



- (51) International Patent Classification:
A61B 7/04 (2006.01) *H04R 1/46* (2006.01)
- (21) International Application Number:
PCT/IB2018/056336
- (22) International Filing Date:
22 August 2018 (22.08.2018)
- (25) Filing Language: English
- (26) Publication Language: English
- (30) Priority Data:
62/554,041 05 September 2017 (05.09.2017) US
15/953,502 16 April 2018 (16.04.2018) US
- (71) Applicant: **BAT CALL D. ADLER LTD.** [IL/IL]; 19
Hamesila Street, 3688519 Neshet (IL).
- (72) Inventors: **ADLER, Doron**; 8 Denia Street, Haifa (IL).
KAGAN, Igor; 12a Yakinton Street, Kiriat Bialik (IL).

SALOMON, Ezra; 22 Alon Street, Rakefet (IL). **ADLER, Omri**; 8 Denia Street, Haifa (IL). **LINHARD, David**; 15 Hanadiv Avenue, Haifa (IL). **GILAD, Hod**; 901/12 Segel Zutar Dorms, Technion City, Haifa (IL).

(74) Agent: **D. KLIGLER I. P. SERVICES LTD.**; P. O. Box 57651, 61576 Tel Aviv (IL).

(81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BN, BR, BW, BY, BZ, CA, CH, CL, CN, CO, CR, CU, CZ, DE, DJ, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IR, IS, JO, JP, KE, KG, KH, KN, KP, KR, KW, KZ, LA, LC, LK, LR, LS, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SA, SC, SD, SE, SG, SK, SL, SM, ST, SV, SY, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW.

(54) Title: DIAGNOSIS OF PATHOLOGIES USING INFRASONIC SIGNATURES

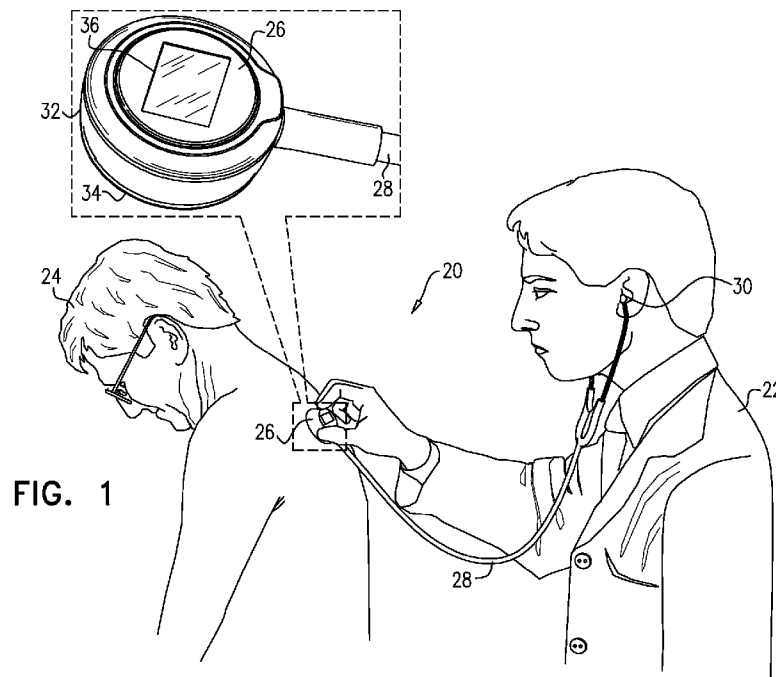


FIG. 1

(57) Abstract: A medical device (20) includes an acoustic transducer (34), which is configured to sense infrasonic waves emitted from a body of a living subject (24) with a periodicity determined by a periodic physiological activity and to output an electrical signal in response to the sensed waves. At least one speaker (30) is configured to output audible sounds in response to an electrical input. Processing circuitry (44) is configured to process the electrical signal so as to generate a frequency-stretched signal in which infrasonic frequency components of the electrical input are shifted to audible frequencies while preserving the periodicity of the periodic physiological activity in the frequency-stretched signal, and to input the frequency-stretched signal to the at least one speaker.



WO 2019/048961 A1

(84) Designated States (*unless otherwise indicated, for every kind of regional protection available*): ARIPO (BW, GH, GM, KE, LR, LS, MW, MZ, NA, RW, SD, SL, ST, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, RU, TJ, TM), European (AL, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MK, MT, NL, NO, PL, PT, RO, RS, SE, SI, SK, SM, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, KM, ML, MR, NE, SN, TD, TG).

Published:

— *with international search report (Art. 21(3))*

DIAGNOSIS OF PATHOLOGIES USING INFRASONIC SIGNATURES

FIELD OF THE INVENTION

The present invention relates generally to methods, apparatus and software for medical diagnosis, and particularly to techniques for diagnosis of medical conditions based on processing
5 of acoustic signals.

BACKGROUND

Auscultation has been a key technique in medical diagnosis for centuries. In auscultation, the medical practitioner listens to the internal sounds of the body, typically using a stethoscope. Auscultation is most commonly performed for the purpose of examining the
10 circulatory and respiratory systems, and thus diagnosing conditions of the heart and lungs in particular. In more recent years, electronic stethoscopes and methods of digital processing of body sounds have become available, in order to enhance and supplement the practitioner's auditory capabilities.

For example, U.S. Patent 5,853,005 describes an acoustic monitoring system in which a
15 transducer in communication with fluid in a pad is held in close contact against a sound or movement source and monitors acoustic signals transferred into the fluid. The signal pattern is monitored aurally and/or compared to predetermined reference patterns.

As another example, U.S. Patent 6,699,204 describes a device for analyzing auscultation sounds, in particular respiratory sounds. The device comprises an input receiving a sound signal
20 sampled in intensity levels each associated with a selected time, and storage means comprising a processing module for evaluating, in cooperation with computing means, a set of transformed intensity levels, each associated with a predetermined sound frequency. It further comprises an output connected to the storage means for delivering each transformed intensity level in
25 correspondence with an associated frequency, and means for representing intensity levels transformed on the basis of frequencies, to obtain a spectral representation of the auscultation sound.

U.S. Patent Application Publication 2005/0222515 describes methods of analyzing patient's heart, using a cardiovascular sound signature in diagnosis at the early stages of cardiac
30 dysfunctions. The invention is said to present cardiovascular sounds in time and frequency while keeping signal resolution equally strong in both directions.

SUMMARY

Embodiments of the present invention that are described hereinbelow provide methods, apparatus and software for processing and analysis of acoustic signals received from the body.

There is therefore provided, in accordance with an embodiment of the invention, a
5 medical device, including an acoustic transducer, which is configured to sense infrasonic waves emitted from a body of a living subject with a periodicity determined by a periodic physiological activity and to output an electrical signal in response to the sensed waves. At least one speaker is configured to output audible sounds in response to an electrical input. Processing circuitry is configured to process the electrical signal so as to generate a frequency-stretched signal in which
10 infrasonic frequency components of the electrical input are shifted to audible frequencies while preserving the periodicity of the periodic physiological activity in the frequency-stretched signal, and to input the frequency-stretched signal to the at least one speaker.

In some embodiments, the processing circuitry is configured to receive and digitize the electrical signal, to transform the digitized signal to a frequency domain, to shift the infrasonic
15 frequency components to the audible frequencies in the frequency domain so as to generate a frequency-shifted transformed signal, and to retransform the frequency-shifted transformed signal to a time domain in order to generate the frequency-stretched signal. In a disclosed embodiment, the processing circuitry is configured to shift the infrasonic frequency components by interpolating the transformed signal by a predefined factor, and to decimate the retransformed
20 signal in the time domain by the predefined factor in order to generate the frequency-stretched signal while preserving the periodicity of the periodic physiological activity.

In some embodiments, the device includes a user interface, which is configured to receive a user input from a user of the device indicative of a desired stretch of the infrasonic frequency components, and the processing circuitry is configured to adjust the predefined factor
25 responsively to the user input. The user interface can be configured to select a value of the predefined factor from among a first value for transformation of respiratory sounds and a second value, greater than the first value, for transformation of cardiac sounds.

In a disclosed embodiment, the infrasonic frequency components sensed by the acoustic transducer have frequencies extending at least down to 5 Hz, and the processing circuitry is
30 configured to stretch the infrasonic frequency components so that a component at 5 Hz is shifted to a frequency of at least 20 Hz.

In one embodiment, the device includes a case, which contains the acoustic transducer and at least a part of the processing circuitry, and which is configured to be brought into contact with the body of the living subject, wherein the at least one speaker includes earphones extending from the case and configured to output the audible sounds.

5 There is also provided, in accordance with an embodiment of the invention, a medical device, including an acoustic transducer, which is configured to sense acoustic waves emitted from a body of a living subject in response to a periodic physiological activity and to output an electrical signal in response to the sensed waves. Processing circuitry is configured to compute respective autocorrelations of the electrical signal for a plurality of different times within a
10 period of the physiological activity, to transform the respective autocorrelations to a frequency domain, and to render to a display, responsively to the transformed autocorrelations, a graphical representation of a spectral distribution of an energy of the acoustic waves over the period.

 In some embodiments, the acoustic waves include an infrasonic component, and the spectral distribution includes the energy at infrasonic frequencies. In one embodiment, the
15 device includes at least one speaker, which is configured to output audible sounds in response to an electrical input, wherein the processing circuitry is configured to shift the infrasonic component to audible frequencies so as to generate a frequency-stretched signal, and to generate the electrical input to the at least one speaker responsively to the frequency-stretched signal.

 In some embodiments, the physiological activity includes respiration, and the graphical
20 representation is indicative of the energy of the acoustic waves due to respiratory activity of the body. In a disclosed embodiment, the device includes a motion sensor and a case, which contains the acoustic transducer and the motion sensor and is configured to be brought into contact with the body of the living subject, wherein the processor is configured to detect a respiratory motion responsively to an output of the motion sensor, and to extract the period of the
25 respiratory activity from the detected respiratory motion.

 Typically, the graphical representation includes a plot having a frequency axis and a time axis defining a time-frequency plane and presenting a value of the energy at each point in the time-frequency plane.

