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Performance analysis of four nonlinearity analysis methods using a model with variable complexity and application to uterine EMG signals

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Abstract

Several measures have been proposed to detect nonlinear characteristics in time series. Results on time series, multiple surrogates and their z-score are used to statistically test for the presence or absence of non-linearity. The z-score itself has sometimes been used as a measure of nonlinearity. The sensitivity of nonlinear methods to the nonlinearity level and their robustness to noise have rarely been evaluated in the past. While surrogates are important tools to rigorously detect nonlinearity, their usefulness for evaluating the level of nonlinearity is not clear. In this paper we investigate the performance of four methods arising from three families that are widely used in non-linearity detection: statistics (Time reversibility), predictability (Sample Entropy, Delay Vector Variance) and chaos theory (Lyapunov Exponents). We used sensitivity to increasing complexity and the Mean square Error (MSE) of Monte Carlo instances for quantitative comparison of their performances. These methods were applied to a Henon nonlinear synthetic model in which we can vary the complexity degree (*CD*). This was done first by applying the methods directly to the signal and then using the z-score (surrogates) with and without added noise. The methods were then applied to real uterine EMG signals and used to

distinguish between pregnancy and labor contraction bursts. The discrimination performances were compared to linear frequency based methods classically used for the same purpose such as Mean Power Frequency (MPF), Peak Frequency (PF) and Median Frequency (MF). The results show noticeable difference between different methods, with a clear superiority of some of the nonlinear methods (Time reversibility, Lyapunov exponents) over the linear methods. Applying the methods directly to the signals gave better results than using the z-score, except for Sample Entropy.

Keywords

Nonlinear time series analysis, uterine electromyogram, contraction discrimination, surrogates.

I. Introduction

One of the most common ways of obtaining information on neurophysiologic systems is to study the features of the signal(s) using time series analysis techniques. This traditionally rely on linear methods in both time and frequency domains [1]. Unfortunately, these methods cannot give information about purely nonlinear features of the signal. Due to the intrinsic nonlinearity of most biological systems, these nonlinear features may be present in physiological data and even be a characteristic of major interest. Recently, much attention has been paid to the use of nonlinear analysis techniques for the characterization of a biological signal [2]. Indeed, this type of analysis gives information about the nonlinear features of these signals, which arise from the underlying physiological processes, many of which have complex behavior. There is a growing literature reporting nonlinear analysis of various biosignal types (EEG [3], ECG [4], HRV [5] and EMG [6]).

The EHG or electrohysterogram (electrical uterine activity recorded on woman's abdomen) has been widely studied [7], [8], [9], [10], [11]. Nonlinear characteristics have been observed in the EHG and some success has been achieved by using these characteristics to obtain information of potential clinical usefulness. Radomski et al. show that nonlinear analysis of EHG based on the sample entropy statistic could differentiate dynamic states of uterine contractions [12]. A comparison between linear and nonlinear analysis with different conditions was done in [13]. It was concluded that median frequency is the best method among linear methods and that sample entropy is the best method among

nonlinear methods for term/preterm EHG contractions classification. Sample entropy is superior to median frequency, which indicates that nonlinear analysis is more suitable than linear analysis for studying EHG signals. In [14] the progress of labor was evaluated using sample entropy. Our team has examined nonlinear EHG analysis methods. Our results confirm the presence of nonlinearity in EHG signals. This character of the signals is useful in discriminating between pregnancy and labor contractions [15], [2], [16]. Practical disadvantages of the nonlinear analysis methods have been reported in [16]. They include excessive calculation time due to surrogates analysis and promising but inconclusive results due to the small amount of data that can practically be used due to heavy calculation times.

This paper presents work that extends previous work done in our group in comparing Approximate Entropy, Correntropy and Time reversibility [16]. In this work we implemented additional nonlinear analysis methods (delay vector variance, Lyapunov exponents) and new ways of testing them. We also used a larger database of real signals than in the previous work and we investigated the sensitivity of the methods to the varying complexity of signals and their robustness. The kind of sensitivity and robustness analyses of non-linearity measures presented in this paper, are rare or absent in the literature.

