurology* 134, 3398–3407.  
569 doi:10.1093/brain/awr253
- 570 Morbiducci, U., Scalise, L., De Melis, M., and Grigioni, M. (2007a). Optical vibrocardiography: A novel  
571 tool for the optical monitoring of cardiac activity. *Annals of biomedical engineering* 35, 45–58
- 572 Morbiducci, U., Scalise, L., Melis, M. D., and Grigioni, M. (2007b). Optical vibrocardiography: A  
573 novel tool for the optical monitoring of cardiac activity. *Annals of Biomedical Engineering* 35, 45–58.  
574 doi:10.1007/s10439-006-9202-9
- 575 Mueen, A., Keogh, E., Zhu, Q., Cash, S., and Westover, B. (2009). Exact discovery of time series motifs.  
576 *Society for Industrial and Applied Mathematics - 9th SIAM International Conference on Data Mining*  
577 *2009, Proceedings in Applied Mathematics* 1, 469–480. doi:10.1137/1.9781611972795.41
- 578 Nick, T. and Campbell, K. (2012). Logistic regression. *Discovering Statistics Using SPSS* 404, 731–735.  
579 doi:\textcolor{blue}{10.1007/978-1-59745-530-5\_14}
- 580 Omachi, S. and Omachi, M. (2007). Fast Template Matching With Polynomials. *IEEE TRANSACTIONS*  
581 *ON IMAGE PROCESSING* 16
- 582 O'Rourke, M. and Kelly, R. (1993). Wave reflection in the systemic circulation and its implications in  
583 ventricular-function. *Journal of Hypertension* 11, 327–337. doi:{10.1097/00004872-199304000-00001}
- 584 Pereira, T., Correia, C., and Cardoso, J. (2015). Novel methods for pulse wave velocity measurement.  
585 *Journal of medical and biological engineering* 35, 555–565
- 586 Pinotti, M., Paone, N., Santos, F. A., and Tomasini, E. P. (1998). Carotid artery pulse wave measured by a  
587 laser vibrometer. In *Third International Conference on Vibration Measurements by Laser Techniques:*  
588 *Advances and Applications* (International Society for Optics and Photonics), vol. 3411, 611–616
- 589 Rockwell, R. and Goldman, L. (1974). engResearch on human skin laser damage thresholds. *Report*  
590 *DERM-LL-74-1003, USAF School of Aerospace Medicine, Brooks Air Force Base TX*
- 591 Rohrbaugh, J. W., Sirevaag, E. J., and Richter, E. J. (2013). Laser doppler vibrometry measurement of the  
592 mechanical myogram. *Review of Scientific Instruments* 84, 121706. doi:10.1063/1.4845435

- 593 Scalise, L. and Morbiducci, U. (2008). Non-contact cardiac monitoring from carotid artery using optical  
594 vibrocardiography. *Medical Engineering and Physics* 30, 490–497. doi:10.1016/j.medengphy.2007.05.  
595 008
- 596 Segers, P., Rietzschel, E. R., and Chirinos, J. A. (2020). How to measure arterial stiffness in humans.  
597 *Arteriosclerosis, thrombosis, and vascular biology* 40, 1034–1043
- 598 Vlachopoulos, C., Aznaouridis, K., and Stefanadis, C. (2010). Prediction of cardiovascular events and  
599 all-cause mortality with arterial stiffness: a systematic review and meta-analysis. *Journal of the American*  
600 *College of Cardiology* 55, 1318–1327
- 601 Westerhof, N., Lankhaar, J.-W., and Westerhof, B. E. (2009). The arterial windkessel. *Medical & biological*  
602 *engineering & computing* 47, 131–141
- 603 Wolinsky, H. and Glagov, S. (1967). A lamellar unit of aortic medial structure and function in mammals.  
604 *Circulation research* 20, 99–111
- 605 Won-Du, C. and Chang-Hwan, I. (2014). Enhanced Template Matching Using Dynamic Positional Warping  
606 for Identification of Specific Patterns in Electroencephalogram. *Journal of Applied Mathematics* 2014
- 607 Yeh, C.-C. M., Zhu, Y., Ulanova, L., Begum, N., Ding, Y., Dau, H. A., et al. (2018). Time series joins,  
608 motifs, discords and shapelets: a unifying view that exploits the matrix profile. *Data Mining and*  
609 *Knowledge Discovery* 32, 83–123. doi:10.1007/s10618-017-0519-9
- 610 Zhang, J. (2007). EFFECT OF AGE AND SEX ON HEART RATE VARIABILITY IN HEALTHY  
611 SUBJECTS , 374–379doi:10.1016/j.jmpt.2007.04.001
- 612 Zhu, Y., Gharghabi, S., Silva, D. F., Dau, H. A., Yeh, C.-C. M., Shakibay Senobari, N., et al. (2020). The  
613 Swiss army knife of time series data mining: ten useful things you can do with the matrix profile and ten  
614 lines of code. *Data Mining and Knowledge Discovery* 34, 949–979. doi:10.1007/s10618-019-00668-6
- 615 Zhu, Y., Imamura, M., Nikovski, D., and Keogh, E. (2017a). Matrix Profile VII: Time Series Chains: A  
616 New Primitive for Time Series Data Mining. *Ieee Icdm*
- 617 Zhu, Y., Zimmerman, Z., Senobari, N. S., Yeh, C. C. M., Funning, G., Mueen, A., et al. (2017b).  
618 Matrix profile II: Exploiting a novel algorithm and GPUs to break the one hundred million barrier for  
619 time series motifs and joins. *Proceedings - IEEE International Conference on Data Mining, ICDM* ,  
620 739–748doi:10.1109/ICDM.2016.126
- 621 Zimmerman, Z., Shakibay Senobari, N., Funning, G., Papalexakis, E., Oymak, S., Brisk, P., et al. (2019).  
622 Matrix profile XVIII: Time series mining in the face of fast moving streams using a learned approximate  
623 matrix profile. *Proceedings - IEEE International Conference on Data Mining, ICDM 2019-November*,  
624 936–945. doi:10.1109/ICDM.2019.00104

## CONFLICT OF INTEREST STATEMENT

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

## AUTHOR CONTRIBUTIONS

627 The Author Contributions section is mandatory for all articles, including articles by sole authors. If an  
628 appropriate statement is not provided on submission, a standard one will be inserted during the production  
629 process. The Author Contributions statement must describe the contributions of individual authors referred  
630 to by their initials and, in doing so, all authors agree to be accountable for the content of the work. Please  
631 see here for full authorship criteria.