 There is additionally provided, in accordance with an embodiment of the invention, a
30 method for medical diagnosis, which includes receiving an electrical signal from an acoustic transducer in response to infrasonic waves emitted from a body of a living subject with a periodicity determined by a periodic physiological activity. The electrical signal is processed so as to generate a frequency-stretched signal in which infrasonic frequency components of the

electrical input are shifted to audible frequencies while preserving the periodicity of the periodic physiological activity in the frequency-stretched signal. An electrical input to at least one speaker is generated responsively to the frequency-stretched signal, whereby the at least one speaker outputs audible sounds in response to the electrical input.

5 There is further provided, in accordance with an embodiment of the invention, a method for medical diagnosis, which includes receiving an electrical signal from an acoustic transducer in response to acoustic waves emitted from a body of a living subject due to a periodic physiological activity. Respective autocorrelations of the electrical signal are computed for a plurality of different times within a period of the physiological activity. The respective
10 autocorrelations are transformed to a frequency domain, and responsively to the transformed autocorrelations, a graphical representation of a spectral distribution of an energy of the acoustic waves over the period is rendered to a display.

The present invention will be more fully understood from the following detailed description of the embodiments thereof, taken together with the drawings in which:

15

BRIEF DESCRIPTION OF THE DRAWINGS

Fig. 1 is a schematic pictorial illustration showing a digital stethoscope in clinical use, in accordance with an embodiment of the invention;

Fig. 2 is a block diagram that schematically shows elements of a digital stethoscope, in accordance with an embodiment of the invention;

20 Fig. 3 is a flow chart that schematically illustrates a method for frequency stretching of infrasonic signals, in accordance with an embodiment of the invention;

Fig. 4 is a flow chart that schematically illustrates a method for generating an acoustic signature, in accordance with an embodiment of the invention;

25 Figs. 5A-D are plots that schematically illustrate acoustic signatures, in accordance with an embodiment of the invention;

Figs. 6A-D are plots that schematically illustrate acoustic signatures, in accordance with another embodiment of the invention; and

Figs. 7A-C are plots that schematically illustrate acoustic signatures, in accordance with yet another embodiment of the invention.

DETAILED DESCRIPTION OF EMBODIMENTS

OVERVIEW

The stethoscope is generally the first tool that the medical practitioner deploys in attempting to diagnose conditions of the circulatory and respiratory systems. Stethoscopes that
5 are known in the art, however, including electronic stethoscopes, are limited by the user's auditory capabilities and diagnostic skills. Acoustic vibrations generated by physiological activity contain a wealth of diagnostic information that is not exploited by traditional auscultation.

Embodiments of the present invention that are described herein address these limitations
10 by applying techniques of digital signal processing to acoustic waves that are emitted from the body of a living subject in response to periodic physiological activities, such as heart motion and respiration. These embodiments provide processing circuitry and methods that can be incorporated, for example, into a digital electronic stethoscope, or alternatively used in conjunction with acoustic transducers in other sorts of diagnostic devices and systems (such as
15 those described in the above-mentioned PCT Patent Application PCT/IB2017/050833). The disclosed embodiments take advantage not only of acoustic waves in the audible range (defined as the range between 20 Hz and 20,000 Hz), but also infrasonic waves, at frequencies below 20 Hz.

In some embodiments, the processing circuitry processes electrical signal output by an
20 acoustic transducer so as to generate a frequency-stretched signal, in which infrasonic frequency components of the electrical input are shifted to audible frequencies. Although the signal frequencies are shifted, however, the periodicity of the periodic physiological activity is preserved in the frequency-stretched signal. Consequently, the heart sounds repeat at the original heart rate, and breath sounds repeat at the original respiratory rate, while the infrasonic
25 components (as well as audible components) are shifted to higher tones in the audible range.

The processing circuitry inputs the frequency-stretched signal to one or more speakers, such as the earphones of an electronic stethoscope, which output the corresponding audible sounds. The medical practitioner can use such a device intuitively, as he or she would use a conventional stethoscope, but will be able to hear a wealth of additional sounds (with the
30 expected periodicity) indicative of infrasonic activity. The inventors have found that this approach enhances the practitioner's ability to diagnose cardiac and pulmonary pathologies with only a short learning curve.

In other embodiments, the processing circuitry computes and displays acoustic “signatures” based on the electrical signal output by the acoustic transducer. Such signatures may be applied in conjunction with or separately from the frequency-stretching techniques described above. Specifically, to accommodate the periodic nature of the physiological activity, the processing circuitry computes respective autocorrelations of the electrical signal at multiple different times within the period of the physiological activity. The separation between the autocorrelation peaks reflects the period, for example, the respiratory period, while the autocorrelation amplitude reflects the energy content of the signal.

The processing circuitry transforms the autocorrelation curves to the frequency domain, and renders the transformed autocorrelations in graphical form to a display. This display presents a spectral distribution of the energy of the acoustic waves over the period of the physiological activity, showing how the pattern of frequencies (including infrasonic frequencies) changes over the period, for example over the course of a respiratory cycle. The inventors have found that certain respiratory pathologies are characterized by distinct spectral patterns, referred to as signatures, some of which are illustrated in the figures that follow. Because the autocorrelation function reflects the actual, natural period of the physiological activity, as opposed to computing signal energy over some other, arbitrary window, the acoustic signature based on the autocorrelation provides a self-consistent, intuitive picture of the activity and associated pathologies.

SYSTEM DESCRIPTION

Fig. 1 is a schematic pictorial illustration showing the use of a digital stethoscope 20, in accordance with an embodiment of the invention. A practitioner 22 brings a head 26 of stethoscope 20 into contact with the body of a patient 24. Processing circuitry in head 26 outputs an audio signal via a cable 28 extending from head 26 to one or more speakers, such as earphones 30, which open apart and fit into the practitioner’s ears as would the earphones of a conventional stethoscope.

As shown in the inset, head 26 comprises a case 32, which contains an acoustic transducer, such as a suitable microphone 34, which contacts the body of patient 24 and thus senses acoustic waves emitted from the body. Microphone 34 may be of any suitable type that is known in the art, for example a piezoelectric sensor, which is sensitive not only to audible frequencies, but also to infrasonic frequencies going down to 5 Hz. On the other side of head 26, a user interface, such as a touch-sensitive display 36, enables practitioner 22 to control the

functions of stethoscope 20 and displays data, such as acoustic signatures of heart and/or breath sounds sensed by microphone 34.

Fig. 2 is a block diagram that schematically shows functional elements of stethoscope 20, in accordance with an embodiment of the invention. In the pictured embodiment, the elements shown in Fig. 2 are assumed to be contained inside head 26, within case 32. Alternatively, however, some of these components may be housed in an external processing unit.

An analog front end 40 performs analog processing functions, including filtering, buffering and amplification of the electrical signals output by microphone 34. Optionally, head 26 also comprises an inertial sensor, such as an integrated circuit accelerometer 41, which measures motion and low-frequency vibrations of head 26. Analog front end 40 may then process the signals output by the inertial sensor, as well. An analog/digital converter (ADC) 42 digitizes the acoustic and inertial signals.

A digital preprocessing circuit 44 transforms the digitized signals to the frequency domain, for example by computing a short-time Fourier transform (STFT) over successive time windows of the signals. In addition, circuit 44 can perform digital filtering functions, such as noise suppression, and “frequency stretching”: shifting infrasonic frequency components to the audible frequency range, as described further hereinbelow. Following these filtering and frequency stretching steps, circuit 44 converts the frequency-domain samples back to the time domain. For these purposes, circuit 44 typically comprises digital processing logic, which may be hard-wired or programmable; but alternatively, at least some of the functions of circuit 44 may be carried out on a programmable processor under the control of software or firmware. A digital/analog converter (DAC) 46 converts the stream of time-domain samples to analog form. An analog output circuit 48 filters and amplifies the analog signal to generate an electrical input to earphones 30.

A programmable processor 50 receives the stream of samples – in either the time domain or the frequency domain, or both – from digital preprocessing circuit 44. Processor 50 is coupled to a memory 52, which typically comprises non-volatile memory, such as flash memory, containing software or firmware to control the operations of processor 50. In addition, memory 52 typically comprises volatile random-access memory (RAM), which is used by processor 50 to store the digital samples received from circuit 44, as well as processing results.

Based on the digitized samples, processor 50 can compute acoustic signatures, representing the spectral distribution of the energy of the acoustic waves over the period of the cardiac or respiratory cycle, as described further hereinbelow. Processor 50 renders a graphical

representation of the acoustic signature to display 36 and/or outputs the signature information via a communication link (not shown). In addition, processor 50 may receive and carry out user instructions, for example in response to finger gestures on the touch screen of display 36.

The processing components shown Fig. 2, including analog front end 40, ADC 42, digital preprocessing circuit 44, DAC 46, analog output circuit 48, processor 50 and memory 52, are collectively and individually referred to herein as “processing circuitry.” These components are typically implemented in integrated circuits, as are known in the art, which are mounted together on a printed circuit board within case 32. Alternatively, other implementations of these functional components will be apparent to those skilled in the art after reading the present description and are considered to be within the scope of the present invention. Although Fig. 2 shows a certain arrangement of functional blocks for the sake of conceptual clarity, the functions of at least some of these blocks may be combined into a single integrated circuit chip or, alternatively, split among multiple chips.

FREQUENCY STRETCHING OF INFRASONIC SIGNALS

Fig. 3 is a flow chart that schematically illustrates a method for frequency stretching of infrasonic signals, in accordance with an embodiment of the invention. The method is described here, for the sake of concreteness and clarity, with reference to the components of digital stethoscope 20. As noted earlier, the object of this method is to generate a frequency-stretched signal in which infrasonic frequency components of the electrical input from microphone 34 are shifted to audible frequencies, while preserving the periodicity of periodic physiological activities (such as heartbeat and/or respiration) in the frequency-stretched signal. Digital preprocessing circuit 44 generates the frequency-stretched signal in real time, for output via earphones 30. Alternatively, the principles of the method of Fig. 3 may be implemented in substantially any system that includes a suitable acoustic transducer, speaker and processing resources.