Four nonlinear methods: Time reversibility [17], Sample Entropy [18], Delay Vector Variance [19] and Lyapunov Exponents [20] were used in this work. Sensitivity of these methods to the complexity degree (CD) of a signal as well as robustness analysis were done on Henon model synthetic signals where CD can be controlled. The sensitivity to CD was first studied using the direct value provided by the method. It was then studied using surrogates and z-score, as the measure permitting evaluation of the nonlinearity. One objective of this study is to show which method(s) is most sensitive to the change of signal complexity. A second objective is to determine whether the use of surrogates gives better overall results than the direct application of the methods. This is of major practical importance for clinical application, as the generation of surrogates is very computationally expensive. The methods are also compared using the Mean square error (MSE) of the method results for 30 Monte Carlo instances of the signal. Finally, these non-linear methods are compared to three linear frequency

based characteristics of the signal, MPF, PF and MF, when applied to real EHG signals, in order to discriminate pregnancy and labor contractions.

II. Materials and Methods

A) Data

1. Synthetic signals

The Henon map is a well-known two-dimensional discrete-time system given by:

$$Y_{t+1} = c - Y_t^2 + CD * X_t,$$

$$X_{t+1} = Y_t,$$

where Y_t and X_t represent dynamical variables, CD is the complexity degree and c is the dissipation parameter. In this paper we use $c = 1$ as in [21] and $CD \in [0, 1]$ to change the model complexity [22] (Figure 1). The number of generated points is fixed to 1000. For the robustness analysis, we add to the synthetic signals a white Gaussian noise with the same duration, with a fixed 5db SNR with CD varying between 0 and 1 with a step 0.1. In the Monte Carlo analysis, we use 30 signals generated for each CD value.

2. Real signals

EHG signals were recorded from 38 subjects using a 4x4 electrode matrix located on the subject's abdomen (Figure 2), during one hour either at rest (woman lying on a bed) or during labor. One signal channel (bipolar vertical 7: BP7), located on the median vertical axis of the uterus was used for subsequent analysis (see [23] for details). After segmentation we obtained 115 labor bursts (recorded during delivery) and 174 pregnancy bursts (recorded more than 24 hours before delivery).

B) Non-linear Analysis Methods

1. Statistics family

98 a) Time reversibility

99 A time series is said to be reversible only if its probabilistic properties are invariant with respect to
100 time reversal. Time irreversibility can be taken as a strong signature of nonlinearity [17]. In this paper
101 we used the simplest method, described in [24] to compute the time reversibility of a signal S_n :

$$Tr(\tau) = \left(\frac{1}{N - \tau}\right) \sum_{n=\tau+1}^N (S_n - S_{n-\tau})^3$$

102 where N is the signal length and τ is the time delay.

103 2. Chaos theory family

104 a) Lyapunov Exponents

105 Lyapunov exponent (LE) is a quantitative indicator of system dynamics, which characterizes the
106 average convergence or divergence rate between adjacent tracks in phase space [20]. We used the
107 method described in [13] to compute LE:

$$\lambda = \lim_{t \rightarrow \infty} \lim_{\| \Delta_{y_0} \| \rightarrow 0} \left(\frac{1}{t}\right) \log(\| \Delta_{y_t} \| / \| \Delta_{y_0} \|),$$

108 Where $\| \Delta_{y_0} \|$ and $\| \Delta_{y_t} \|$ represent the Euclidean distance between two states of the system,
109 respectively to an arbitrary time t_0 and a later time t .

110 3. Predictability family

111 a) Sample Entropy

112 Sample Entropy (*SampEn*) is the negative natural logarithm of the conditional probability that a
113 dataset of length N , having repeated itself for m samples within a tolerance r , will also repeat itself for
114 $m+1$ samples. Thus, a lower value of *SampEn* indicates more regularity in the time series [18]. We
115 used the method described in [12] to compute *SampEn* :

For a time series of N points, $x_1, x_2, \dots, x_N$, we define subsequences, also called template vectors, of length m , given by: $y_i(m) = (x_i, x_{i+1}, \dots, x_{i+m-1})$ where $i = 1, 2, \dots, N-m+1$.

