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AdvFPCG-Delineator: Advanced delineator for fetal phonocardiography / Tomassini, S.; Sbröllini, A.; Strazza, A.; Sameni, R.; Marcantoni, I.; Morettini, M.; Burattini, L.. - In: BIOMEDICAL SIGNAL PROCESSING AND CONTROL. - ISSN 1746-8094. - ELETTRONICO. - 61:(2020). [10.1016/j.bspc.2020.102021]

*Availability:*

This version is available at: 11566/282806 since: 2024-05-08T14:31:17Z

*Publisher:*

*Published*

DOI:10.1016/j.bspc.2020.102021

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5  
6 TITLE - AdvFPCG-Delineator: Advanced delineator for fetal phonocardiography  
7 AUTHORS - Selene Tomassini, Agnese Sbrollini, Annachiara Strazza, Reza Sameni, Ilaria  
8 Marcantoni, Micaela Morettini, Laura Burattini.  
9 PUBLISHED ONLINE: 22 June 2020  
10 YEAR - 2020  
11 DOI - 10.1016/j.bspc.2020.102021  
12 JOURNAL - Biomedical Signal Processing and Control  
13 VOLUME - 61  
14 ISSUE August 2020  
15 ARTICLE NUMBER - 102021  
16 PUBLISHER- Elsevier  
17 Full text link - <https://www.sciencedirect.com/science/article/pii/S1746809420301774?via%3Dihub>

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3 **AdvFPCG-Delineator:**  
4 **Advanced Delineator for Fetal Phonocardiography**  
5

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1 **Abstract**

2 *Fetal phonocardiogram (FPCG) consists in the recording of fetal heart sounds by means of a sensor*  
3 *placed on the mother’s abdominal surface. Usually, FPCG includes two major sounds for each fetal*  
4 *cardiac cycle: S1, produced by the sudden closure of mitral and tricuspid valves, and S2 produced*  
5 *by the closure of aortic and pulmonary valves. The aim of the present study was to propose AdvFPCG-*  
6 *Delineator for automatic fetal S1 and S2 identification and to demonstrate its reliability in different*  
7 *clinical conditions. The method consists of a wavelet-based filtering procedure followed by the*  
8 *computation of the scalogram, from which S1 and S2 were identified using a threshold-based*  
9 *algorithm. AdvFPCG-Delineator was tested on the “Simulated Fetal PCGs database” (37 FPCG*  
10 *signals) and on the experimental “Shiraz University fetal heart sounds database” (119 FPCG*  
11 *signals), both available at PhysioNet (<https://physionet.org>). Manual S1 and S2 annotations and*  
12 *simultaneously acquired cardiocographic recordings were used to compute reference fetal heart*  
13 *rate (FHR) for the simulated and experimental databases, respectively. No statistically significant*  
14 *difference was observed between estimated vs reference FHR (median: 140 bpm vs 140 bpm,*  
15 *respectively) for the simulated database, for which AdvFPCG-Delineator was also be able to track*  
16 *beat-to-beat variability (correlation over 92%). Additionally, no statistically significant difference*  
17 *was observed between estimated vs reference FHR (141 bpm vs 140 bpm, respectively) for the*  
18 *experimental database, even when stratifying by clinical conditions (maternal age, gestational age,*  
19 *etc.). In conclusion, AdvFPCG-Delineator proved to be a reliable method to automatically identify*  
20 *S1 and S2 from fetal phonocardiograms.*

21  
22 **Keywords:** Fetal heart-rate extraction; Fetal heart-sound identification; Fetal phonocardiography;  
23 Phonocardiographic features; Scalogram; Wavelet denoising

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25

## 1. Introduction

Fetal phonocardiography (FPCG) consists in the recording of fetal heart sounds (FHSs) occurring during each fetal cardiac cycle by means of a sensor placed on the mother's abdominal surface [1-4]. FHSs are produced by the mechanical action of the cardiac muscles and valves and by the blood motion through the heart chambers [2,4]. Under normal conditions, a FPCG signal includes two major FHSs during each cardiac cycle [2-8]: the first sound (S1), produced by the sudden closure of mitral and tricuspid valves, *i.e.* atrio-ventricular valves, and the second sound (S2), produced by the closure of aortic and pulmonary valves, *i.e.* semilunar valves. S1 represents the longest and the loudest sound since the mitral valve closes with greater force.