In the method of Fig. 3, analog front end 40 receives the electrical signal output by microphone 34, and ADC 42 digitizes the signal. The time-domain digital samples are input to digital preprocessing circuit 44, which transforms the digitized signal to the frequency domain, using a short-time Fourier transform (STFT), for example. Formally, the STFT $F(m,k)$ of the time-domain signal $f[n]$, is computed by applying the formula:

$$F(m, k) = \sum_n f[n] * w[n - m] * e^{-jkn}$$

Here m and k are both discrete variables (m being the time variable and k being the frequency variable), and $w[n]$ is a suitable window function, such as a Hamming, Gaussian, or Hann window function.

Next, circuit 44 interpolates the frequency spectrum $F(m, k)$ in the frequency domain by a factor R in order to yield interpolated spectrum $F_r(m, k)$, in which the infrasonic components are shifted to audible frequencies, at an interpolation step 62. Generally speaking, R may take any rational value, but typical values of R are in the range between 2 and 20, in order to shift infrasonic frequency components to the low end of the audible range. Assuming that the infrasonic frequency components sensed by microphone 34 have frequencies extending at least down to 5 Hz, and circuit 44 will use a value of R such that the infrasonic frequency component at 5 Hz will be shifted to a frequency of at least 20 Hz.

Different values of R can be used depending upon whether practitioner 22 wishes to hear respiratory or cardiac sounds. For example, respiratory sounds typically fall within the range of 12-2000 Hz, which can be shifted to the audible range of 24-4000 Hz by setting $R=2$. As another example, heart sounds typically fall within the range of 5-200 Hz, which can be shifted to the audible range of 40-1600 Hz by setting $R=8$. Optionally, practitioner 22 may select the value of R (for example, toggling between preset values for different types of physiological activity) by entering an appropriate user input to stethoscope 20, using the touch screen of display 36, for example.

Stretching the frequency content of this acoustic signal, so as to slow down or accelerate the temporal evolution of a sound without altering its time period, requires an explicit separation of temporal and spectral information. As a result of the spectral interpolation performed at step 62, spectral changes will occur more slowly in the synthesized sound than in the input signal. Simply spacing the frequencies further apart means that phase changes now occur over a longer time interval. Hence, the periodicity of the signal will be inadvertently altered. The solution in the present embodiment is to rescale the phase by precisely the same factor by which the sound is expanded in the frequency domain. This rescaling ensures that the signal in any given filter band has the same frequency variation in the re-synthesis as in the original (though it occurs more slowly), and thus the periodicity of the sounds is preserved.

To carry out the required time domain preservation, digital preprocessing circuit 44 first computes $f_r[n]$, the inverse STFT of $F_r(m,k)$, at an inverse transformation step 64. For this purpose, circuit 44 applies the formula:

$$f_r[n] = \frac{1}{2\pi} \sum_m \sum_k F_r(m, k) * e^{jkn}$$

5

Circuit 44 then decimates the sequence of time-domain samples $f_r[n]$ by the same factor R as was used in interpolation of the frequency spectrum, at a decimation step 65. The result is a frequency-stretched signal $f_s[n]$, which has the same number of samples, and hence the same periodicity, as the original $f[n]$, but is stretched to higher frequencies. (The decimation performed at step 65 is equivalent to replaying the frequency-shifted sequence of time-domain samples $f_r[n]$ at R times the original sample rate.) DAC 46 converts $f_s[n]$ to analog form, and analog output circuit 48 amplifies and inputs this signal to earphones 30, at an audio output step 66.

15 The frequency-domain signal $F_r(m,k)$ contains the frequency-stretched spectrum in each of a succession of STFT windows. The interpolation performed at step 62 can add an incoherent phase offset as a function of frequency between successive windows. Therefore, prior to computing the inverse STFT at step 64, circuit 44 typically rotates the peak phase of each frequency component in each time window in order to match the phase in the preceding time window.

20

COMPUTATION AND DISPLAY OF ACOUSTIC SIGNATURES

Fig. 4 is a flow chart that schematically illustrates a method for generating an acoustic signature, in accordance with an embodiment of the invention. As in the preceding embodiment, the method of Fig. 4 is described here, for the sake of concreteness and clarity, with reference to the components of digital stethoscope 20; but this method may alternatively be implemented in substantially any system that includes a suitable acoustic transducer, display and processing resources.

25

The method of Fig. 4 makes use of the autocorrelation spectrum of acoustic signals received from microphone 34 over the period of a physiological activity of interest, such as the autocorrelation of heart sounds over the cardiac cycle or breath sounds over the respiratory cycle. For this purpose, it can be useful to extract the period before computing the signature. The

30

period can be extracted from the acoustic signals themselves, using methods that are known in the art. Additionally or alternatively, a motion sensor within case 32, such as accelerometer 41, can be used to sense the respiratory motion of the thorax of patient 24 while head 26 is in contact with the thorax. Processor 50 detects the respiratory motion using the output of the motion sensor and thus extracts the period of the patient's respiratory activity.

The period of the activity, based on the digitized signal $x(n)$ from microphone 34 or accelerometer 41, can itself be computed using the autocorrelation function:

$$y(k) = \frac{\sum_n x(n)x(n+k)}{\sum_n x^2(n)}$$

Processor 50 identifies significant peaks in the autocorrelation $y(k)$, for example, peaks whose amplitudes exceed a certain threshold and are separated by at least a minimum amount of time, such as 0.5 sec for cardiac activity. The time between each pair of successive peaks is taken to be a candidate period. Processor 50 discards outlying values of the candidate period and averages the remaining values to find the actual period for purposes of computing the acoustic signature. Alternatively or additionally, a k-nearest neighbor (KNN) approach can be used in classifying successive periods of the autocorrelation function and thus measure the period accurately.

As noted earlier, this method of identifying the time period of the respiratory activity can be used together with an additional sensor, such as a motion sensor to detect the chest motion.

In the description that follows, the method of Fig. 4 is applied to chest sounds (including infrasonic components) due to respiratory activity, and the resulting acoustic signatures are thus indicative of normal or pathological respiratory patterns. In alternative embodiments, not shown in the figures, the principles of this method can be applied, *mutatis mutandis*, to cardiac activity. To compute the acoustic signature of the respiratory activity, processor 50 builds a two-dimensional (2D) autocorrelation matrix based on the digitized samples $x(n)$ of the electrical signals from microphone 34, at a matrix construction step 70. The elements of the autocorrelation matrix $y(n,l)$ have the form:

$$y(n,l) = x(n+l)x^*(n-l)$$

In other words, processor 50 computes the autocorrelation as a function of a variable time shift l for each time n within a period of the signal.

Processor 50 transforms the 2D autocorrelation matrix to a matrix representation of the spectral (frequency) distribution of the energy of the acoustic waves over the respiratory period, at a frequency transformation step 72. This step can be carried out by calculating a one-dimensional fast Fourier transform (FFT) over the l -dimension of the autocorrelation matrix, which yields the time/frequency chest sounds energy distribution (CSED):

$$CSED(n, k) = \sum_{l=-N/2}^{\frac{N}{2}-1} y(n, l) e^{-2\pi i k l / N}$$

10

Here N is the number of samples in a respiratory period. Because it is tied to the actual period of respiration, the CSED provides a consistent representation of the spectral distribution of the energy of the acoustic waves, notwithstanding shifts in time and frequency.

Processor 50 outputs the CSED for used by practitioner 22 and/or further offline recording and analysis, at a CSED output step 74. For example, processor 50 may render a graphical representation of the CSED to display 36 or to a separate, external display (not shown). Typically, this graphical representation comprises a plot having a frequency axis and a time axis, which together define a time-frequency plane. The value of the energy is presented in the plot at each point in the time-frequency plane. A number of modes of display of this sort are shown in the figures that follow.

Figs. 5A-D are plots that schematically illustrate acoustic signatures, in accordance with an embodiment of the invention. In these pseudo-3D plots, the time-frequency plane is horizontal, and the energy is represented by peaks extending above and below the plane. The time axis is marked in seconds, reflecting the length of an average respiratory period, while the frequency axis is marked in Hertz. The energy scale is relative.

Fig. 5A shows the CSED of normal breath sounds, which is characterized by a sharp ridge 86 of frequencies, in the range of 50-100 Hz, extending continuously in time over the central part of the respiratory cycle. By contrast, Fig. 5B shows a CSED with a wider, more extended ridge 80, reflecting wheezes in the breath sounds of a patient suffering from bronchiectasis due to cystic fibrosis.

Figs. 5C and 5D illustrate the effect of crackles in the breath sounds of patients with bronchiectasis and pulmonary edema, respectively. The crackles are reflected in the CSED by ridges 82 and 84 covering wide ranges of frequencies, repeating multiple times over the course of the respiratory cycle.

5 Figs. 6A-D are plots that schematically illustrate acoustic signatures, in accordance with another embodiment of the invention. These plots show the same CSED computations, for the same patients, as in Figs. 5A-D, respectively. In this embodiment, however, the energy as a function of time and frequency is shown as an intensity value, with darker shades of gray corresponding to higher energy values.

10 Figs. 7A-C are plots that schematically illustrate acoustic signatures, in accordance with yet another embodiment of the invention. These plots similarly show energy as an intensity value over the time-frequency plane, except that in this embodiment, the frequency scale is logarithmic.

Fig. 7A shows the company energy spectrum of a normal patient with healthy lungs. By contrast, Fig. 7B shows the CSED computed over the breath sounds of a patient with chronic obstructive pulmonary disease (COPD), with frequency bands extending down to the infrasonic range over the duration of the respiratory cycle. As another example, Fig. 3C shows the CSED of a patient suffering from idiopathic pulmonary fibrosis (IPF), in which crackles are represented by vertical lines in the low-frequency range, again extending down to infrasonic frequencies.

20 The distinctions between normal respiration and various, specific sorts of pathologies are visually apparent in the acoustic signatures shown above. These acoustic signatures can thus assist the practitioner in making fast, accurate diagnoses and assessing changes in the patient's condition thereafter.

It will be appreciated that the embodiments described above are cited by way of example, and that the present invention is not limited to what has been particularly shown and described hereinabove. Rather, the scope of the present invention includes both combinations and subcombinations of the various features described hereinabove, as well as variations and modifications thereof which would occur to persons skilled in the art upon reading the foregoing description and which are not disclosed in the prior art.