Then the following quantity is defined: $B_i^m(r)$ as $(N-m-1)^{-1}$ times the number of vectors X_j^m within r of X_i^m , where j ranges from 1 to $N-m$, and $j \neq i$, to exclude self-matches, and then define:

$$B^m(r) = \frac{1}{N-m} \sum_{i=1}^{N-m} B_i^m(r)$$

Similarly, we define $A_i^m(r)$ as $(N-m-1)^{-1}$ times the number of vectors X_j^{m+1} within r of X_i^{m+1} , where j ranges from 1 to $N-m$, where $j \neq i$, and set

$$A^m(r) = \frac{1}{N-m} \sum_{i=1}^{N-m} A_i^m(r)$$

The parameter $SampEn(m, r)$ is then defined as $\lim_{N \rightarrow \infty} \{-\ln[A^m(r)/B^m(r)]\}$, which can be estimated by the statistic:

$$SampEn(m, r, N) = -\ln[A^m(r)/B^m(r)]$$

where N is the length of the time series, m is the length of sequences to be compared, and r is the tolerance for accepting matches.

b) Delay Vector Variance

We use the measure of unpredictability σ^{*2} described in [25]:

Time series can be represented in phase space using time delay embedding. When time delay is embedded into a time series, it can be represented by a set of delay vectors (DVs) of a given dimension m . The dimension of the delay vectors can then be expressed as $X(k) = [x_{(k-m\tau)} \dots x_{(k-\tau)}]$, where τ is the time lag. For every DV $X(k)$, there is a corresponding target, namely the next sample x_k . A set $\beta_k(m, d)$ is generated by grouping those DVs that are within a certain Euclidean distance d to DV $X(k)$. This Euclidean distance will be varied in a standardized manner with respect to the

distribution of pairwise distances between DVs. For a given embedding dimension m , a measure of unpredictability σ^{*2} (target variance) is computed over all sets of β_k .

The mean μ_d and the standard deviation σ_d are computed over all pair wise Euclidean distances between DVs given by $\|x(i) - x(j)\| (i \neq j)$. The sets $\beta_k(m, d)$ are generated such that $\beta_k = \{x(i) \setminus \|x(k) - x(j)\| \leq d\}$ i.e., sets which consist of all DVs that lie closer to $X(k)$ than a certain distance d , taken from the interval $[\mu_d - n_d \sigma_d, \mu_d + n_d \sigma_d]$ where n_d is a parameter controlling the span over which to perform DVV analysis.

For every set $\beta_k(m, d)$ the variance of the corresponding targets $\sigma_k^2(m, d)$ is computed. The average over the N sets $\beta_k(m, d)$ is divided by the variance of the time series signal σ_x^2 , σ_k gives the inverse measure of predictability, namely target variance σ^{*2} .

$$\sigma^{*2} = \frac{(1/N) \sum_{k=1}^N \sigma_k^2}{\sigma_x^2}$$

C) Surrogates and z-score.

The most commonly used null hypothesis considers that a given time series is generated by a Gaussian linear stochastic process and collected through a nonlinear measurement static function. Thus surrogates must have the same linear properties (autocorrelation and amplitude distribution) as the original signal. However, any underlying nonlinear dynamic structure within the original data is altered in the surrogates by phase randomization [16].

The statistics of significance z-score is,

$$Z_{score} = \frac{|q_0 - \langle q_s(i) \rangle|}{\sigma_q(i)}$$

where q_0 stands for the statistic on the original time series, $\langle q_s(i) \rangle$ for the mean and $\sigma_q(i)$ for the standard deviation of the surrogate, for $i=1, 2, \dots, M$ (number of generated surrogate). The critical value of z-score is 1.96 [26].

III. Results

A) Results on synthetic signals

In this section we study the evolution of the values generated by the four methods with variable complexity degree (CD) of the Henon synthetic model in four cases; 1) direct application of the method with no added noise, 2) using surrogates with no added noise, 3) direct application of the method with added noise and 4) using surrogates with added noise. The added noise is a white Gaussian noise ($SNR=5$ db) while CD varies between 0 and 1, for the Henon model. Our first objective is to test the sensitivity of the methods to varying CD for signals with and without noise. The use of surrogates is computationally very expensive and therefore our second objective is to test if the use of surrogates improves the method sensitivity or not. .

We compare the methods using two criteria, the method's sensitivity to the change of CD (slope of the curve "value of the method" vs. " CD ") and the MSE of the method for different values of CD .

Figure 3-A1 presents the mean value for each method (direct method value) as a function of CD computed from the 30 Monte Carlo instances of the signal generated by the Henon model. Figure 3-A2 presents the MSE of the methods for each CD . We see in Figure 3-A1 that in the direct case without noise, the four methods evolve well but with differences in their sensitivity (slopes). Tr and LE are more sensitive than the other methods. In Figure 3-A2 we observe that Tr has a much lower MSE than LE.