## FUNDING

632 The research was funded by the European Union’s Horizon 2020 Research and Innovation Programme  
633 under grant agreement No 644798 (CARDIS) and 871547 (INSIDE)

## ACKNOWLEDGMENTS

634 We acknowledge the key contributions of Louise Marais, Hakim Khettab and Michael Vanslembrouck in  
635 setting up the CARDIS data and acquiring the data.

## TABLES

Quality score $Q_{vis}$	Quality	Description
Score 1	Bad	Acquisition with no evidence of repeatable features that may be linked to the detection of a pulse
Score 2	Poor	Very noisy acquisition not suitable for analysis, but with identifiable pulses within the noisy trace
Score 3	Bordeline	Acquisition affected by noise but presenting clear repeatable patterns. Advanced signal processing algorithms could remove the noise and allow to detect the foot of the pulse wave with reasonable affordability
Score 4	Good	Acquisition with sharp and pronounced peaks at the foot (and dicrotic notch), with relatively low noise levels between successive pulse peaks
Score 5	Excellent	Acquisition with very sharp and pronounced peaks at the foot (and dicrotic notch), with low noise levels in between the peaks. Signals of textbook quality

**Table 1.** The 5-levels grade scale taking values  $Q_{vis}$

## TABLES

	Template of 200ms		Template of 400ms		Template of 600ms	
Carotid recordings	TM score 0	TM score 1	TM score 0	TM score 1	TM score 0	TM score 1
score 1	97%	3%	97%	3%	97%	3%
score 2	86%	14%	86%	14%	82%	18%
score 3	58%	42%	55%	45%	44%	56%
score 4	29%	71%	30%	70%	15%	85%
score 5	8%	92%	19%	81%	6%	94%

**Table 2.** Confusion matrices of signal classification done by the hand-engineered classification model constructed with template matching. Signals classified in this table were measured at the carotid and the templates used were the carotid population templates.

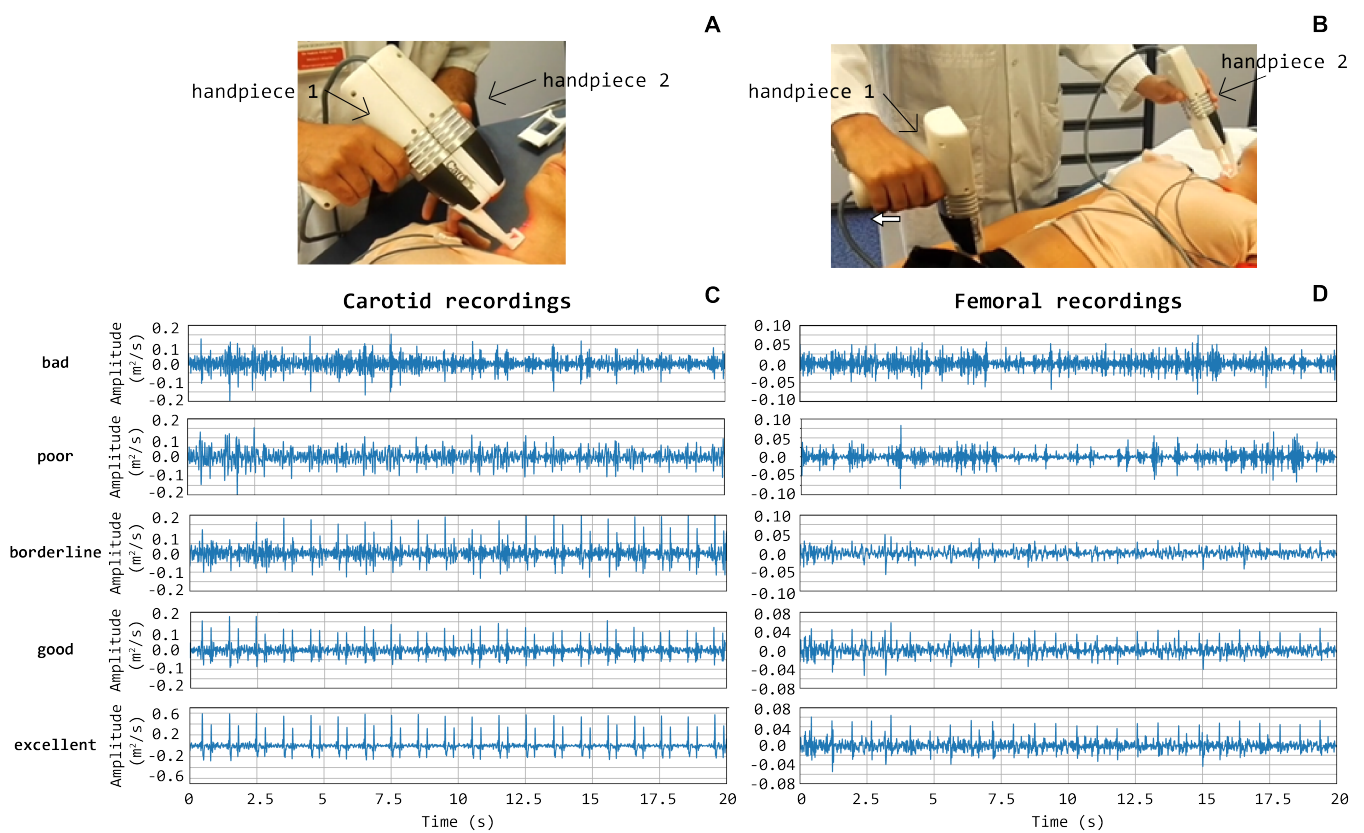
	Template of 300ms		Template of 400ms		Template of 500ms	
Femoral recordings	TM score 0	TM score 1	TM score 0	TM score 1	TM score 0	TM score 1
score 1	96%	4%	96%	4%	96%	4%
score 2	91%	9%	89%	11%	91%	9%
score 3	68%	32%	64%	36%	65%	35%
score 4	33%	67%	28%	72%	27%	73%
score 5	8%	92%	6%	94%	7%	93%

**Table 3.** Confusion matrices of signal classification done by the hand-engineered classification model constructed with template matching. Signals classified in this table were measured at the femoral and the templates used were the femoral population templates.