30

CLAIMS

1. A medical device, comprising:
 - an acoustic transducer, which is configured to sense infrasonic waves emitted from a body of a living subject with a periodicity determined by a periodic physiological activity and to
 - 5 output an electrical signal in response to the sensed waves;
 - at least one speaker, which is configured to output audible sounds in response to an electrical input; and
 - processing circuitry, which is configured to process the electrical signal so as to generate a frequency-stretched signal in which infrasonic frequency components of the electrical input are
 - 10 shifted to audible frequencies while preserving the periodicity of the periodic physiological activity in the frequency-stretched signal, and to input the frequency-stretched signal to the at least one speaker.
2. The device according to claim 1, wherein the processing circuitry is configured to receive and digitize the electrical signal, to transform the digitized signal to a frequency domain, to shift
- 15 the infrasonic frequency components to the audible frequencies in the frequency domain so as to generate a frequency-shifted transformed signal, and to retransform the frequency-shifted transformed signal to a time domain in order to generate the frequency-stretched signal.
3. The device according to claim 2, wherein the processing circuitry is configured to shift the infrasonic frequency components by interpolating the transformed signal by a predefined
- 20 factor, and to decimate the retransformed signal in the time domain by the predefined factor in order to generate the frequency-stretched signal while preserving the periodicity of the periodic physiological activity.
4. The device according to claim 3, and comprising a user interface, which is configured to receive a user input from a user of the device indicative of a desired stretch of the infrasonic
- 25 frequency components, and wherein the processing circuitry is configured to adjust the predefined factor responsively to the user input.
5. The device according to claim 4, wherein the user interface is configured to select a value of the predefined factor from among a first value for transformation of respiratory sounds and a second value, greater than the first value, for transformation of cardiac sounds.
- 30 6. The device according to any of claims 1-5, wherein the infrasonic frequency components sensed by the acoustic transducer have frequencies extending at least down to 5 Hz, and wherein

the processing circuitry is configured to stretch the infrasonic frequency components so that a component at 5 Hz is shifted to a frequency of at least 20 Hz.

7. The device according to any of claims 1-5, and comprising a case, which contains the acoustic transducer and at least a part of the processing circuitry, and which is configured to be brought into contact with the body of the living subject,

wherein the at least one speaker comprises earphones extending from the case and configured to output the audible sounds.

8. A medical device, comprising:

an acoustic transducer, which is configured to sense acoustic waves emitted from a body of a living subject in response to a periodic physiological activity and to output an electrical signal in response to the sensed waves;

a display; and

processing circuitry, which is configured to compute respective autocorrelations of the electrical signal for a plurality of different times within a period of the physiological activity, to transform the respective autocorrelations to a frequency domain, and to render to the display, responsively to the transformed autocorrelations, a graphical representation of a spectral distribution of an energy of the acoustic waves over the period.

9. The device according to claim 8, wherein the acoustic waves comprise an infrasonic component, and wherein the spectral distribution includes the energy at infrasonic frequencies.

10. The device according to claim 9, and comprising at least one speaker, which is configured to output audible sounds in response to an electrical input, wherein the processing circuitry is configured to shift the infrasonic component to audible frequencies so as to generate a frequency-stretched signal, and to generate the electrical input to the at least one speaker responsively to the frequency-stretched signal.

11. The device according to any of claims 8-10, wherein the physiological activity comprises respiration, and wherein the graphical representation is indicative of the energy of the acoustic waves due to respiratory activity of the body.

12. The device according to claim 11, and comprising a motion sensor and a case, which contains the acoustic transducer and the motion sensor and is configured to be brought into contact with the body of the living subject, wherein the processor is configured to detect a

respiratory motion responsively to an output of the motion sensor, and to extract the period of the respiratory activity from the detected respiratory motion.

13. The device according to any of claims 8-10, wherein the graphical representation comprises a plot having a frequency axis and a time axis defining a time-frequency plane and presenting a value of the energy at each point in the time-frequency plane.

14. A method for medical diagnosis, comprising:
receiving an electrical signal from an acoustic transducer in response to infrasonic waves emitted from a body of a living subject with a periodicity determined by a periodic physiological activity;
10 processing the electrical signal so as to generate a frequency-stretched signal in which infrasonic frequency components of the electrical input are shifted to audible frequencies while preserving the periodicity of the periodic physiological activity in the frequency-stretched signal; and
generating an electrical input to at least one speaker responsively to the frequency-
15 stretched signal, whereby the at least one speaker outputs audible sounds in response to the electrical input.

15. The method according to claim 14, wherein processing the electrical signal comprises:
digitizing the electrical signal;
transforming the digitized signal to a frequency domain;
20 shifting the infrasonic frequency components to the audible frequencies in the frequency domain so as to generate a frequency-shifted transformed signal; and
retransforming the frequency-shifted transformed signal to a time domain in order to generate the frequency-stretched signal.

16. The method according to claim 15, wherein shifting the infrasonic frequency components comprises interpolating the transformed signal by a predefined factor, and preserving the periodicity comprises decimating the retransformed signal in the time domain by the predefined factor.

17. The method according to claim 16, and comprising receiving a user input indicative of a desired stretch of the infrasonic frequency components, and adjusting the predefined factor
30 responsively to the user input.

18. The method according to claim 17, wherein receiving the user input comprises selecting a value of the predefined factor from among a first value for transformation of respiratory sounds and a second value, greater than the first value, for transformation of cardiac sounds.
19. The method according to any of claims 14-18, wherein the infrasonic frequency components sensed by the acoustic transducer have frequencies extending at least down to 5 Hz, and wherein processing the electrical signal comprises stretching the infrasonic frequency components so that a component at 5 Hz is shifted to a frequency of at least 20 Hz.
20. A method for medical diagnosis, comprising:
receiving an electrical signal from an acoustic transducer in response to acoustic waves emitted from a body of a living subject due to a periodic physiological activity;
computing respective autocorrelations of the electrical signal for a plurality of different times within a period of the physiological activity;
transforming the respective autocorrelations to a frequency domain; and
rendering to a display, responsively to the transformed autocorrelations, a graphical representation of a spectral distribution of an energy of the acoustic waves over the period.
21. The method according to claim 20, wherein the acoustic waves comprise an infrasonic component, and wherein the spectral distribution includes the energy at infrasonic frequencies.
22. The method according to claim 21, and comprising stretching the infrasonic component to audible frequencies so as to generate a frequency-stretched signal, and generating an electrical input to at least one speaker responsively to the frequency-stretched signal, whereby the at least one speaker outputs audible sounds in response to the electrical input.
23. The method according to claim 20, wherein the physiological activity comprises respiration, and wherein the graphical representation is indicative of the energy of the acoustic waves due to respiratory activity of the body.
24. The method according to claim 23, and comprising detecting a respiratory motion of the body using a motion sensor in contact with the body, and extracting the period of the respiratory activity from the detected respiratory motion.
25. The method according to any of claims 20-24, wherein the graphical representation comprises a plot having a frequency axis and a time axis defining a time-frequency plane and presenting a value of the energy at each point in the time-frequency plane.