Figure 3-B1 presents the effect of adding noise ($SNR=5$ db) on the methods. We notice no significant slope for the LE and SampEn. The sensitivity of Tr and DVV also decreases with the addition of noise. In the other hand we find, Figure 3-B2, that DVV and SampEn give the lowest MSE. However SampEn does not demonstrate any sensitivity to the variation of CD so this method is useless for the noisy signal. Tr gives an intermediate MSE and the highest sensitivity when compared to the other methods when applied to noisy signals.

We then applied the methods to the synthetic signals with surrogates using the z-score as measure, in order to test if the use of surrogates improves the results or not. Figure 3-C1 presents the z-score for each method versus CD . We note that all the methods reflect the non-linearity of the signal generated by the Henon model as theirs z-score are always above 1.96. In terms of sensitivity to CD variation, SampEn is the best, but with the highest MSE (Figure 3-C2). Tr presents an acceptable evolution for lower CD . But beyond $CD = 0.4$ an unexpected decrease occurs in the curve and the Tr value remains constant after $CD=0.7$. This method however, gives the lowest MSE (Figure 3-C2). The DVV method presents an intermediate slope, contrary to the LE that presents no change with CD . Both DVV and LE have low MSE under these conditions.

The methods were then applied to the signals using again surrogates and z-score but with added noise (SNR=5db). All the methods still reveal the nonlinearity of the model. Indeed z-score is above 1.96 for all the methods, except for DVV where it gives a z-score value lower then 1.96 for CD between 0.4 and 0.6. We can clearly notice an increase in the sensitivity of Tr, Figure 3-D1, compared to the case in Figure 3-C1. SampEn has a good evolution beyond $CD = 0.4$ but, on the other hand, it presents a rapid increase in MSE (Figure 3-D2). The LE and DVV do not evolve as a function of CD (Figure 3-D1) and give similar MSE as Tr (Figure 3-D2).

B) Results on real signals

The different nonlinear methods were applied to real uterine EMG signals (EHG), first direct application of the method, and then with surrogates. We also computed three classical linear frequency based parameters from these real signals. The values were then used to discriminate the pregnancy and labor contractions. We used ROC curves in order to test the discriminating power of each case.

Our first objective was to test if the use of surrogates improves the discrimination of EHG bursts recorded during Pregnancy or Labor. Our second objective was to compare the performances of linear and nonlinear methods and to verify that the nonlinear methods reveal the evolution of EHG characteristics better than the linear ones. The ROC curves obtained with the different methods without and with use of surrogates are presented Figure 4-B and Figure 4-C respectively. The

characteristics of all the ROC curves without and with use of surrogates are presented in Table I and Table II respectively. From these data, it is clear that nonlinear methods improve the discrimination of pregnancy and labor signals. Indeed, the highest Area Under Curve AUC (0.842), sensitivity (0.86) and specificity (0.72) are obtained for the Tr method whatever the nonlinear or linear methods used. The MPF and LE methods also give an acceptable performance (Figure 4-B) with AUC=0.778 and AUC=0.758 respectively. The performances in correct discrimination of labor varies markedly from AUC=0.478 with SampEn to AUC=0.842 with Tr. When surrogates are used, all ROC curves present approximately the same appearance with the highest AUC=0.650 obtained for SampEn. Using surrogates we notice that the performance of SampEn improves while that of DVV remains approximately the same. On the other hand, the performance of Tr and LE seem to decrease with the use of surrogates. Finally, we can conclude from Figure 4 and Table I that nonlinear methods can provide better discrimination between pregnancy and labor contractions compared to the linear methods. Furthermore, even if the use of surrogates improves the performance of some methods, it does not generally improve the discrimination results.