Accuracy	Template Matching		Matrix Profile	
	Average	Std	Average	Std
Carotid-carotid hp1	80%	1.75%	82%	1.64%
Carotid-carotid hp2	85%	1.63%	88%	1.53%
Femoral-carotid hp1	87%	1.31%	86%	1.43%
Femoral-carotid hp2	81%	1.96%	85%	1.71%

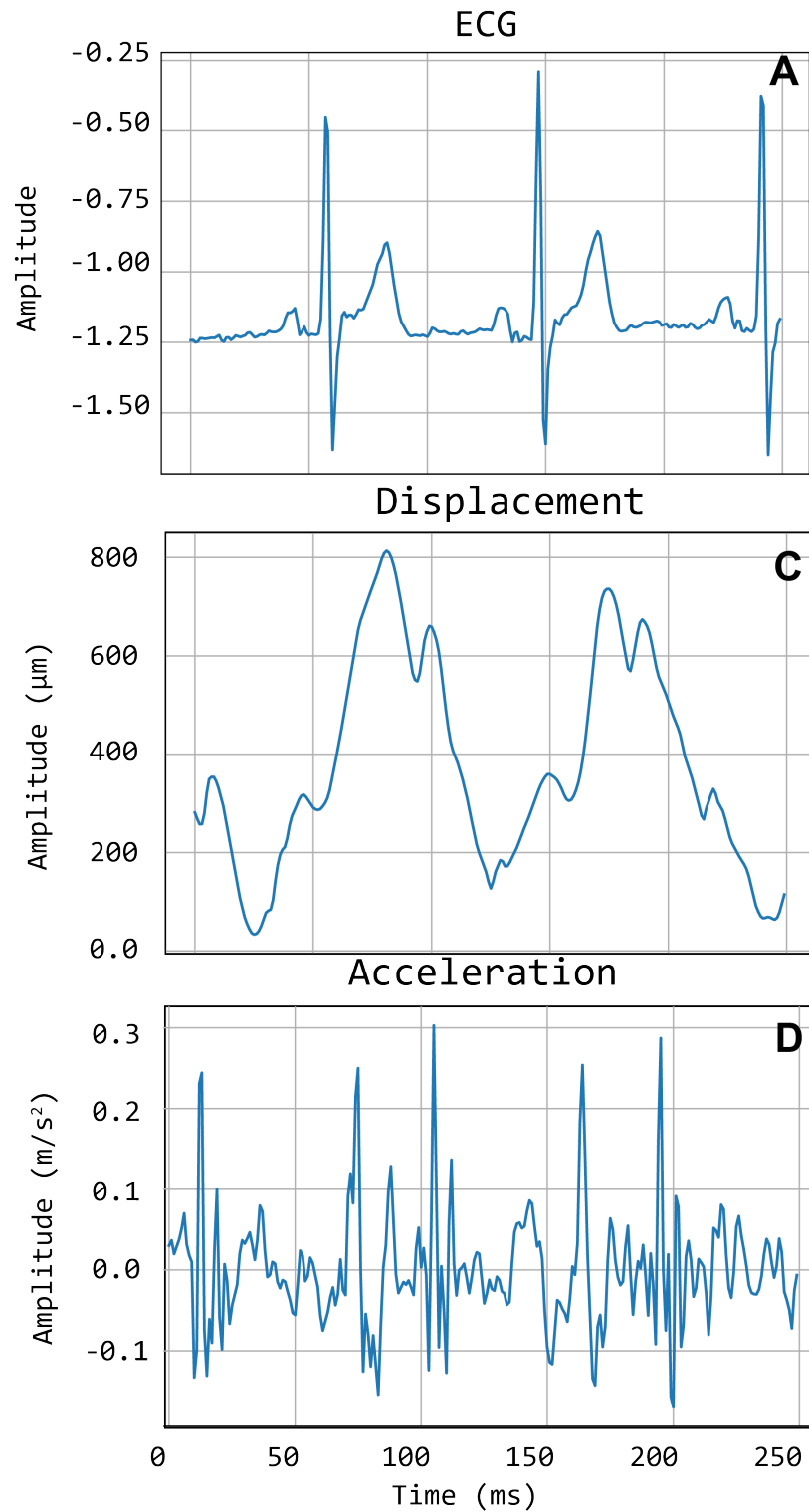
**Table 4.** Table containing the average performance of the logistic regression models trained on features derived by both template matching and matrix profile methods. Results are shown per handpiece of the measuring device. Average classification accuracy as well as its standard deviation are given.