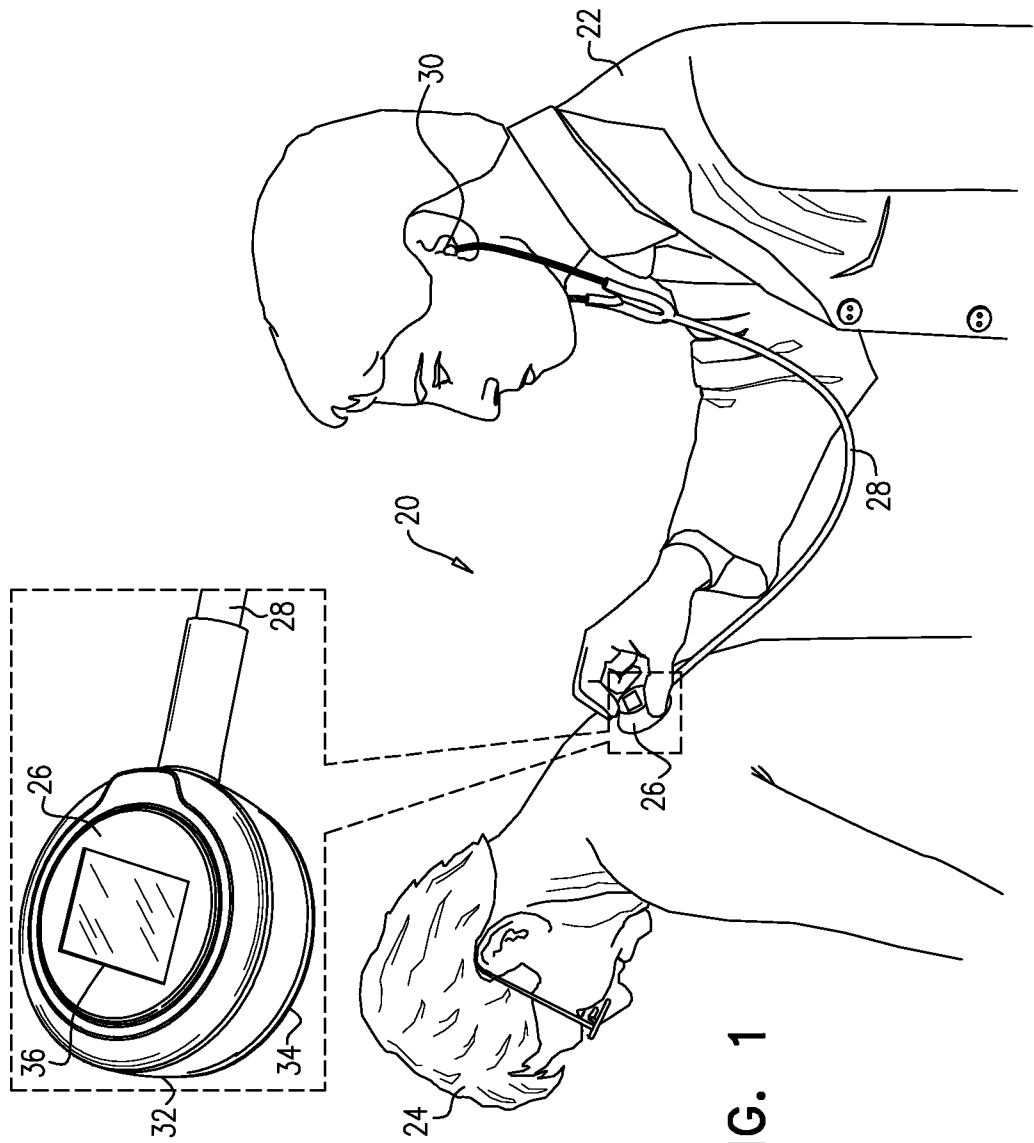


FIG. 1

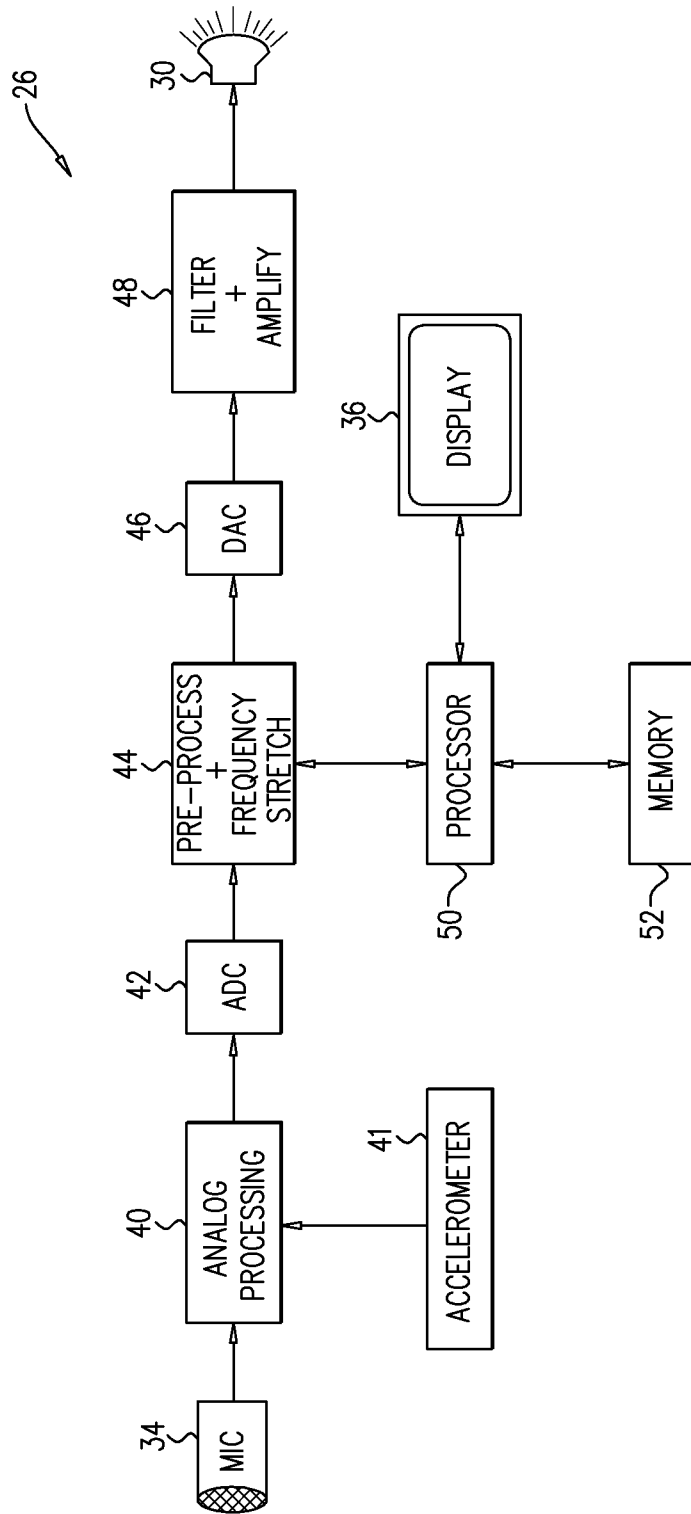


FIG. 2

3/10

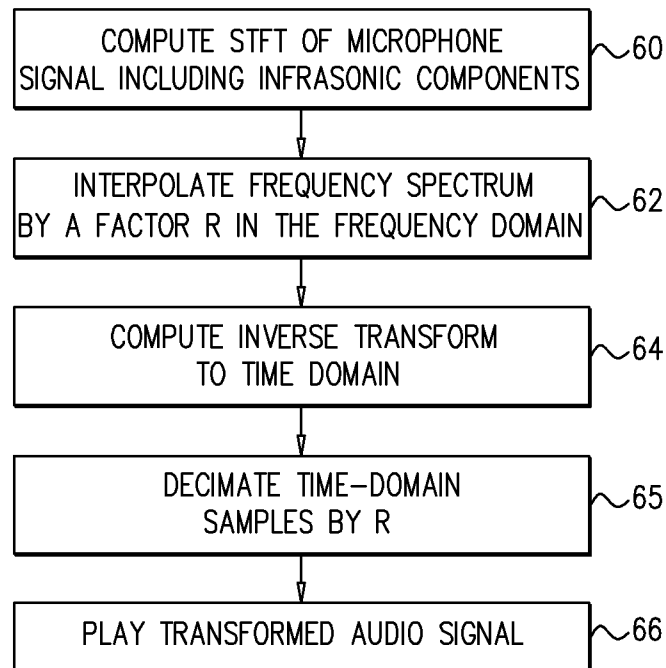


FIG. 3

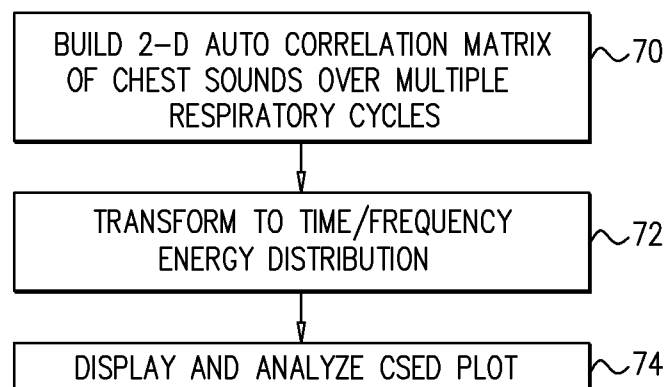