IV. Discussion and conclusion

We analyzed, quantitatively and as comprehensively as possible, four different nonlinear analysis methods (Tr, SampEn, DVV and LE). These methods were applied on synthetic signals, in order to test their sensitivity to the change in signal complexity, in normal and noisy conditions, with or without using surrogates. All four methods were found to reflect correctly the increasing complexity of the signals in the noise free case, but with different sensitivities. In the case of added noise and direct application of the method, as expected, a decrease in the sensitivity of all methods occurred at a low Signal to Noise Ratio (SNR=5db). Indeed, at this low SNR, none of the methods detected the varying complexity of the signal, except for Tr, which clearly reflected the increasing non-linearity. The sensitivity of SampEn increased with the use of surrogates and it gave the highest sensitivity of all the methods, in the case of surrogate use with no added noise. Indeed SampEn has previously been shown to be sensitive to many aspects of the signal characteristics, including the sampling rate of the signal [14], [11]. Unexpected results were obtained in the case of surrogate use and with added noise.

Tr was more sensitive when compared to the previous case, and SampEn still presented a good sensitivity. We noticed that in the case of surrogate use, SampEn gave the highest sensitivity but also had the highest MSE, making it unreliable.

In this paper we also presented results obtained using nonlinear and linear methods for discrimination of EHG bursts recorded during pregnancy and labor. Comparison between the methods indicated that Tr, which is a nonlinear method, applied without using surrogates is clearly better in discriminating correctly pregnancy and labor contractions than the other methods. We can see also that the use of surrogates improves the performance of some methods like SampEn. These results confirm the results obtained during the study on synthetic signal, since the sensitivity of SampEn increases if surrogates are used, a posteriori justifying the use of the Henon model.

To sum up, the main findings of this study are the following: (i) Some of the studied methods are insensitive to varying signal complexity; (ii) SampEn performance depends on the use of surrogates; (iii) Generally speaking, none of the studied methods performed best in all the studied situations; (iv) Tr is very sensitive to change of model complexity, giving average or good performances, associated with the lowest MSE in most situations.

This leads to the conclusion that, of the four methods tested, Tr performed best for our application on real EHG. Indeed Tr deals robustly with real, usually noisy, signals and has a good sensitivity to complexity, one of the EHG characteristics that permits discrimination of uterine contraction efficiency. Using surrogates and the z-score, as a measure of nonlinearity, does not seem to bring any improvement to Tr. Therefore we will not use them for further work on EHG when using Tr.

There are some weaknesses in our study of which we are aware and aim to improve. Tr is dependent on the length of the signal and on the choice of the time delay (τ) and we aim to find a method to optimize these parameters. In further work we also aim to use all of the available bipolar channels (VA1,...,VA12) instead of only one channel, as in this work. This has been shown to dramatically increase the discrimination rate as evidenced in prior work [27].

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٢٦٠ **Conflict of interest**

٢٦١ No conflict of interest.

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٢٦٤ **Ethical Approval**

٢٦٥ The measurements in Iceland were approved by the relevant ethical committee (VSN 02-0006-V2),
٢٦٦ those in France approved by the regional ethical committee (ID-RCB 2011-A00500-41) of Amiens
٢٦٧ Hospital.

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Fig. 1. Simulated signal generated using Henon model with different complexity degrees (CD). Top: $CD = 0.1$, Bottom: $CD = 0.9$

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Fig. 2. Electrode placement (left), monopolar configuration and the corresponding bipolar signals BPi (right).

Fig. 3. Results obtained for Henon model using Monte-carlo simulation. On the left: Evolution of the methods with variable complexity in different cases. On the right: MSE of the methods function of complexity degree in different cases. (A) Direct method with no added noise, (B) Direct method with added noise, (C) With surrogate use and no added noise, (D) With surrogate use and added noise.

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Fig. 4. Example of ROC curves obtained for the detection of labor with the different linear and nonlinear methods. (A) Real Pregnancy and Labor contractions, (B) Direct method, (C) With surrogate use.

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TABLE I (Direct method)

Comparison of ROC curves for labor detection

Parameter	AUC	Specificity	Sensitivity
Time reversibility	0.842	0.721	0.860
Sample Entropy	0.478	0.382	0.643
Lyapunov Exponent	0.758	0.643	0.756
Delay Vector Variance	0.615	0.582	0.600
Mean Power Frequency	0.778	0.678	0.730
Peak Frequency	0.561	0.582	0.600
Median Frequency	0.654	0.556	0.704

TABLE II (with surrogate use)
Comparison of ROC curves for labor detection

Parameter	AUC	Specificity	Sensitivity
Time reversibility	0.560	0.513	0.626
Sample Entropy	0.650	0.593	0.643
Lyapunov Exponent	0.614	0.591	0.530
DVV	0.642	0.573	0.669