Since representing specific phenomena of the cardiac cycle, FPCG is considered a potentially useful technique for fetal monitoring [9]. Indeed, it provides important diagnostic information for the assessment of fetal well-being during pregnancy [2] in a non-invasive, harmless (or passive, since no energy is emitted to the mother's abdomen and thus to the fetus), affordable (low-cost instrumentation), reliable and simple fashion, even as a user-independent home monitoring tool [2,4]. FPCG analysis allows measurement of the cardiac cycle, defined by the time interval between two consecutive S1 (S1S1 interval), from which fetal heart rate (FHR) can be assessed. In turn, from the 24<sup>th</sup> gestational week on, continuous and long-term FHR monitoring represents the most frequently used diagnostic tool for fetal health assessment [10]. Additionally, FPCG analysis provides information about the systolic time interval, defined as the time interval between an S1 and the subsequent S2 (S1S2 interval) [7-8], and the diastolic time interval, defined as the time interval between an S2 and the subsequent S1 (S2S1 interval) [7], with S1S2 interval being rather constant and normally shorter than S2S1 interval ( $S1S2 > 0.1$  s;  $S2S1 > 0.2$  s) [11]. FPCG analysis is however not trivial. FPCG signals are highly non-stationary and weak because of the high attenuation along the sound transmission pathway due to the physical distance between the acoustic source and the transducer [4,12-13]. In addition, there could be impedance mismatch between the transducer and the maternal abdominal surface during the acquisition [5]. Consequently, FPCG signals are typically heavily corrupted by several sources of noise, *i.e.* internal noise from the mother (such as mother respiratory sound) and external noise from the environment (such as electrode motion artifact) [2-3,10,14-16], thus requiring filtering procedures to be applied before S1 and S2 identification [2-4,6,15-17]. Moreover, quality of FPCG signals highly depends on fetal position, also with respect to FPCG sensor location [18], which is highly uncertain and variable, even though usually, starting from the 30<sup>th</sup>-35<sup>th</sup> weeks of gestation, the fetus presents the vertex position [18].

Development of accurate techniques to automatically analyze FPCG is currently very desirable [9]. Several algorithms proposed in the literature are mainly focused on S1 identification in order to evaluate FHR [1-2,9,11]; others also challenged to detect S2 [4,6,9,19] in order to have additional information on the systolic and diastolic intervals and on cardiac valves functioning. Still, a reliable automatic software able to denoise FPCG signals and to characterize FHSs when applied to different clinical conditions is yet nonexistent [9]. In a recent study, we proposed PCG-Delineator as an efficient algorithm for automatic identification of both S1 and S2 in FPCG [11], which included a procedure based on the wavelet transform (4<sup>th</sup>-order Coiflet mother wavelet with thresholding settings consisting of soft universal thresholding rule and 7 decomposition levels) for noise suppression. When tested on the "Simulated Fetal PCGs database" [20] of PhysioNet [21], PCG-Delineator provided promising results, being able to significantly reduce noise (median signal-to-noise ratio increased from 0.15 dB to 15.86 dB) and to accurately detect S1 (sensitivity: 88%; positive predictive value: 91%). Sensitivity of S2 detection, however, was under 80% (sensitivity: 77%; positive predictive value: 99%). Additionally, PCG-Delineator was tested only on simulated and not on experimental data. In a successive study reporting a comparative analysis of wavelet-transform

1 filtering procedures [22], we concluded that the filter based on wavelet transform, obtained by  
2 combining the 4<sup>th</sup>-order Coiflet mother wavelet with the thresholding settings constituted of the soft  
3 universal rule and 7 decomposition levels, is optimal for FPCG filtering according to evaluation  
4 criteria based on both noise and clinical features. The aim of the present study was to propose  
5 AdvFPCG-Delineator (Advanced FPCG-Delineator) as an improved version of PCG-Delineator  
6 integrating all our most recent findings on FPCG filtering and fetal S1 and S2 identification, and to  
7 demonstrate its reliability in different clinical conditions. The scope is to make available a clinically  
8 useful tool able to reliably provide information on FHR as well as fetal cardiac valves functioning  
9 based on FPCG. To this aim, both PCG-Delineator and AdvFPCG-Delineator were tested on two  
10 FPCG databases (the “Simulated Fetal PCGs database” [20] and the “Shiraz University fetal heart  
11 sounds database” [13,15]) and results were compared. Eventually, AdvFPCG-Delineator will be  
12 freely available to researches for additional testing and evaluations.

13

## 14 **2. Methods**

15 In this work, FPCG signals from two databases (the “Simulated Fetal PCGs database” [20]  
16 and the “Shiraz University (SU) fetal heart sounds database” [13,15]) were submitted to PCG-  
17 Delineator [11] and AdvFPCG-Delineator for FPCG denoising and S1 and S2 identification.  
18 Performances of both methods were evaluated in terms of correctness of FHR computed from both  
19 S1S1 and S2S2 intervals.

20

### 21 **2.1. Data**

#### 22 **2.1.1. Simulated data**

23 The “Simulated Fetal PCGs database” [20], freely available at PhysioNet/PhysioBank [21],  
24 contains 37 simulated FPCG signals (8-min long) related to different fetal states and recording  
25 conditions. Specifically, simulated FPCG signals were obtained by summation of a sequence of  
26 simulated S1 and S2 waveforms with different levels of noise. Noise was obtained as superimposition  
27 of several contributions simulating internal noise (maternal heart sounds, maternal body organs  
28 sounds and fetal movements) and external noise (surrounding environments and white Gaussian  
29 noise). Signal-to-noise ratio values ranged from -26.7 to -4.4 dB and sampling frequency was 1 kHz.  
30 S1 and S2 were manually annotated.