## FIGURE CAPTIONS

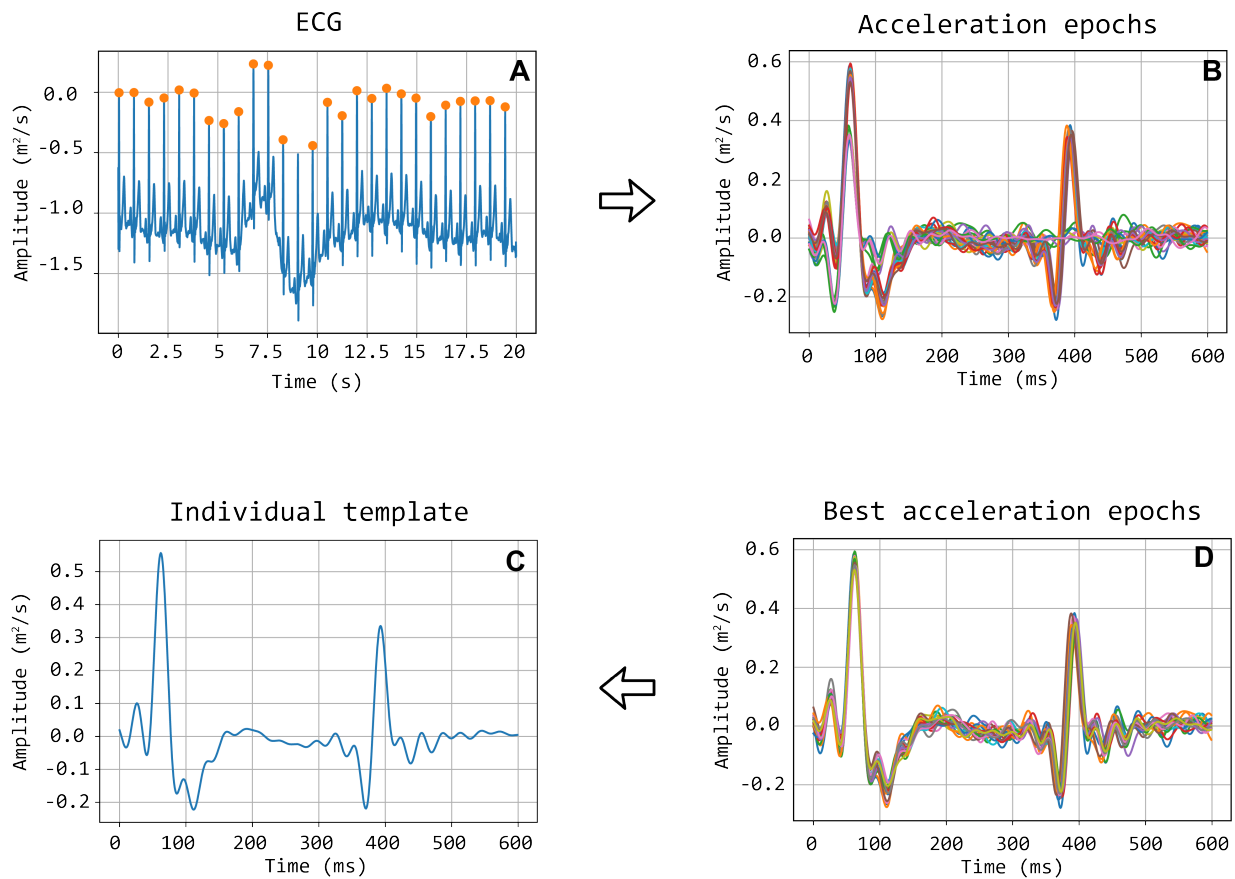


**Figure 1.** CARDIS device in configuration to measure carotid-femoral (A) and local carotid PWV (B). (C) and (D) display representative tracings on the carotid (C) and femoral (D) measuring site receiving a visual grading score of 1 to 5. Especially in the excellent tracings, the foot-of-the-wave waveforms are clearly visible, with the same for the dicotic notch waveforms in the local carotid case.