FIG. 4

4/10

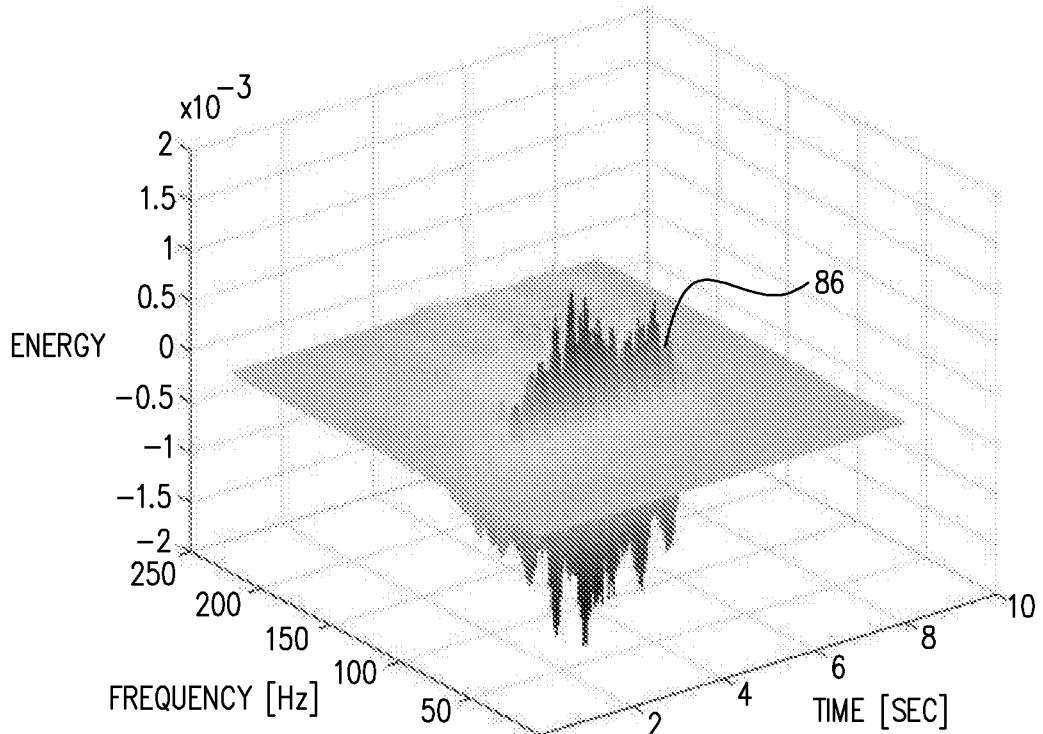


FIG. 5A

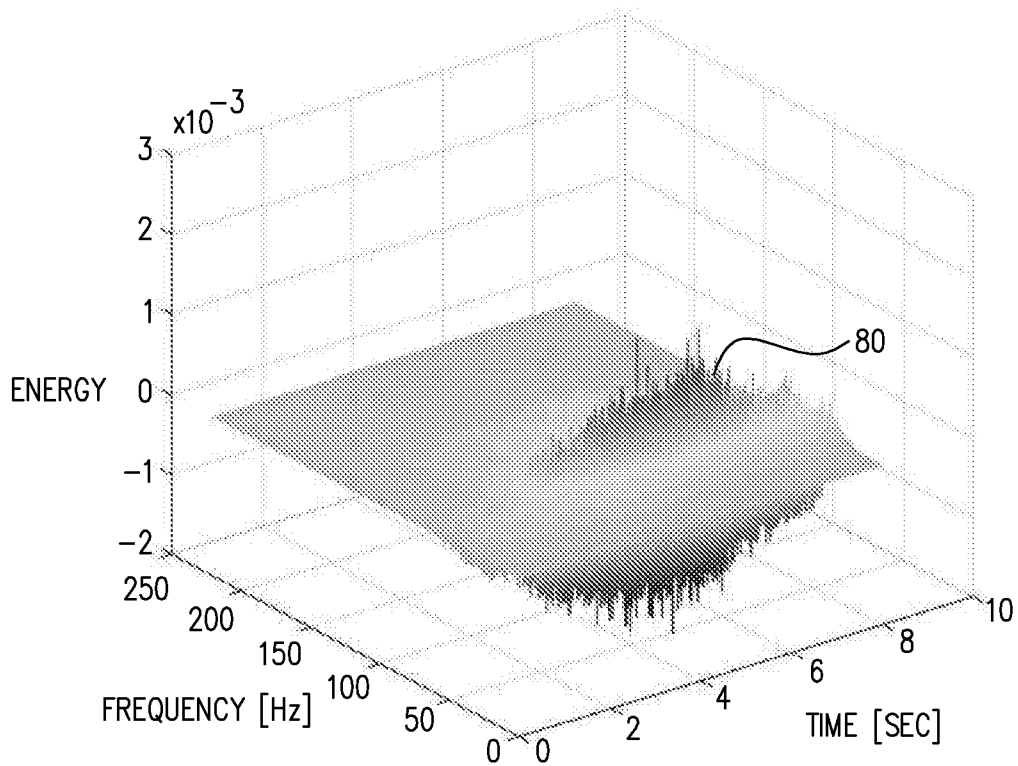


FIG. 5B

5/10

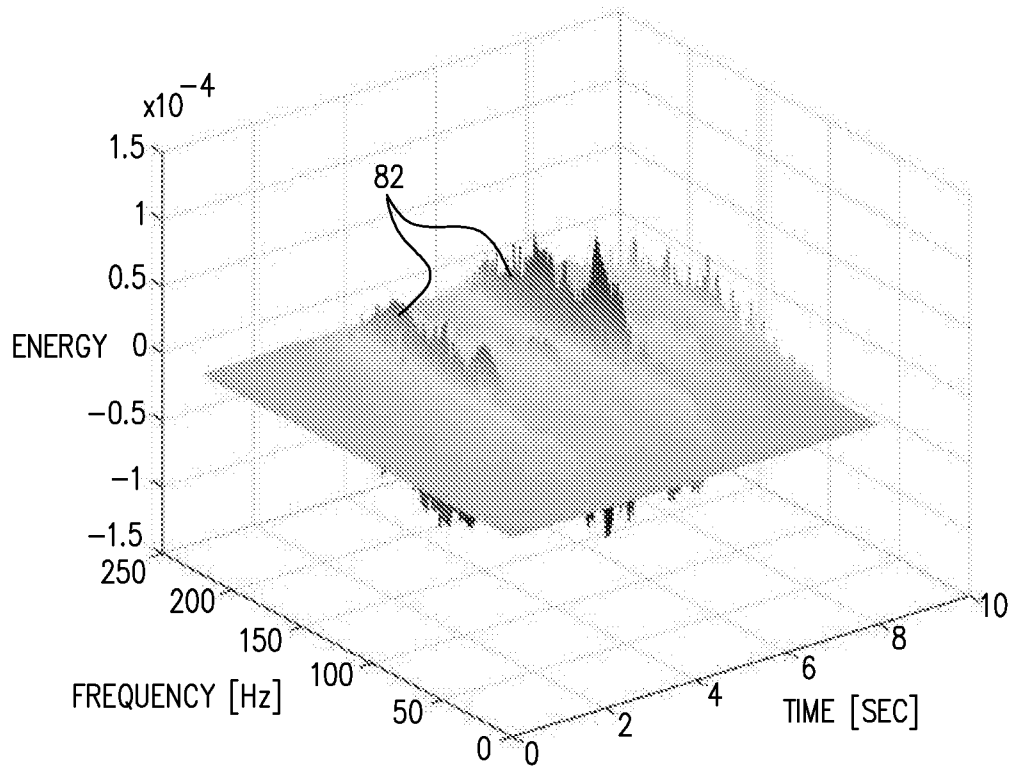


FIG. 5C

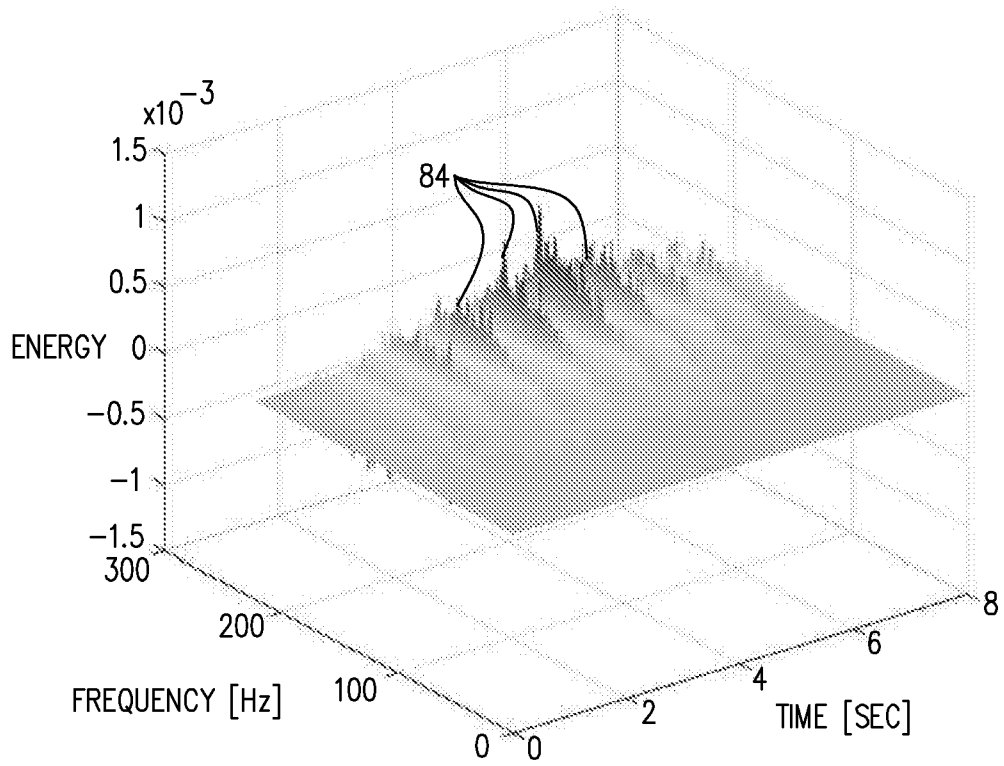