31

#### 32 **2.1.2. Experimental data**

33 The “Shiraz University (SU) fetal heart sounds database” [13,15], freely available at  
34 PhysioNet/PhysioBank (and thus usable without any further institutional review board approval) [21],  
35 contains 119 raw experimental FPCG signals acquired from 109 pregnant women. Out of the 109  
36 pregnancies, 102 were single pregnancies (double recording available in 3 cases) and 7 were twin  
37 pregnancies (one recording for each fetus), corresponding to 116 fetuses. Of these, 60 were males, 54  
38 females and 2 with unknown gender. Additionally, 102 fetuses were healthy and 14 were not healthy  
39 (4 with decrease of amniotic fluid, 3 with abnormality in fetus non-stress test, 1 with severe change  
40 in FHR, 2 with high FHR, 3 with slow fetal growth and 1 with both decrease of amniotic fluid and  
41 slow fetal growth). Maternal age (MA) was 29±6 years; specifically, 59 pregnant women were young  
42 (MA<30 years), 43 middle-aged (30≤MA≤39 years) and 7 were old (MA>39 years) [23]. Maternal  
43 body mass index (BMI) was 29±4 kg/m<sup>2</sup>; specifically, 12 pregnant women were normal (BMI<25  
44 kg/m<sup>2</sup>), 55 overweight (25≤BMI≤30 kg/m<sup>2</sup>) and 42 were obese (BMI>30 kg/m<sup>2</sup>) [24]. Gestational  
45 age (GA) was 36±3 weeks; specifically, 75 pregnancies were early-term (GA<39 weeks), 32 full-  
46 term (39≤GA<41 weeks) and 2 late-term (GA≥41 weeks) [25]. All pregnant women were volunteers  
47 and data were fully deidentified before being placed in the public domain.

1 FPCG signals were recorded with the JABES<sup>TM</sup> electronic stethoscope, placed on the lower  
2 maternal abdominal parts, according to the locations advised by an expert gynecologist [15]. The  
3 FPCG length ranged from 29 s to 133 s (mean duration 90 s). Sampling frequency was 16 kHz except  
4 for few cases (termed f9-1, f63, f64, f65, f88 in the database) for which it was 8 kHz. FPCG signals  
5 originally sampled at 8 kHz were oversampled at 16 kHz in order to have all experimental FPCG  
6 signals equally sampled. No S1 and S2 annotations were available.

## 8 **2.2. Automatic fetal phonocardiogram delineation**

9 Both simulated and experimental data were submitted to AdvFPCG-Delineator and PCG-  
10 Delineator in order to automatically identify S1 and S2. Both procedures were implemented in  
11 Matlab®.

### 13 **2.2.1. AdvFPCG-Delineator**

14 AdvFPCG-Delineator integrates a wavelet-based filtering procedure, finalized to reduce the  
15 level of noise affecting FPCG signals, followed by the scalogram computation and by the S1 and S2  
16 identification procedures. The block diagram of the algorithm is reported in Figure 1.

17 According to the wavelet-based filtering procedure, each FPCG signal was normalized by its  
18 maximum amplitude and rescaled so that its amplitude could vary between  $\pm 100$ . Normalized FPCG  
19 signals were pre-filtered by application of a 6<sup>th</sup>-order bandpass bidirectional Butterworth filter with  
20 lower and upper cut-off frequencies of 20 Hz and 120 Hz, respectively [11,16,22,26], before being  
21 passed through a filter based on the wavelet transform [22], for further noise removal. Such filter  
22 consisted in a combination of the 4<sup>th</sup>-order Coiflet mother wavelet [4,11-12,15-16,22,26-28] with soft  
23 universal thresholding rule [2-4,11,22,27-29], using 7 decomposition levels [11,16,22]. According to  
24 the standard dyadic structure of the wavelet transform, at first level, the pre-filtered FPCG signal was  
25 decomposed into approximation coefficients (*i.e.* low-frequencies FPCG components) and detail  
26 coefficients (*i.e.* high-frequency FPCG components); then, at every level, the approximation  
27 coefficients were further decomposed into approximation and detail coefficients through a cascade of  
28 low-pass and high-pass filters followed by down-sampling, while the detail coefficients were not  
29 decomposed any further [12,30]. To denoise FPCG signal, the detail coefficients were thresholded at  
30 each level. Finally, the detail coefficients of all the levels and the approximation coefficient of the  
31 last level were summed up to reconstruct the wavelet-transform filtered FPCG signal [12,30].

32 The scalogram [4,31-32] of filtered FPCG signal (Figure 1) was obtained using the 4<sup>th</sup>-order  
33 Coiflet mother wavelet; the continuous wavelet coefficients were computed using the conventional  
34 scale interval 1 to 100 [33].

35 Eventually, S1 and S2 identification was performed on the scalogram using widely accepted  
36 FPCG physiological properties. At first, S1 were identified, one after another, with a procedure based  
37 on a temporal threshold. An S1, to be physiologically acceptable, had to occur at least 300 ms from  
38 the previous S1. Successively, S2 were identified. S2 identification was also based on a temporal  
39 threshold. An S2 had to satisfy the following conditions: there is only one S2 between two consecutive  
40 S1; and S2 had to fall at least 100 ms after the preceding S1 and at most 200 ms before the successive  
41 S1, in order to have the S2S1 interval longer than the S1S2 interval [5,7-8,11].

### 43 **2.2.2. PCG-Delineator**

44 Details of the PCG-Delineator algorithm can be found elsewhere [11]. Briefly, PCG-  
45 Delineator integrates the same wavelet-based filtering procedure in AdvFPCG-Delineator directly  
46 followed by the S1 and S2 identification procedures (no scalogram computation is performed).