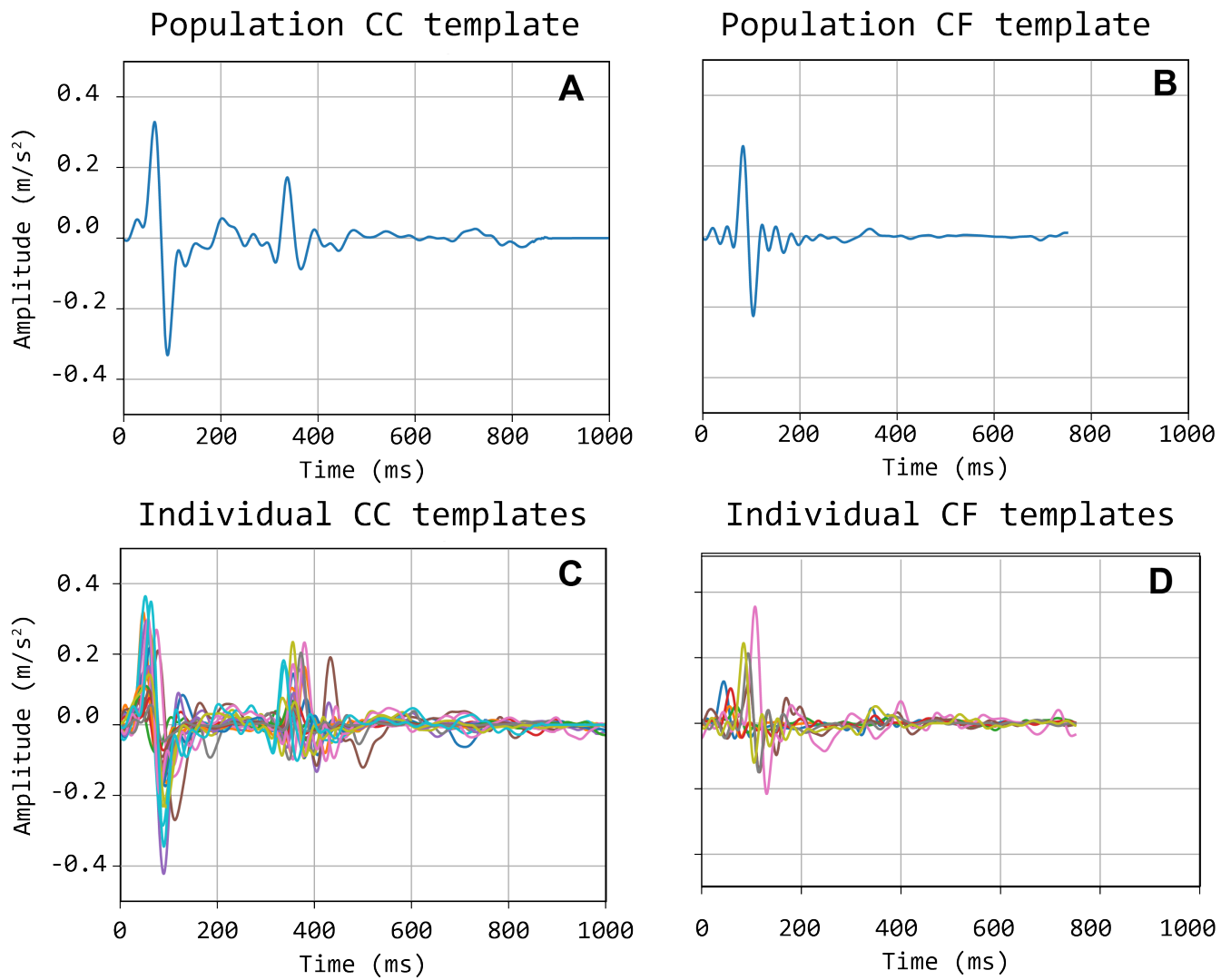




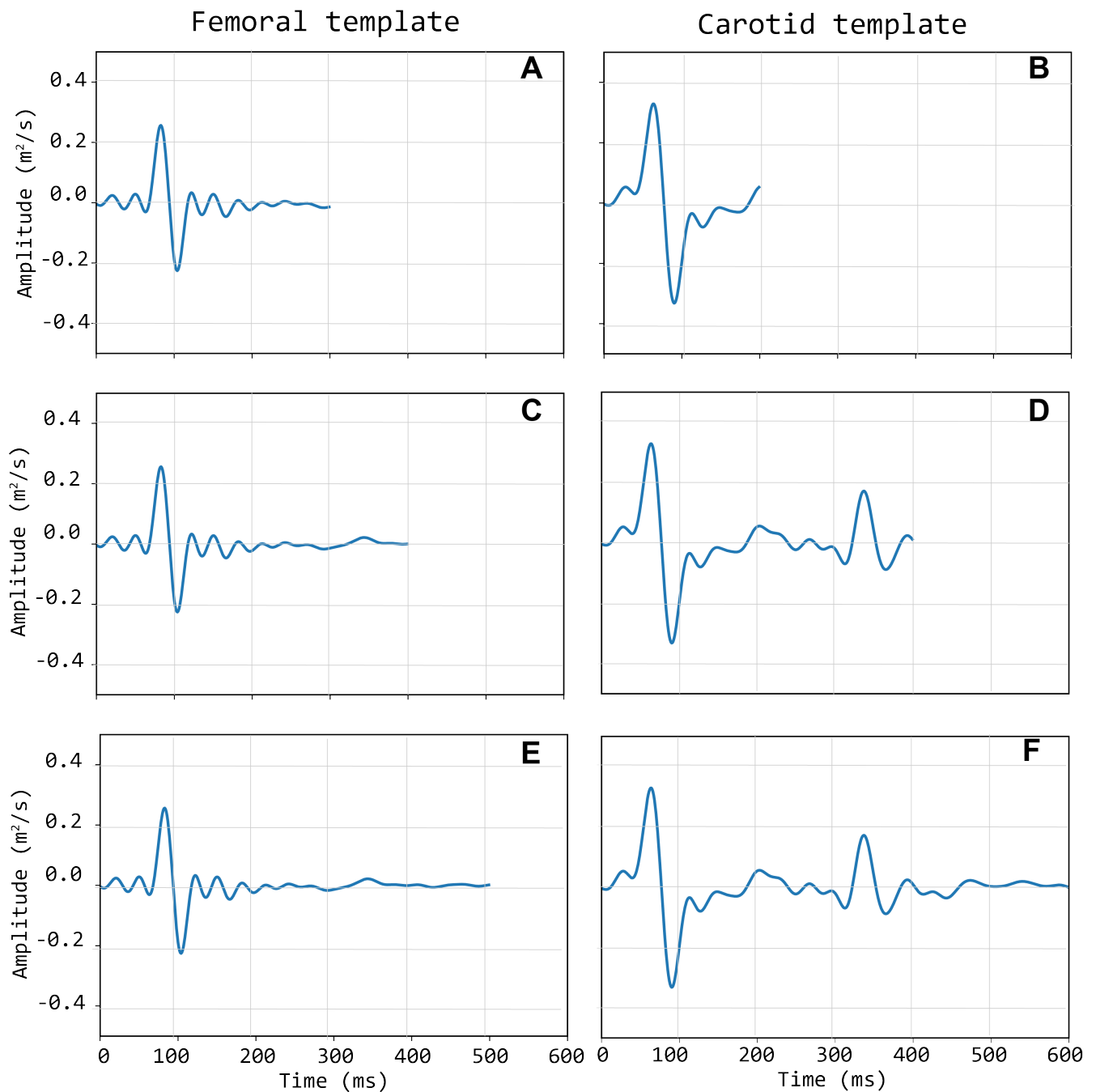
**Figure 2.** An example of Cardis data. (A) shows the ECG signals, (B) shows the displacement signal and (C) the corresponding acceleration signal.



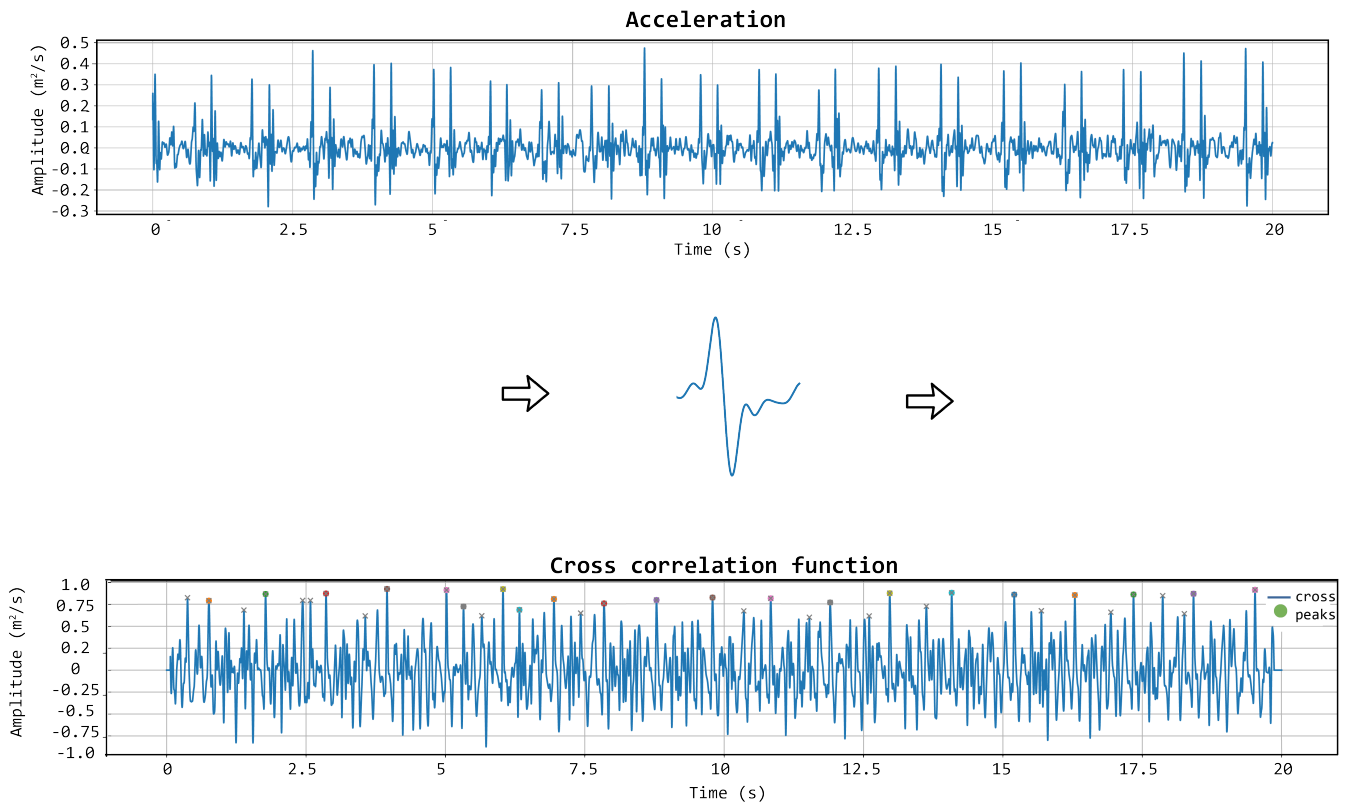
**Figure 3.** Workflow detailing the construction of the template. Illustration of selection of good quality epochs using the correlation coefficient. (A) ECG signal with detected R peaks, which are used to segment the acceleration signal into heartbeat epochs (B). After the correlation matrix analysis, only the good epochs are maintained (D). In (C) the final individual template, calculated as the average of the good epochs, is displayed.



