

Telemedicine Digital Phonocardiography: Cost-Effective Strategies in Heart Failure Screening and Monitoring

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Abstract

Three studies were performed assessing the clinical value of digital electro- and phonocardiography (dECG, dPCG) with telemedicine application. In the first study, some Doppler echocardiographic parameters (ejection fraction, aortic Vmax, the grade of mitral and tricuspid regurgitation) were estimated from the spectral amplitude value of 170 time-frequency cells of the TriTest dPCG using multivariate discriminant analysis of 584 cardiac patients (292 for the training, and 292 for the test set). A cost analysis of heart failure (HF) screening in various populations was performed on 452 subjects. The greatest cost-savings (Euro / one HF patient) was found in the combined use of dECG and dPCG compared with the TE screening alone (mean: 82.4 CI-95%: 69.5-96.4 versus mean: 230.1 CI-95%: 196.5-254.4; $p < 0.001$). In the third study, during the 24 months telemonitoring of serious heart failure patients, 124 hospital days charge was saved, comparing the two, 29-29 patients' groups.

1. Introduction

In the last two decades, Tavel has written almost everything about the clinical use of auscultation and phonocardiography [1,2]. Our works started from this point, introducing more sophisticated computer application, and expanding the patients data set with digital phonocardiograms (dPCG), electrocardiograms (dECG), and the traditional transesophageal echocardiograms (TTE) [3,4]. From the first work on spectral analysis of the PCG [5] to the last [6], many efforts were made [7-10], mainly in pediatrics [11-13], without any significant change in everyday clinical use. The aim of our studies was to expand the clinical use of dPCG with advanced computer technique (signal pre-, and post-processing, 3D visualization of the heart sounds and murmurs, communication) combined with telemedicine application. By the cardiology point of view, the complex management of heart failure (HF) patients is essential, but time and cost consuming. The

cost of the hospitalization is the dominant component, thus every cost-efficacy/efficiency analysis is based on it. The wide range of telemedicine applications in cardiology could be found in the literature, lesser in the area of signal processing (almost all in computerized electrocardiography), and a few of using phonocardiograms [14-16]. The telemonitoring of (mostly advanced) heart failure patients is a very promising way in the complex and cost-saving care [17-19]. Screening the left ventricular dysfunction would be very important, because the heart failure is associated with high morbidity, mortality, and cost [20-22]. In this paper the cost analysis of heart failure management (screening and monitoring) with or without telemedicine application is performed.

2. Methods

In the first study 584 cardiac patients (292 for the training and 292 for the test set) were enrolled. The Doppler echocardiographic parameters (ejection fraction, aortic Vmax, the grade of mitral and tricuspid regurgitation (I-III)) were estimated from the spectral amplitude values of 170 time-frequency cells of the dPCG.

In the second study, the patients were recruited from the general population of a small town in Hungary (7400 inhabitants). The study population consists of a general population (GPg; N=192), a low-risk group (LRg; N=113), and a high-risk group (HRg; N=147). High risk patients: any of IHD, hypertension (blood pressure > 160/100), diabetes, peripheral/cerebral vascular disease, and heavy alcohol use (alcoholism or 40 units alcohol per week). Low risk patients are defined as patients without target organ damage, but are on higher cardiovascular risk using the European Risk Score (value > 5.0). Four screening strategies were compared by sensitivity and cost-effectiveness analysis: Strategy 1 (Str_1): all subjects to undergo TE (gold-standard strategy); Strategy 2 (Str_2): all subjects with an abnormal ECG were undergone TE; Strategy 3 (Str_3): all subjects with an

abnormal dPCG were undergone TE; Strategy 4 (Str_4): all subjects with an abnormal ECG and abnormal dPCG were undergone TE. In the statistical analysis the screening characteristics and cost-effectiveness were calculated in the three groups. The screening characteristics and cost-effectiveness defined as the cost per case of sHF and of dHF found. Strategy 1, or TE alone, was the gold-standard strategy. To account for multiple testing, $P < 0.01$ was taken as significant.

58 patients were enrolled in the third study, where the cost-effective strategies of telemedicine monitoring were analyzed. All of these patients had serious (NYHA III-IV) heart failure (HF). Twenty-nine patients were monitored in every 4 weeks by the telemedicine method (12-lead ECG, 30-minute ECG monitoring, dPCG; either in the GP's office, or at home), and 29 age-matched patients consist the control group.

The echocardiographic parameters were measured by the standard mode; heart failure was defined according to European guidelines. The following dPCG parameters were determined by windowed FFT analysis of dPCG (TriTest device, sampling rate 1 kHz, in the range of 20-12000 kHz): third, fourth heart sound, systolic murmur at the apex. The dPCG recordings were stepwise processed: general bandpass, adaptive by the segments of cardiac cycle, and median filtering were used.

Data were derived from the nationwide administrative dataset of the National Health Insurance Fund Administration (OEP); the cost of is calculated from the annual resource use (DRG cost-weights, heart failure cases and hospital days).