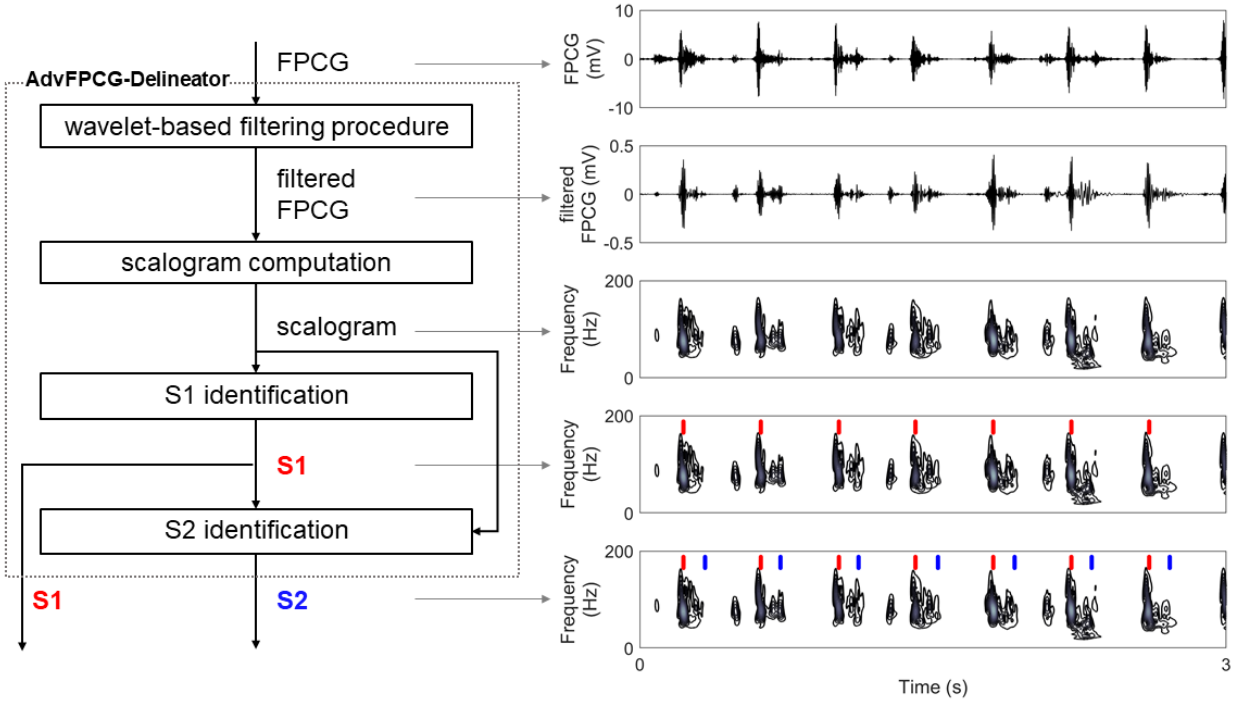


Figure 1: The block diagram of AdvFPCG-Delineator.

### 2.3. Performances evaluation

S1 and S2 annotations were available for the simulated data but not for the experimental data. Thus, while for simulated FPCG signals reference FHR ( $FHR_{ref}$ ) could be computed by means of S1S1 intervals obtained using the annotations, for the experimental FPCG signals  $FHR_{ref}$  had to be indirectly obtained from the simultaneously acquired cardiocotographic (CTG) recordings [15] available at Shiraz University. However, cardiocotography provides mean FHR values obtained by averaging over 10-second windows. Thus, for consistency, in all cases we referred to  $FHR_{ref}$  as the mean FHR over 10-second windows, obtained using S1 annotations for the simulated data, and CTG data for the experimental data.

Performances of PCG-Delineator and AdvFPCG-Delineator were evaluated in terms of number of S1 and S2 identifications (which should be equal), S1S1, S2S2, S1S2 and S2S1 intervals (which should be physiologic), and of mean FHR over 10-second windows ( $FHR_{10s}$ , which should be equal to  $FHR_{ref}$ ). In particular, performances evaluation in terms of FHR was performed by estimating  $FHR_{10s}$  through S1S1 interval ( $FHR_{S1S1}$  in bpm) and S2S2 interval ( $FHR_{S2S2}$  in bpm):

$$FHR_{S1S1} = 60 / \text{mean}(S1S1), \quad (1)$$

$$FHR_{S2S2} = 60 / \text{mean}(S2S2). \quad (2)$$

In Equations (1) and (2), 60 is the number of seconds in a minute, and  $\text{mean}(S1S1)$  and  $\text{mean}(S2S2)$  are mean intervals (in s) over 10-second windows.

Next, FHR-estimation errors with respect to reference were computed as in Equations (3) and (4) (by definition, reference errors were equal to zero):

$$\varepsilon_{S1S1} = FHR_{ref} - FHR_{S1S1}, \quad (3)$$

$$\varepsilon_{S2S2} = FHR_{ref} - FHR_{S2S2}. \quad (4)$$



For simulated data only, performances of AdvFPCG-Delineator and PCG-Delineator in evaluating beat-to-beat variations was performed by comparing estimated S1S1 and S2S2 intervals against corresponding intervals computed using reference annotations, and by computation of the correlation coefficient ( $\rho$ ) as well as slope ( $m$ ) and intercept ( $q$ ) of regression line between estimated vs reference S1S1 and S2S2 intervals (in case of perfect automatic S1 and S2 identification  $\rho=1$ ,  $m=1$  and  $q=0$ ).