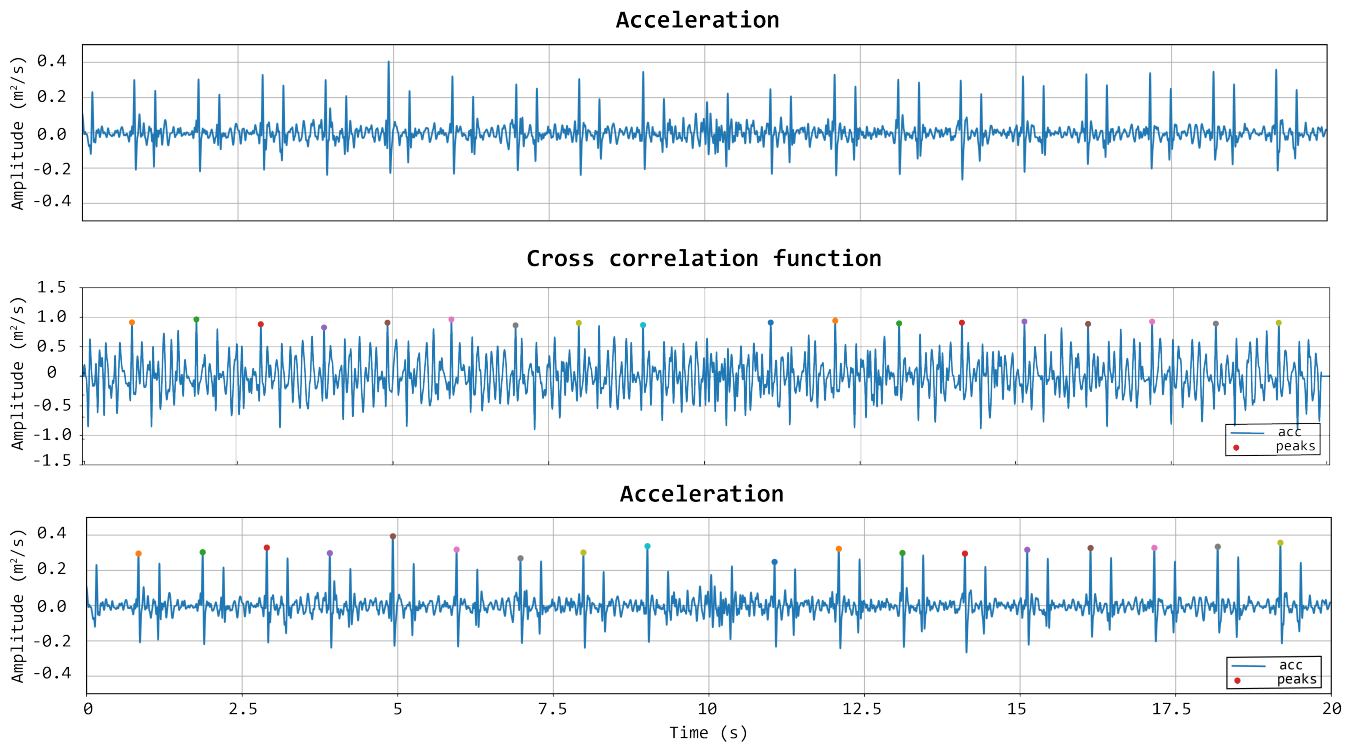
**Figure 4.** Bottom (C & D): individual carotid (CC) and femoral (CF) templates; Top (A & B): population-average carotid and femoral templates.



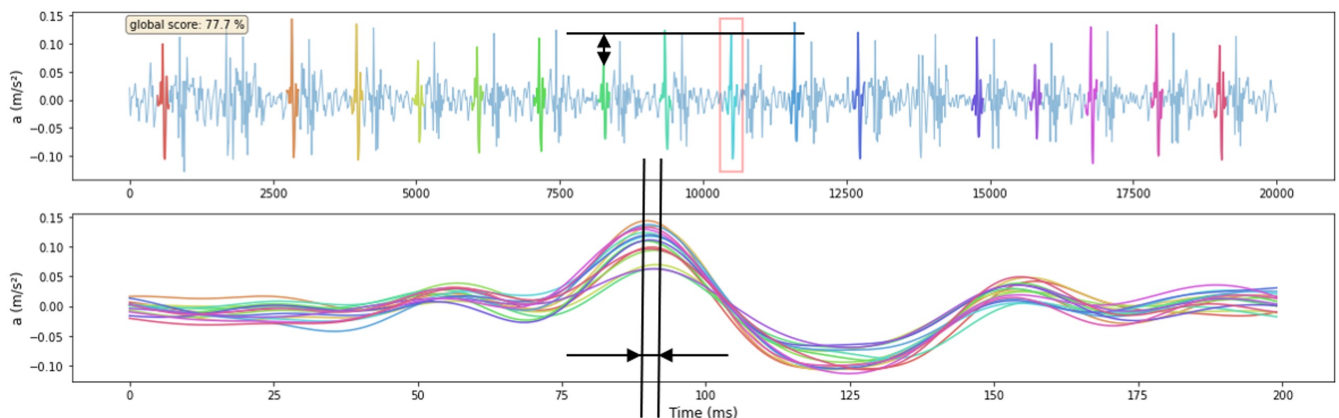
**Figure 5.** Left column (A, C & E): three different length of the femoral template; Right column (B, D & F): three different lengths of the carotid template.