3. Results

The echocardiographic aortic Vmax (detecting systolic failure), the ejection fraction (EF), the rate of mitral and tricuspid regurgitation (MR, TR) were estimated using SPSS (V15.0) multivariate discriminant analysis (MDA) module. These output variables converted into three discrete values, and the process was the following (example of the estimation of the three grades of MR). The model (MDA, Wilks' method) chooses the best input parameters from the 170 time-frequency amplitude values from one cardiac cycle of dPCG. The Wilks's statistic selects the best parameters (found seven) and the unstandardized canonical discriminant function coefficients were determined. The linear multivariate discriminant score equation is calculated from the measured parameters, and these coefficients, either the training, or the test sets. The model chooses 17 input parameters for the Vmax, 9 for the EF, and 6 for the TR. The statistics of classification: for Vmax: Wilks's lambda: 0.111, chi-square: 274.6, degree of freedom: 34; for EF: 0.349, 154.1, 15; for MR: 0.416, 113.0, 14; for

TR: 0.535, 81.6, 12, respectively; all $p < 0.0001$.

Table 1. shows the classification results of MDA for Vmax, EF, MR, and TR.

Table 1. Classification results of the four TTE parameters. Orig%= original groups, Pred%= predicted groups.

| Orig% | Pred % | | | |
|-------|--------|------|------|------|
| | Group | 1 | 2 | 3 |
| Vmax | 1 | 97.5 | 2.5 | 0 |
| | 2 | 30.8 | 69.2 | 0 |
| | 3 | 0 | 0 | 100 |
| EF | 1 | 78.4 | 10.4 | 11.2 |
| | 2 | 29.3 | 67.9 | 4.8 |
| | 3 | 6.6 | 23.9 | 69.5 |
| MR | 1 | 85.0 | 12.4 | 2.7 |
| | 2 | 33.3 | 66.7 | 0 |
| | 3 | 14.3 | 14.3 | 71.4 |
| TR | 1 | 75.0 | 21.6 | 3.4 |
| | 2 | 38.5 | 61.5 | 0 |
| | 3 | 14.3 | 0 | 85.7 |

The prevalence of sHF was 0.34% in the LRg vs. 9.9% in the HRg ($P < 0.001$), and of dHF was 1.1% and 12.4%, respectively ($P < 0.001$). The screening characteristics of combined dECG and dPCG (sensitivity, specificity, positive and negative predictive value were: in HRg with sHF: 93.0, 68.8, 91.5, 73.3; with dHF: 82.4, 45.2, 67.3, 65.1; LRg with sHF: 74.9, 55.0, 89.7, 42.3; with dHF: 74.7, 29.4, 65.1, 29.6; in GPg with sHF: 78.5, 41.7, 64.6, 58.8; with dHF: 67.5, 38.4, 60.4, 52.2, respectively.

The unit costs were estimated at 175 € for TE, 34 for dPCG, and 15 for the dECG. The costs per case of systolic and/or diastolic HF found using the most likely current estimate of test costs. The cost per case of sHF found of Str 1. Was 8387.5 € of low-risk patients and 230.1 € of high-risk patients. The cost per case value of dHF in Str 1. were 7994.3 € in the LRg and 221.9 € in the HRg. Comparing the four strategies, the costs were (expressed in percentage of Str 1.) for Str 2.: 23.6%, Str 3.: 19.1%, Str 4.: 42.5%, Str 5.: 29.0%. The same values for dHF were in the LRg 19.4, 16.8, 38.6, 14.4; in the HRg were 21.4, 14.2, 36.7, and 16.8, respectively.

The telemedicine screening of the patients by ECG and dPCG could produce great cost-savings compared with the TE screening alone (Str4: mean: 82.4 CI-95%: 69.5-96.4 versus Str1: mean: 230.1 CI-95%: 196.5-254.4; $p < 0.001$).

In the third study, there were not any significant differences between the two groups (G2: Usual Care, G1: Telemedicine): in Group G2: age 64.8 ± 8.5 , female: 13 (44.8%), sHF: 21(72.4%), dHF: 8(27.6%), prior myocardial infarction: 19(65.5), NYHA II: 8(27.6),

NYHA III: 18(62.1), NYHA IV: 3 (10.3%), therapy with beta blocker 17(58.6%), with ACEI 24(82.7%), with ARB 5(17.2%); in Group G1: age 63.3 ± 8.1, female: 14 (48.3%), sHF: 21(72.4%), dHF: 8(27.6%), prior myocardial infarction: 17(58.6), NYHA II: 7(24.1), NYHA III: 18(62.1), NYHA IV: 4 (13.8%), therapy with beta blocker 18(62.1%), with ACEI 25(86.2%), with ARB 5(17.2%). The primary outcome measurements of the 2-years follow-up of the telemedicine study are showed in Table 2.