For the experimental data only, robustness of AdvFPCG-Delineator was also evaluated in relation to MA, BMI, GA, single vs twin pregnancies, gender of the fetus and fetus clinical conditions.

In all cases, to compare FPCG features (either intervals, FHR values and errors), normality of distributions were first assessed by means of the Lilliefors test [34]; then, non-normal distributions were described in terms of 50<sup>th</sup> [25<sup>th</sup>; 75<sup>th</sup>] percentiles and compared by means of the Wilcoxon Rank-Sum test [35]. Statistical level of significance ( $P$ ) was set at 0.05.

### 3. Results

#### 3.1. Simulated data

FPCG features obtained using reference S1 and S2 annotations and S1 and S2 identifications by PCG-Delineator and AdvFPCG-Delineator over the simulated FPCG database are reported and compared in Table 1. Overall, AdvFPCG-Delineator identified more S1 and S2 than PCG-Delineator so that the number of missed S1 and especially S2 identifications was lower for AdvFPCG-Delineator (343 for S1 and 380 for S2) than for PCG-Delineator (627 for S1 and 3583 for S2). Additionally, no statistically significant difference was observed between FPCG intervals obtained using AdvFPCG-Delineator and the reference ones; instead, FPCG intervals obtained using PCG-Delineator were all significantly longer than the reference ones, and thus than those obtained using AdvFPCG-Delineator.

	Reference	PCG-Delineator	AdvFPCG-Delineator
Number of S1	41440	40813	41097
Number of S2	41440	37857	41060
S1S1 interval (ms)	410[410;410]	420[419;436]*	413[406;417]
S2S2 interval (ms)	411[411;411]	424[423;443]*	418[417;426]
S1S2 interval (ms)	138[138;138]	140[129;160]*	138[120;145]§
S2S1 interval (ms)	272[272;272]	294[245;305]*	274[240;286]§

Table 1. Phonocardiographic features of FPCG signals in the “Simulated Fetal PCGs database” according to reference annotations, PCG-Delineator and AdvFPCG-Delineator.

n.a. not applicable.

\*:  $P<0.05$ , when comparing PCG-Delineator and AdvFPCG-Delineator FPCG features vs reference;

§:  $P<0.05$ , when comparing PCG-Delineator vs AdvFPCG-Delineator FPCG features.

Figure 2 displays beat-to-beat associations between S1S1 and S2S2 intervals obtained using PCG-Delineator and AdvFPCG-Delineator vs reference S1S1 and S2S2 intervals. Compared to PCG-Delineator, AdvFPCG-Delineator provided much higher  $\rho$  ( $\rho=0.94$  vs  $0.40$  for S1S1;  $\rho=0.92$  vs  $0.38$  for S2S2), a value of  $m$  closer to 1 ( $m=0.93$  vs  $0.64$  for S1S1;  $m=0.91$  vs  $0.73$  for S2S2) and a value of  $q$  closer to 0 ( $q=28$  vs  $156$  for S1S1;  $q=36$  vs  $117$  for S2S2).