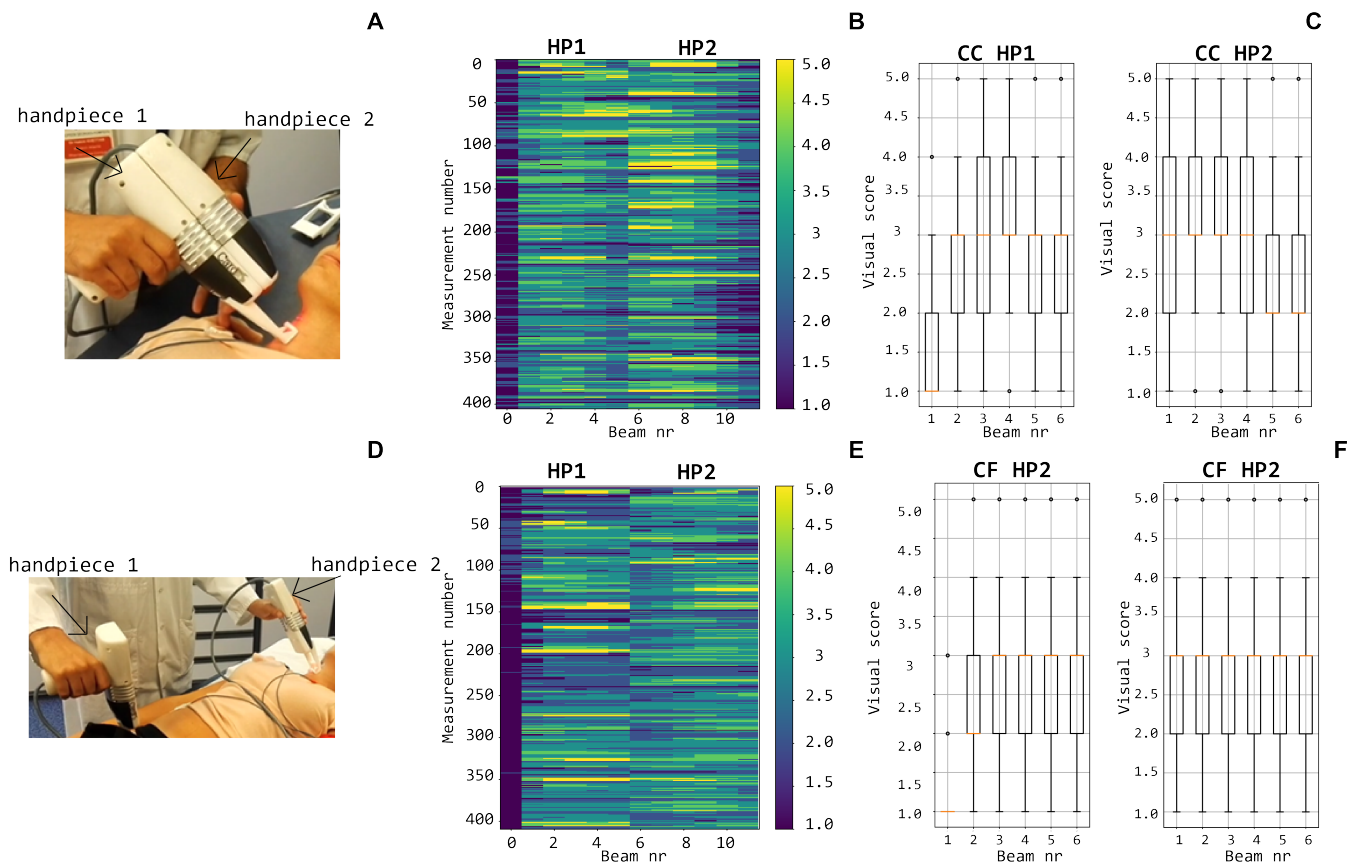
**Figure 6.** The template matching algorithm is shown. The chosen template (displayed in the middle row of the figure) is iteratively correlated with the acceleration signal to get the cross correlation function. In that function, the appropriate peaks are then identified.



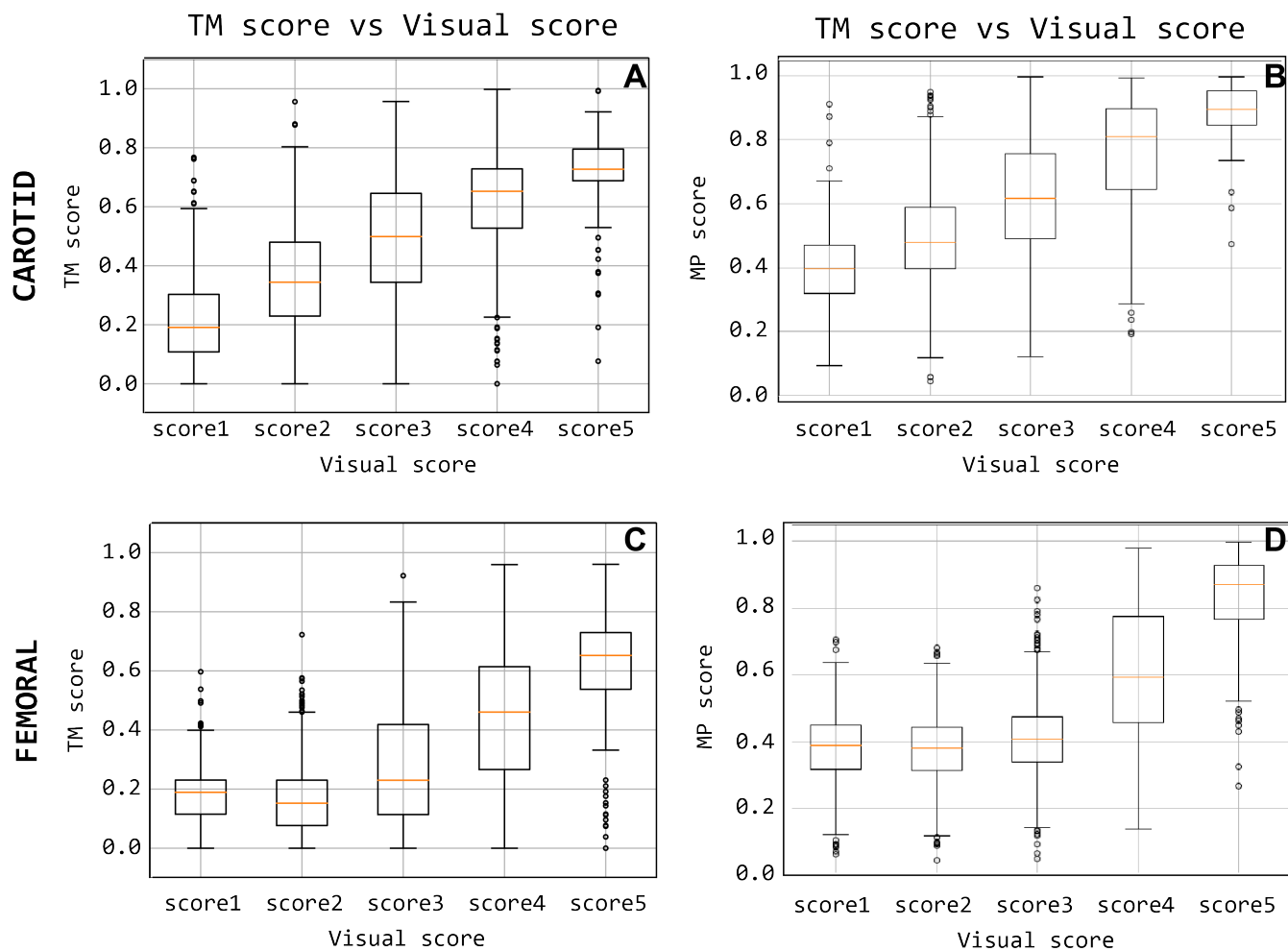
**Figure 7.** First row: acceleration signal. Second row: normalised cross-correlation function and its maximum values of the acceleration signal with the femoral template of 500 ms. Last row: acceleration signal with the detected peaks using the template matching method.