FIG. 5D

6/10

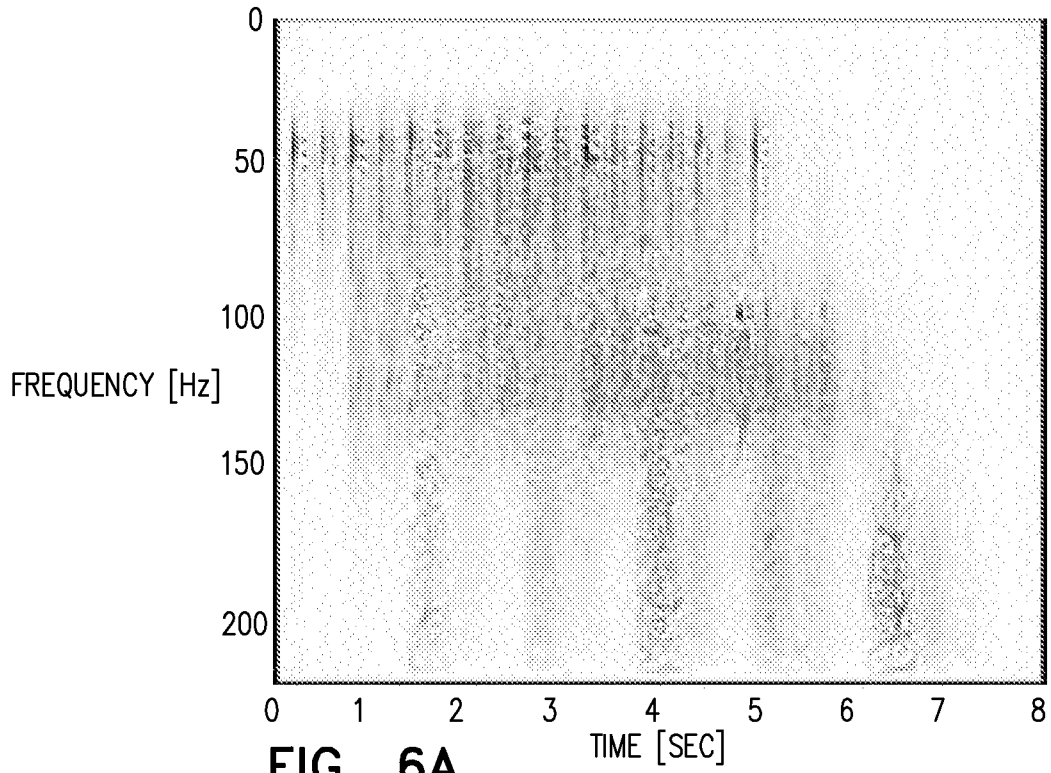


FIG. 6A

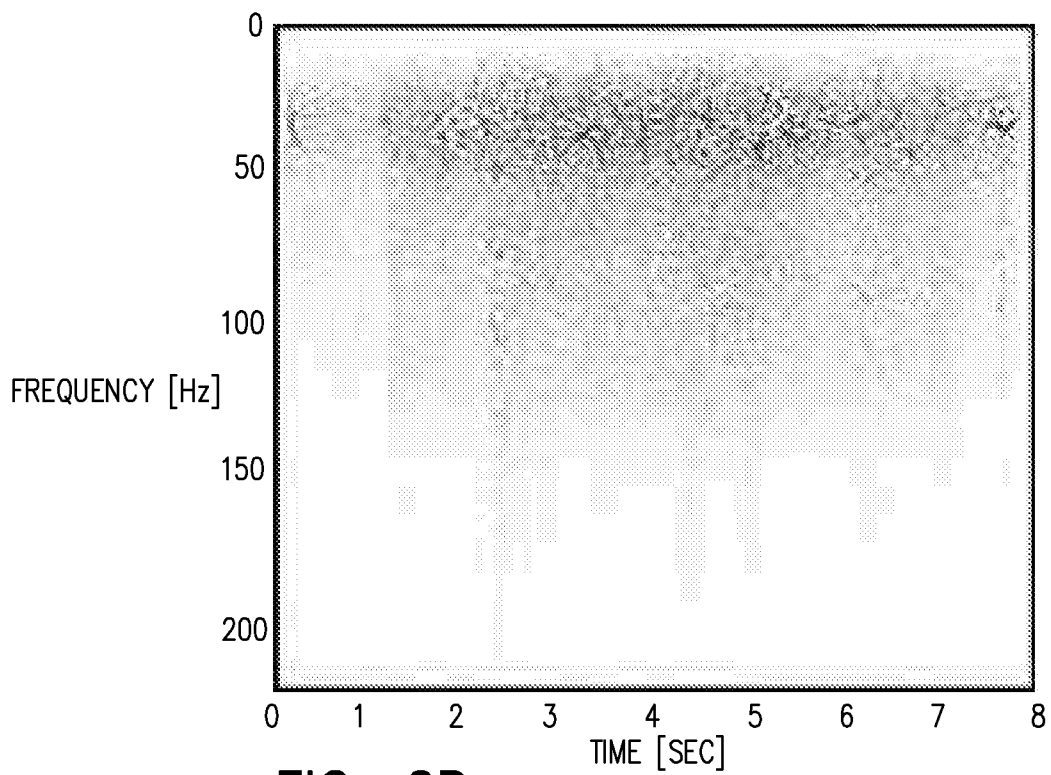
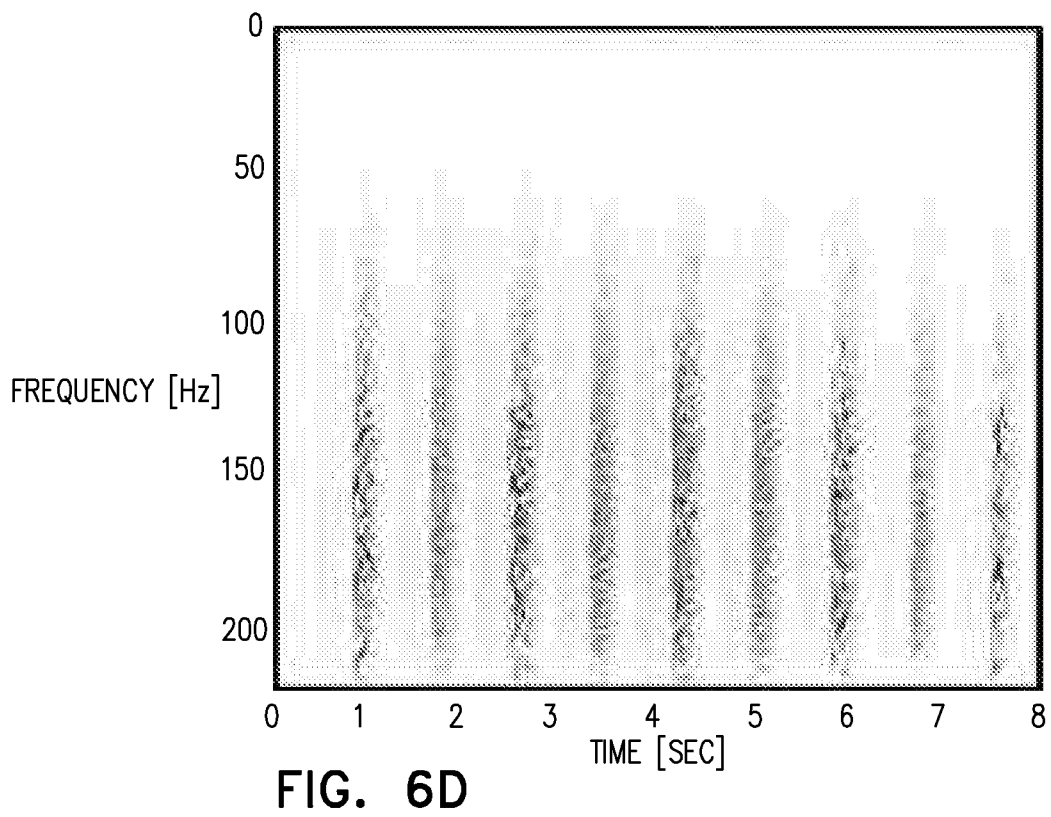
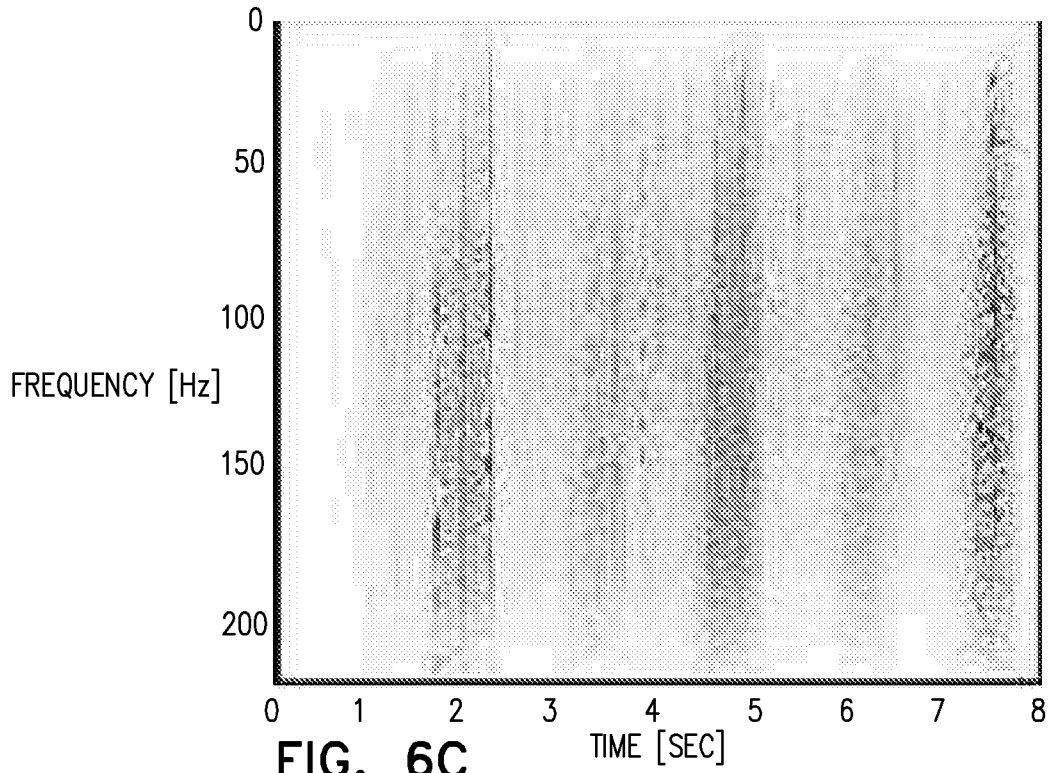