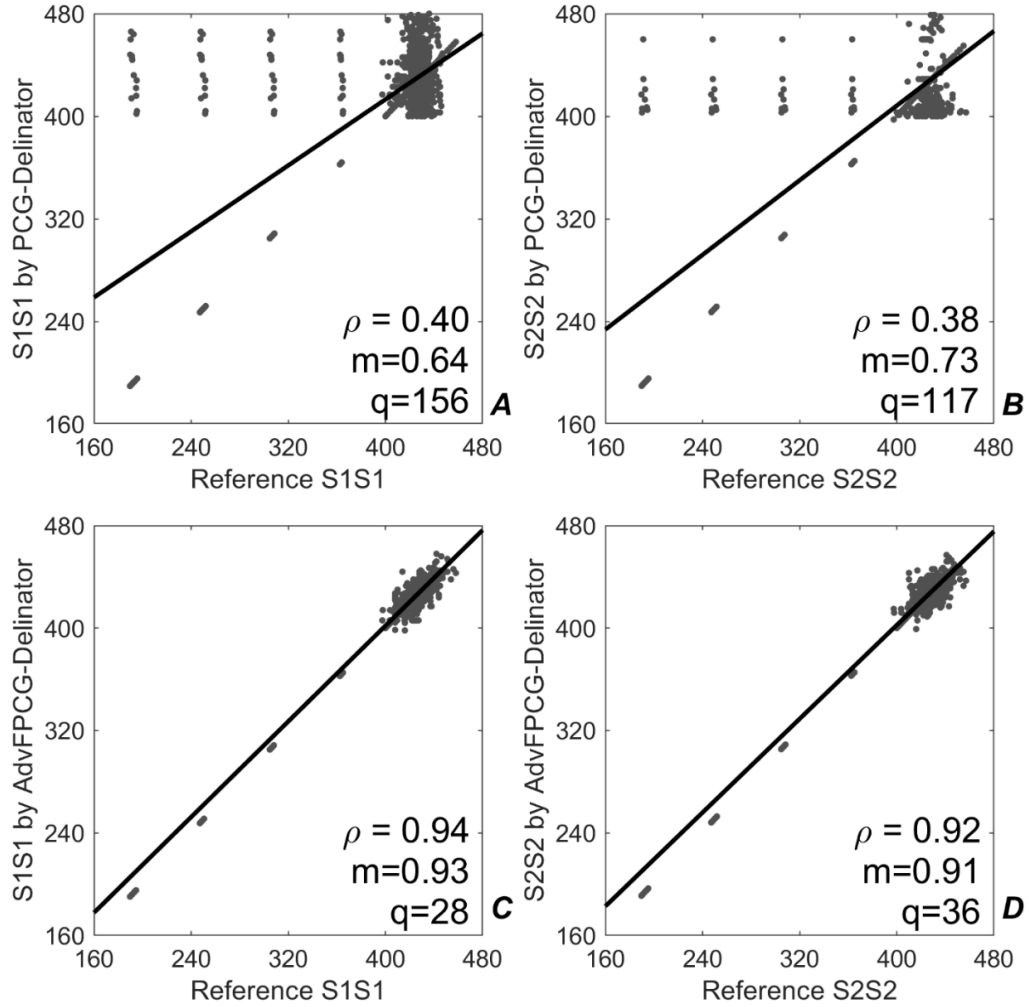


Figure 2. Beat-to-beat associations between S1S1 and S2S2 intervals obtained using PCG-Delineator (panel A and panel B) and AdvFPCG-Delineator (panel C and panel D) vs reference S1S1 and S2S2 intervals;  $\rho$ ,  $m$  and  $q$  indicate correlation coefficients and slopes and intercepts of the regression lines, respectively.

Table 2 reports the comparison of  $FHR_{10s}$  and related errors obtained using reference annotations, PCG-Delineator and AdvFPCG-Delineator. AdvFPCG-Delineator correctly estimated  $FHR_{10s}$  using both S1S1 and S2S2 intervals, so that errors were zero in all cases. Instead, PCG-Delineator provided correct estimate of  $FHR_{10s}$  when using S1S1 interval but not when using S2S2 interval; errors were different from zero in both cases.

FPCG feature	Reference	PCG-Delineator		AdvFPCG-Delineator	
		S1S1	S2S2	S1S1	S2S2
$FHR_{10s}$ (bpm)	140[140;140]	140[140;141]	134[127;139]*	140[139;140]	140[139;140]
$\epsilon$ (bpm)	0[0;0]	-1[-1;-1]	6[1;13]*	0[0;0]	0[0;0]

Table 2. Comparison between  $FHR_{10s}$  estimated using PCG-Delineator and AdvFPCG-Delineator vs reference for the “Simulated Fetal PCGs database”.

n.a.: not applicable;

\*:  $P < 0.05$ , when comparing PCG-Delineator and Advanced PCG-Delineator vs reference.

### 3.2. Experimental data

FPCG features obtained using S1 and S2 identifications by PCG-Delineator and AdvFPCG-Delineator over the experimental database are reported and compared in Table 3. AdvFPCG-Delineator identified more S1 and S2 than PCG-Delineator; moreover, the difference between the number of S1 and the number of S2 (ideally equal to zero) is much smaller for AdvFPCG-Delineator (124) than for PCG-Delineator (3357). No statistically significant difference was observed only between S1S1 intervals, while S2S2, S1S2 and S2S1 intervals obtained using AdvFPCG-Delineator were all significantly shorter than the ones obtained using PCG-Delineator.

	<b>PCG-Delineator</b>	<b>AdvFPCG-Delineator</b>
Number of S1	20383	20829
Number of S2	17026	20705
S1S1 interval (ms)	429[421;435]	421[413;428]
S2S2 interval (ms)	473[461;494]	422[413;429]*
S1S2 interval (ms)	147[137;170]	140[129;152]*
S2S1 interval (ms)	284[255;343]	283[268;292]*

Table 3. Phonocardiographic features of the “Shiraz University (SU) fetal heart sounds database” according to PCG-Delineator and AdvFPCG-Delineator.

n.a. not applicable.

\*:  $P < 0.05$ , when comparing AdvFPCG-Delineator vs PCG-Delineator.

Overall, CTG recordings provided  $FHR_{ref}$  values relative to 653 (61%) out of 1071 10-second windows available from experimental FPCG signals (the other windows were too noisy). The number of windows for which it was possible to evaluate  $FHR_{S1S1}$  and  $FHR_{S2S2}$  was 825 (77%) for AdvFPCG-Delineator and 931 (87%) for PCG-Delineator, both including the above-mentioned 653 windows for which CTG  $FHR_{ref}$  were available.

Table 4 reports the comparison of  $FHR_{10s}$  and related errors according to PCG-Delineator and AdvFPCG-Delineator vs CTG reference over the 653 10-second windows. AdvFPCG-Delineator provided  $FHR_{10s}$  estimates (*i.e.*  $FHR_{S1S1}$  and  $FHR_{S2S2}$ ) not significantly different from reference and thus corresponding errors not significantly different from 0. Instead, PCG-Delineator provided  $FHR_{10s}$  estimate by S2S2 interval (*i.e.*  $FHR_{S2S2}$ ) significantly different from reference and corresponding error significantly greater than zero.