**Figure 8.** Example of one signal being scored by the features that are derived from the Matrix-Profile-identified motif. The amplitude feature of one subsequence is shown in the upper figure, the reference is indicated with a red square. The time-instant of subsequence peak feature is shown in the lower figure where all subsequences shown in the upper figure are time-aligned. Signal score is shown in the upper left corner of the upper figure. The visual score of this signal is 4.

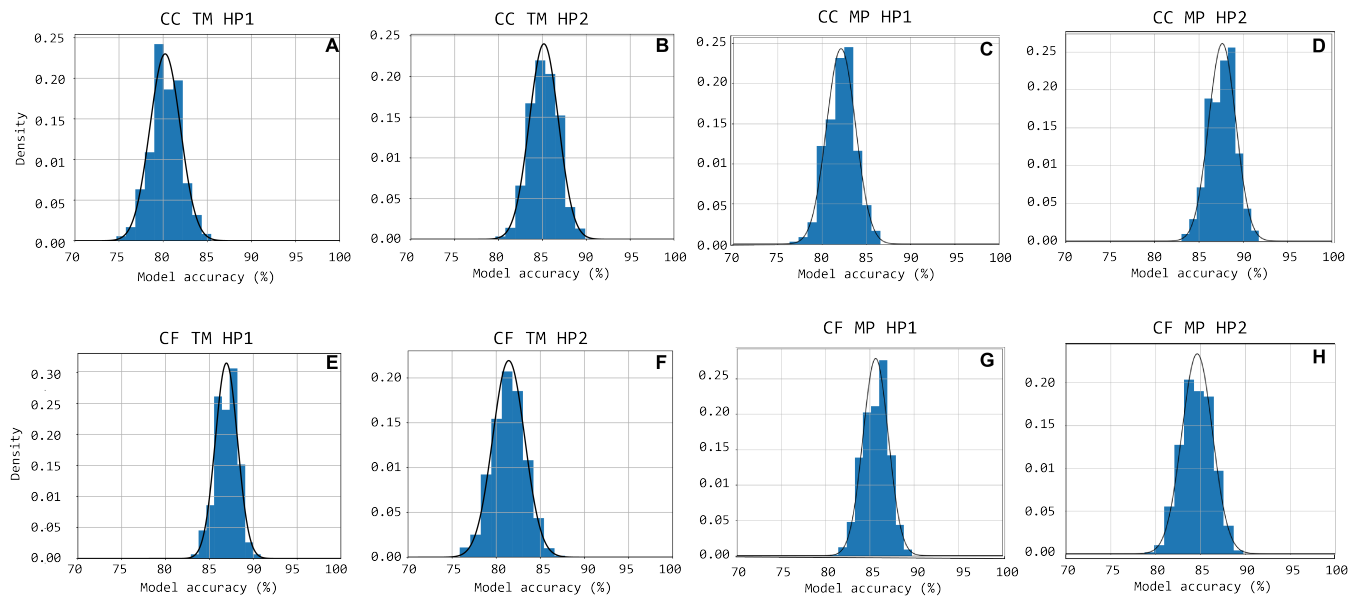


**Figure 9.** Top row: visual scoring of signals measured with handpiece 1 and 2 during local carotid measurements (A) with overall grades shown in (B) and box plots and mean values per channel in (C). Bottom row: visual scoring of signals measured during carotid-femoral PWV measurements (D) with overall grades shown in (E) and box plots and mean values per channel in (F).

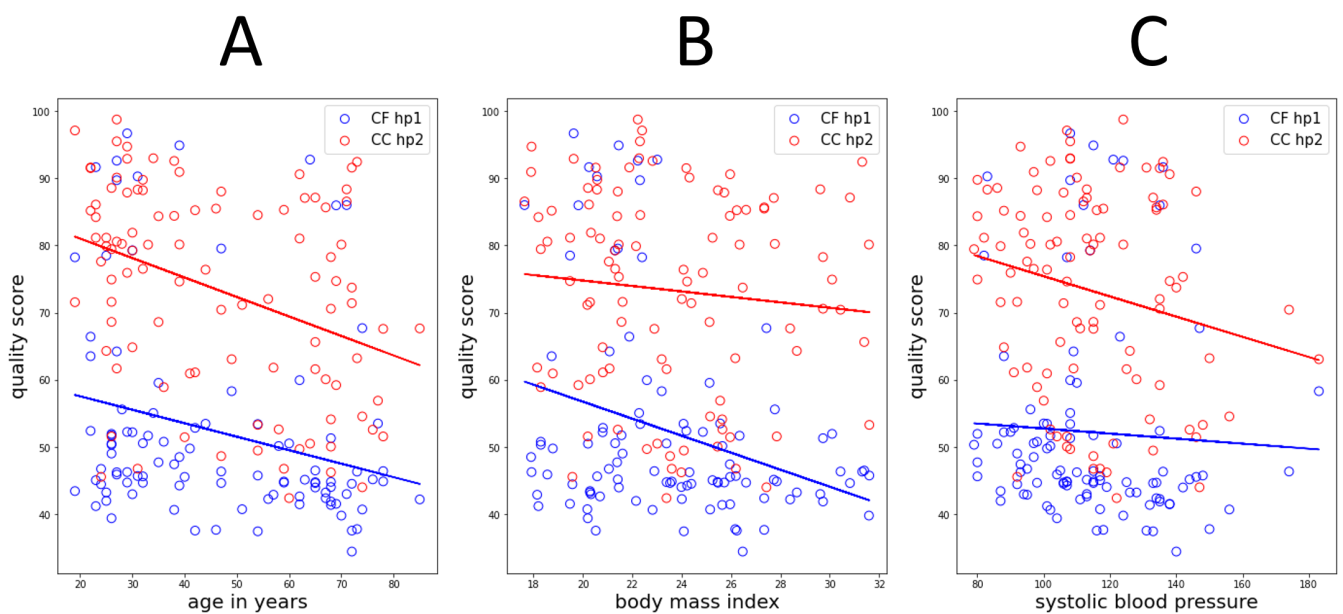


**Figure 10.** Quality score comparison between visual score and the template-matching-derived and matrix-profile-derived. Subfigures (A) and (C) display the score based on the template matching and (B) and (D) quality score based on the matrix profile.





**Figure 11.** Accuracy distributions for 1000 random train-test set partitions and subsequent logistic regression models trained. The accuracy distribution is shown cumulatively through bar-charts with the equivalent Gauss-curve plotted on top of it. Subfigures (A), (B), (C) and (D) show this for CC HP1, CC HP2, CF HP1 and CF HP2 cases respectively.



**Figure 12.** Correlation analyses of the matrix profile-based quality score with age, BMI and systolic blood pressure, shown in subfigures (A), (B) and (C) respectively. Only data in the CC HP2 and CF HP1 cases is shown. Trendlines of the data are also drawn.