Table 2. Primary outcomes of the 2 years follow-up. hosp = hospitalization, days/admission = hospital duration per one admission, percent of days in hospital: days in hospital/potential days.

| Variable | Usual Care | Telemedicine |
|---------------------|------------|--------------|
| Patients | 29 | 29 |
| Patients hosp. | 17 (58.6%) | 12 (41.4%) |
| Number of hosp. | 78 | 55 |
| Days in hospital | 968 (4.5%) | 546 (2.3%) |
| Days/admission | 12.4 | 9.9 |
| Heart failure hosp. | 46 (58.9%) | 31 (56.3%) |
| Other CV hosp. | 21 (26.9%) | 15 (27.4%) |
| Non CV hosp. | 11 (14.2%) | 9 (16.3%) |
| HF hosp. days | 570 (2.7%) | 307 (1.4%) |
| HF hosp. duration | 11.8 | 8.2 |
| Deaths | 10 (34.4%) | 6 (20.6%) |
| Circulatory failure | 7 | 4 |
| Sudden death | 3 | 1 |
| Other death | 2 | 1 |

At 2 years, 79.4% of patients (23 of 29 patients) in telemedicine group (G1) were alive compared with 65.6% of those in usual care group (G2) (19 of 29 patients). A Kaplan-Meier graph indicated, that the curves began to widen after 10 months and remained divergent at 2 years (Figure 1).

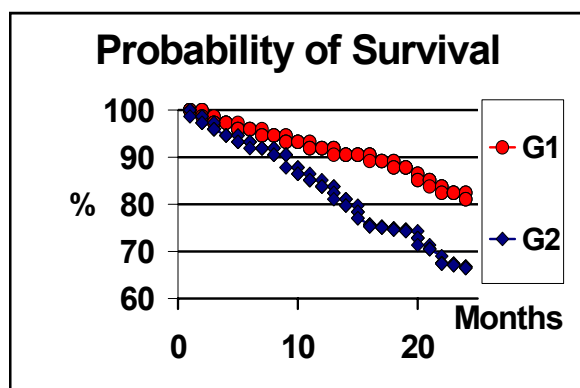


Figure 1. The probability of survival. G1= telemedicine group, G2= usual care group.

The Hungarian DGR cost-weight of one (1.0) cost-weight is 132,600 HUF (526.2 €). The DRG cost-weight of heart failure is 0.915; the normative days of HF are 9.0. Table 3. shows the data of the 2-years telemedicine follow-up cost-analysis of the 58 heart failure patients.

Table 3. Telemedicine cost-analysis. No= number, G1= telemedicine group, G2= usual care group, HC= hospital cost, C= cost, Nu=Nurse, TM= telemedicine, GP= general practitioner, Ca= cardiologist, THC= 2-years total hospital cost, HF= heart failure, NNV= number of nurse visit, NTMV= number of telemedicine visit, NGPV= number of general practitioner visit, NCV= number of cardiologist visit, NTMC= number of cardiologist visit, NTMC= number of telemedicine-driven extra cardiologist visit, TC= 2 years total cost.

| Parameter | No. | Cost HUF | Cost € |
|------------|-----|------------|---------|
| 1 day HC | | 13481 | 53.5 |
| Nu visit C | | 2200 | 8.7 |
| TM visit C | | 2700 | 10.7 |
| GP visit C | | 3600 | 14.3 |
| Ca visit C | | 7100 | 28.2 |
| THC of G1 | | 340404.5 | 29128.6 |
| THC of G2 | | 13038823.2 | 51741.4 |
| G1:HF THC | | 4532312.2 | 17985.4 |
| G2:HF THC | | 9544548 | 37875.2 |
| NNV in G1 | 743 | 1634600 | 6486.5 |
| NNV in G2 | 768 | 1689600 | 6704.8 |
| G1:NTMV | 704 | 1900800 | 7542.9 |
| NGPV in G1 | 468 | 1684800 | 6685.7 |
| NGPV in G2 | 511 | 1839600 | 7300.0 |
| NCV in G1 | 178 | 1263800 | 5015.1 |
| NCV in G2 | 143 | 1015300 | 4029.0 |
| G1:NTMC | 296 | 2101600 | 8339.7 |
| TC of G1 | | 15926004.5 | 63198.4 |
| TC of G2 | | 17583323.2 | 69775.1 |

Comparing the data of the last two rows, the cost-savings of 122.9 hospital days were found.

4. Discussion and conclusions

The first study showed that the method of multiple discriminant analysis could adequately model the time-frequency measurements of dPCG and echocardiographic data in a population of various severity of cardiac disease. The second study has shown that screening high-risk subjects is always more cost-effective than screening general population subjects and much more cost-effective than screening low-risk subjects. Using the ECG and digital phonocardiography methods to predict heart failure in various (general population subjects, high-, and low-risk patients) groups, significant cost-savings would

be achieved comparing echocardiography alone. The third study suggests that telemedicine heart failure service with digital electro- and phonocardiography, can reduce mortality substantially in patients with heart failure. The reduction in mortality is achieved with a decrease in the duration of time spent in hospital. This method a cost-effective solution for the delivery of expert care for patients with heart failure and saves approximately 120 hospital day's charge during two years follow-up of 58 patients. The three studies showed two special methods of internet use in telemedicine application: firstly, how to register dECG and dPCG data far from the cardiologist, secondly how to execute the complex math calculations using the store-and-forward method via the internet. The digital phonocardiography would be a useful method in population screening and monitoring left ventricular dysfunction. This kind of telemedicine application significantly raises the accessibility of general patient population to the adequate management of various heart diseases.

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