FPCG feature	CTG	PCG-Delineator		AdvFPCG-Delineator	
	Reference	S1S1	S2S2	S1S1	S2S2
$FHR_{10s}$ (bpm)	141[135;149]	139[135;145]	122[116;135]*	141[138;145]	141[137;145]
$\epsilon$ (bpm)	0 [0;0]	1[-4;10]	19[11;27]*	-1[-6;6]	-1[-6;6]

Table 4. Comparison between  $FHR_{10s}$  estimated using PCG-Delineator and AdvFPCG-Delineator vs cardiocographic (CTG) reference for the “Shiraz University (SU) fetal heart sounds database” (specifically, for the 653 10-second windows for which CTG  $HR_{10s}$  were available).

n.a.: not applicable;

\*:  $P < 0.05$ , when comparing PCG-Delineator and AdvFPCG-Delineator vs CTG reference.

The results of the robustness of AdvFPCG-Delineator in relation to MA, BMI, GA, single vs twin pregnancies, gender of the fetus and fetus clinical conditions are reported in Table 5. In all cases,

1 no statistically significant differences were obtained when comparing FHR<sub>10s</sub> and related errors  
2 according to AdvFPCG-Delineator vs CTG reference.

3

		N	FPCG Feature	CTG	AdvFPCG-Delineator	
				Reference	S1S1	S2S2
MA (years)	<30 (54%)	59	FHR <sub>10s</sub> (bpm)	141[135;147]	142[136;148]	142[137;147]
			$\epsilon$ (bpm)	0[0;0]	0[-2;0]	0[-2;0]
	30-39 (40%)	43	FHR <sub>10s</sub> (bpm)	143[143;150]	143[136;151]	143[136;151]
			$\epsilon$ (bpm)	0[0;0]	0[-2;1]	0[-2;0]
	>39 (6%)	7	FHR <sub>10s</sub> (bpm)	140[138;144]	140[137;144]	140[137;144]
			$\epsilon$ (bpm)	0[0;0]	0[-1;0]	0[-1;0]
BMI (kg/m <sup>2</sup> )	<25 (11%)	12	FHR <sub>10s</sub> (bpm)	138[132;144]	139[133;145]	139[133;146]
			$\epsilon$ (bpm)	0[0;0]	0[-3;1]	0[-3;0]
	25-30 (50%)	55	FHR <sub>10s</sub> (bpm)	140[133;147]	141[135;148]	141[136;148]
			$\epsilon$ (bpm)	0[0;0]	0[-2;0]	0[-2;0]
	>30 (39%)	42	FHR <sub>10s</sub> (bpm)	143[137;150]	144[138;150]	144[138;150]
			$\epsilon$ (bpm)	0[0;0]	0[-3;1]	0[-3;2]
GA (weeks)	<39 (69%)	75	FHR <sub>10s</sub> (bpm)	141[135;147]	141[136;148]	142[136;148]
			$\epsilon$ (bpm)	0[0;0]	0[-2;0]	0[-2;0]
	39-41 (29%)	32	FHR <sub>10s</sub> (bpm)	143[137;149]	143[137;148]	144[137;149]
			$\epsilon$ (bpm)	0[0;0]	0[-1;1]	0[-2;0]
	$\geq 41$ (2%)	2	FHR <sub>10s</sub> (bpm)	156[139;163]	155[143;162]	155[143;159]
			$\epsilon$ (bpm)	0[0;0]	0[-2;1]	0[-2;3]
Pregnancy type	Single (94%)	102	FHR <sub>10s</sub> (bpm)	141[135;149]	142[136;149]	142[136;148]
			$\epsilon$ (bpm)	0[0;0]	0[-3;1]	0[-3;1]
	Twin (6%)	7	FHR <sub>10s</sub> (bpm)	143[138;149]	143[138;148]	144[139;149]
			$\epsilon$ (bpm)	0[0;0]	0[-1;0]	0[-1;0]
Fetus gender	Male (52%)	60	FHR <sub>10s</sub> (bpm)	143[136;150]	144[137;150]	144[137;150]
			$\epsilon$ (bpm)	0[0;0]	0[-3;1]	0[-3;1]
	Female (46%)	54	FHR <sub>10s</sub> (bpm)	140[133;145]	141[135;147]	141[135;147]
			$\epsilon$ (bpm)	0[0;0]	0[-2;0]	0[-2;0]
	Unknown (2%)	2	FHR <sub>10s</sub> (bpm)	132[131;138]	137[135;140]	137[133;139]
			$\epsilon$ (bpm)	0[0;0]	-2[-7;1]	0[-6;2]
Fetus health condition	Healthy (88%)	102	FHR <sub>10s</sub> (bpm)	140[134;148]	141[135;148]	141[135;148]
			$\epsilon$ (bpm)	0[0;0]	0[-2;1]	0[-2;1]
	Not healthy (12%)	14	FHR <sub>10s</sub> (bpm)	144[138;150]	145[140;150]	145[139;150]
			$\epsilon$ (bpm)	0[0;0]	0[-3;0]	0[-3;0]

4 Table 5. Robustness of AdvFPCG-Delineator in relation to MA, BMI, GA, single vs twin pregnancies, gender  
5 of the fetus and fetus health condition.

6 \*:  $P < 0.05$ , when comparing AdvFPCG-Delineator vs CTG reference.