FIG. 6B

7/10



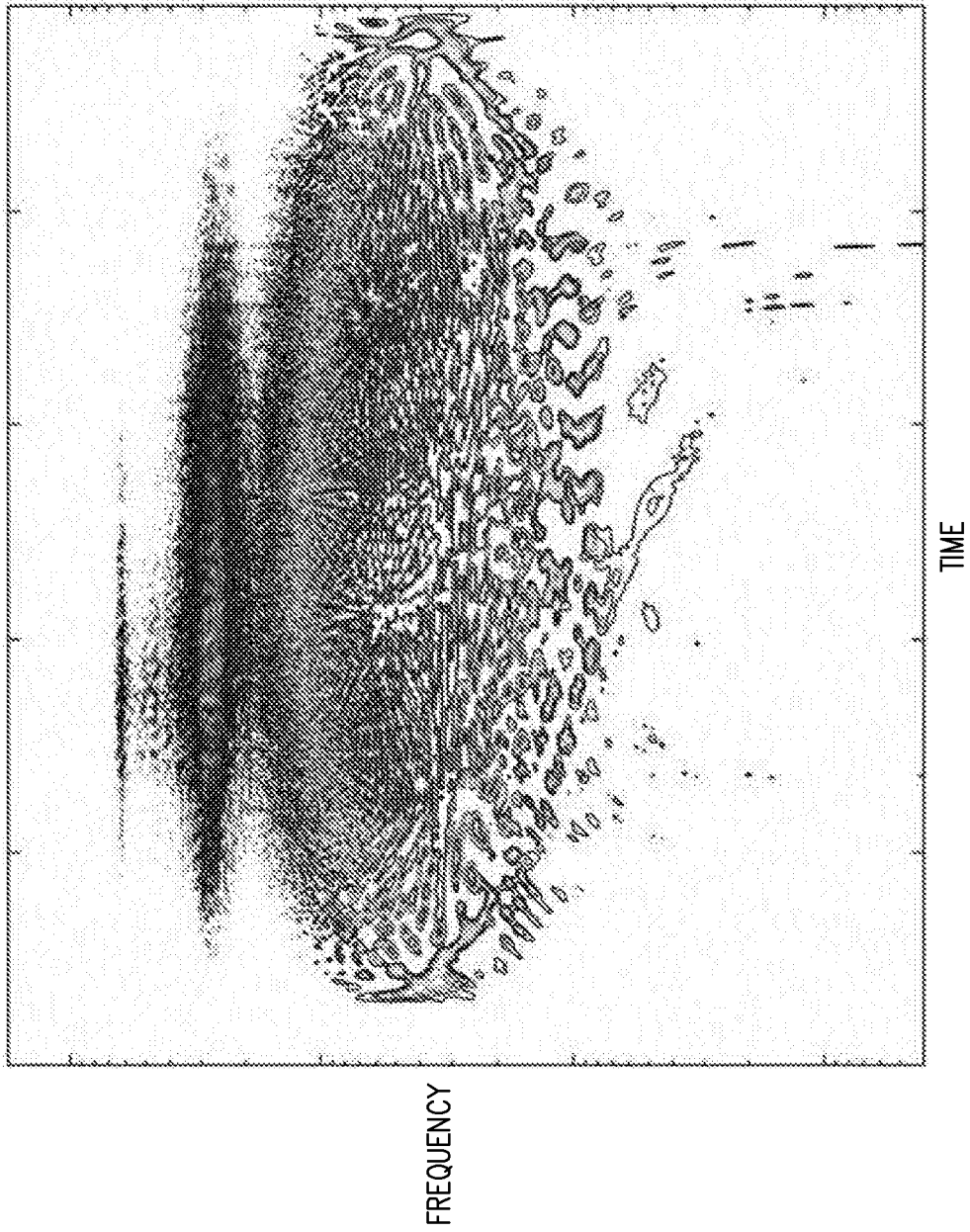


FIG. 7A

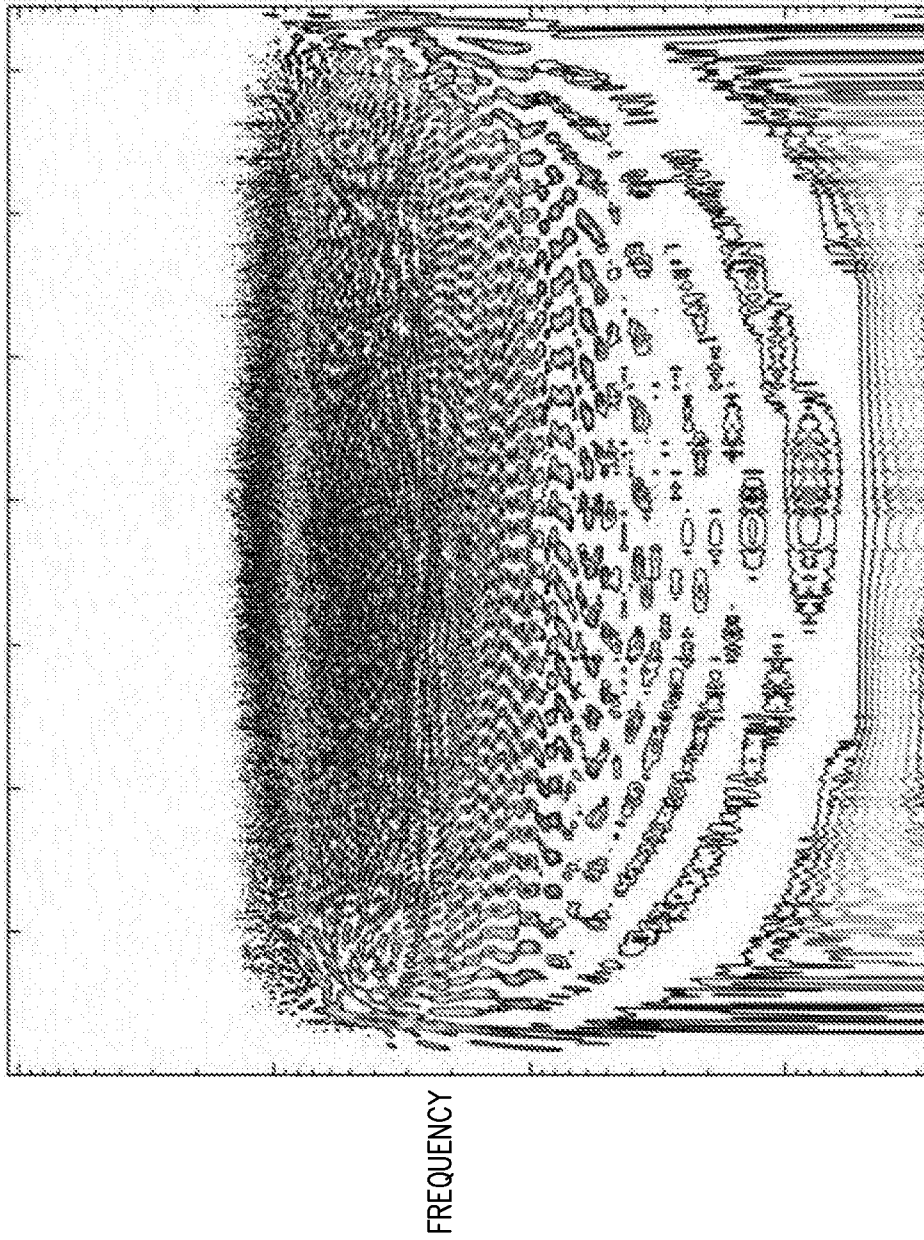
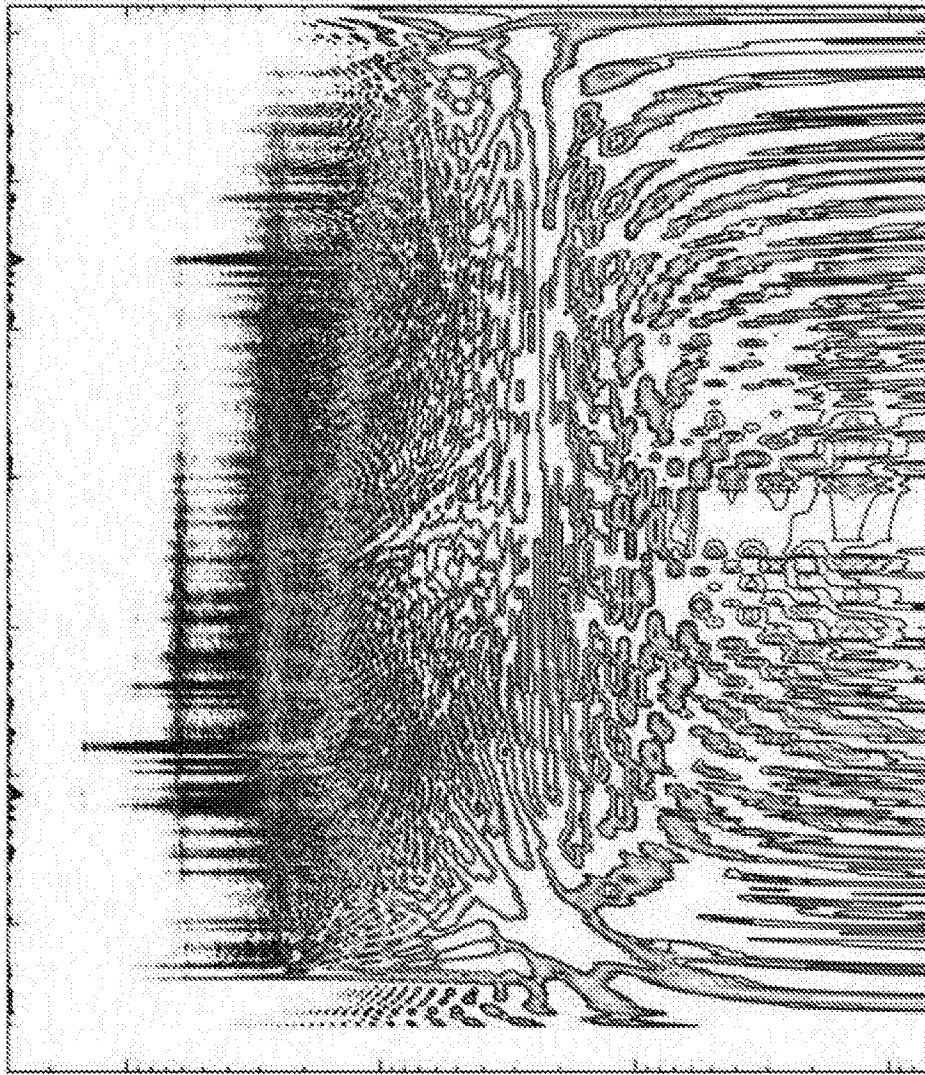


FIG. 7B



TIME

FIG. 7C

FREQUENCY

INTERNATIONAL SEARCH REPORT

International application No.

PCT/IB2018/056336

A. CLASSIFICATION OF SUBJECT MATTER IPC (2018.01) A61B 7/04, H04R 1/46 According to International Patent Classification (IPC) or to both national classification and IPC		
B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) IPC (2018.01) A61B 7/04, H04R 1/46 Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) Databases consulted: Google Patents, Google Scholar, Orbit Search terms used: H04R2430/20 A61B7/04 infrasonic STETHOSCOPE		
C. DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 2017141165 A1 Bat Call D. Adler LTD. 24 Aug 2017 (2017/08/24) fig. 1, 5, 6 , page 12 lines 19-24 page 21 line 5 – page 22 line 27	1-7,12-16
X	US 2014073864 A1 Nellcor Puritan Bennett LLC 13 Mar 2014 (2014/03/13) fig. 1, 2, 4-88, 53, para. 6, 9, 16, 20, 111, 146	8-11,17-25
P,A	US 2018228468 A1 Bat Call D . Adler Ltd . 16 Aug 2018 (2018/08/16) whole doc.	1-25
<input type="checkbox"/> Further documents are listed in the continuation of Box C. <input checked="" type="checkbox"/> See patent family annex.		
* Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier application or patent but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art "&" document member of the same patent family		
Date of the actual completion of the international search 24 Dec 2018		Date of mailing of the international search report 25 Dec 2018
Name and mailing address of the ISA: Israel Patent Office Technology Park, Bldg.5, Malcha, Jerusalem, 9695101, Israel Facsimile No. 972-2-5651616		Authorized officer SHUSHAN Hadas HadasShu@justice.gov.il Telephone No. 972-2-5657813

INTERNATIONAL SEARCH REPORT
Information on patent family members

International application No.
PCT/IB2018/056336

Patent document cited search report	Publication date	Patent family member(s)	Publication Date
WO 2017141165 A1	24 Aug 2017	WO 2017141165 A1 US 2018228468 A1	24 Aug 2017 16 Aug 2018
US 2014073864 A1	13 Mar 2014	US 2014073864 A1	13 Mar 2014
US 2018228468 A1	16 Aug 2018	US 2018228468 A1 WO 2017141165 A1	16 Aug 2018 24 Aug 2017