7

#### 8 4. Discussion

9 This study proposed AdvFPCG-Delineator for FPCG filtering and fetal S1 and S2  
10 identification from fetal phonocardiograms, and demonstrated its reliability in different clinical  
11 conditions.

12 AdvFPCG-Delineator is an improved version of the previously proposed PCG-Delineator.  
13 Improvement of AdvFPCG-Delineator was obtained by adding the scalogram computation in its  
14 algorithm (Figure 1). Filtering is always required because FPCG signal is typically very noisy [2-  
15 3,10,14-16] and here it was performed using a filter based on the wavelet transform obtained by  
16 combining the 4<sup>th</sup>-order Coiflet mother wavelet with soft rule, universal thresholding algorithm and  
17 7 decomposition levels, since this combination was recently found to be optimal for FPCG filtering

1 [22]. Then, the scalogram [4,31-32] was computed to enhance FPCG signals by performing a time-  
2 frequency characterization before S1 and S2 identification. As a result, S1 and especially S2  
3 identification by AdvFPCG-Delineator resulted much more accurate. Indeed, when tested on the  
4 “Simulated Fetal PCGs database” [20] and the “Shiraz University (SU) fetal heart sounds database”  
5 [13,15], the number of identified S1 and S2 increased (+0.7% and +8.5%, respectively, for the  
6 simulated data, and +2.2% and +21.6%, respectively, for the experimental data) and the number of  
7 cardiac beats in which S1 and S2 were both detected increased (from 92.8% to 99.9% for the  
8 simulated data; and from 83.5% to 99.4% for the experimental data).

9 To further assure that the new identifications were not actually false positive identifications,  
10 FHR values were computed from S1S1 and S2S2 intervals and compared against FHR reference  
11 values. S1 and S2 annotations were available for the simulated data but not for the experimental data.  
12 Thus, while for simulated FPCG signals reference FHR could be computed by means of S1S1  
13 intervals obtained using the annotations, for experimental FPCG signals it had to be indirectly  
14 obtained from the simultaneously acquired cardiocographic recordings [14] available at Shiraz  
15 University. However, cardiocography provides mean FHR values obtained by averaging over 10-  
16 second windows. Thus, for consistency, in all cases we referred to reference FHR as the mean FHR  
17 over 10-second windows, obtained using S1 annotations for the simulated data, and cardiocographic  
18 data for the experimental data. For both simulated and experimental data, results indicate that, only  
19 for AdvFPCG-Delineator, FHR estimated using both S1S1 interval and S2S2 interval were not  
20 significantly different from reference (Tables 2 and 4), confirming the outperformance of this method  
21 with respect to PCG-Delineator.

22 Beside on FHR, the two methods were also evaluated on beat-to-beat variations, since these  
23 latter have an important prognostic value [36-37]. Such evaluation was not possible for the  
24 experimental data, since no S1 and S2 annotations were available; thus, it was performed only on the  
25 simulated data. Results indicated that AdvFPCG-Delineator is able to track S1S1 and S2S2 beat-to-  
26 beat variability accurately (high  $\rho$ ,  $m$  close to 1 and  $q$  close to 0). In particular, AdvFPCG-Delineator  
27 overcame the major limit of PCG-Delineator, which consists in the tendency to miss S1 and S2  
28 identifications (especially those belonging to short cardiac beats) and thus to provide estimated S1S1  
29 and S2S2 intervals too long with respect to reference (Figure 2, panel A and B).

30 AdvFPCG-Delineator also demonstrated to be reliable when evaluated in different clinical  
31 conditions. Indeed, its performance was independent on maternal age, maternal body mass index,  
32 gestational age, type of pregnancy (single vs twin), gender of the fetus and fetus health condition.  
33 AdvFPCG-Delineator robustness to fetal position and sensors location was not evaluated because  
34 such information was not annotated during recordings, and thus was not available to our knowledge.  
35 However, it should be observed that it is not possible to identify the optimal sensor position since it  
36 depends on fetal position, which is highly uncertain and variable. Consequently, sensor is usually  
37 located based on clinician experience. This solution allows to have good quality signals but implies  
38 having different sensors locations for different women.

39 Eventually, it is worth to observe that AdvFPCG-Delineator was designed exclusively for fetal  
40 monitoring and not for maternal monitoring. Nevertheless, real-time simultaneous monitoring of both  
41 fetus and mother is fundamental during pregnancy, since fetal wellbeing implies maternal wellbeing  
42 and vice versa. Future studies will evaluate possibility of using AdvFPCG-Delineator in real-time  
43 applications. Instead, maternal monitoring requires different tools specifically designed for adult  
44 subjects, since they have a completely different structural and physiological hearth morphology from  
45 fetuses [18].

#### 46 4.1. Conclusions

1 In this study, it was demonstrated that the hereby proposed AdvFPCG-Delineator represents  
2 a reliable method to identify S1 and S2 from fetal phonocardiograms, which is a very practical means  
3 of fetal cardiac status assessment.

#### 5 **Conflict of interest**

7 The authors declare that they have no known competing financial interests or personal relationships  
8 that could have appeared to influence the work reported in this paper.

#### 10 **Funding**

12 This research did not receive any specific grant from funding agencies in the public, commercial, or  
13 not-for-profit sectors.

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