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A Scalable Healthcare Integrated Platform (SHIP) and Key Technologies for Daily Application

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1. Introduction

A ubiquitous era has arisen based on achievements from the development of science and technology over the previous 1,000 years, and especially the past 150 years. Among the numerous accomplishments in human history, four fundamental technologies have laid the foundation for today's pervasive computing environment.

The electromagnetic wave theory, established by James Maxwell in 1864, predicted the existence of waves of oscillating electric and magnetic fields that travel through empty space at a velocity of 310,740,000 m/s. His quantitative connection between light and electromagnetism is considered one of the great triumphs of 19th century physics. Twenty years later, through experimentation, Heinrich Hertz proved that transverse free space electromagnetic waves can travel over some distance, and in 1888, he demonstrated that the velocity of radio waves was equal to the velocity of light. However, Hertz did not realize the practical importance of his experiments. He stated that, "It's of no use whatsoever … this is just an experiment that proves Maestro Maxwell was right – we just have these mysterious electromagnetic waves that we cannot see with the naked eye. But they are there." His discoveries were later utilized in wireless telegraphy by Guglielmo Marconi, and they formed a part of the new "radio communication age".

The second fundamental technology is spread-spectrum telecommunications, whose multiple access capability allows a large volume of users to communicate simultaneously on the same frequency band, as long as they use different spreading codes. This has been developed since the 1940s and used in military communication systems since the 1950s. Realization of spread-spectrum technology requires a large computational capacity and leads to a bulky size and weight. Since the initial commercial use of spread spectrum telecommunications began in the 1980s, it is now widely used in many familiar systems today, such as GPS, Wi-Fi, Bluetooth, and mobile phones. This was made possible by the invention of computing machines and integrated circuits, the third and fourth tremendous triumphs.

The first automatic computing machine, known as ENIAC, was built using 18,000 vacuum tubes, 1,500 relays, 70,000 resistors, and 10,000 condensers. It performed 35,000 additions per

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second and cost US\$487,000 (Nohzawa, 2003). The latest CPU, the Yorkfield XE, contains 820 million transistors on 2×107 mm² dies and features a 1333 MT/s FSB and a clock speed of 3 GHz (Intel Corp., 2007).

Since the first integrated circuit (IC), which contained a single transistor and several resistors on an $11 \times 1.6 \text{ mm}^2$ germanium chip, was fabricated by Jack Kilby in 1958, advanced 45 nm semiconductor technology makes it possible to condense an entire complicated spread-spectrum telecommunication system into a magic box as small as a mobile phone.

Today, we are interconnected through wired and wireless networks, and surrounded by an invisible pervasive computing environment. This makes "information at your fingertips" and "commerce at light speed" possible. We are already acclimatized to enjoy everything worldwide conveniently, wherever we are. We enjoy online shopping and share information with friends from the other side of the Earth in an instant.

However, this is a double-edged sword. Our daily lifestyle has changed dramatically. While we may benefit from the advantages of today's society, at the same time, we face many unprecedented problems in the health domain, which have emerged with all of these changes.

One of the greatest concerns is the ascent of chronic illness that has occurred concurrently with the accompanying lifestyle changes. Figure 1 shows the change in mortality among different diseases from acute to chronic over the past 100 years in Japan. There has not been a large change in conventional causes of death, such as contingency, caducity, and pneumonia. Acute infectious diseases, such as tuberculosis, have disappeared completely since the 1980s. However, death due to chronic conditions is increasing. The leading causes of death are the three "C" top killer diseases: cerebral, cardiovascular, and cancer (malignant neoplasm), which account for 60 per cent of total deaths.

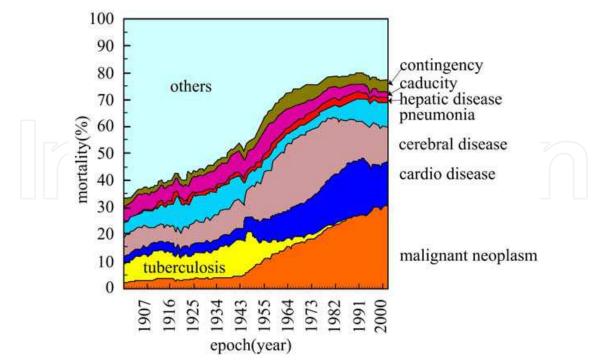


Fig. 1. Change in mortality of different diseases over the previous century in Japan. (Adapted from the Japanese Ministry of Health, Labour, and Welfare).

Rapid changes in both societal environment and daily lifestyle are responsible for most of these chronic illnesses. Treatment of chronic conditions is now recognized as a problem of all society and no longer just a private issue. To elevate all citizens' awareness of the importance of health promotion and disease prevention in response to the steep increase in long-term healthcare requirements, it is indispensable to be involved in every aspect and to take creative action. A variety of innovative strategies and activities are now being explored nationwide in Japan at three levels: macro (administration), meso (community, organization, and company), and micro (personal) levels.

At the macro level, a 12-year health promotion campaign, known as "Healthy Japan 21" (Japan Health Promotion and Fitness Foundation, 2000), has been advocated nationwide since 2000 and is financially supported by the Japanese Ministry of Health, Labour, and Welfare. Furthermore, a "health promotion law" (Japanese Ministry of Health, Labour and Welfare, 2002) was issued by the Japanese parliament. This reconfirmed that the national goal of medical insurance reconstruction was health promotion and disease prevention, and it defined individual responsibility and coordination among citizens, community, and government organizations.

At the meso level, industrial organizations and research institutions have developed many Internet-based systems and related devices for daily healthcare. Professional organizations and academic associations have established a series of educational programs and accreditation systems for professional healthcare promoters. Citizen communities have boosted health promotion campaigns through various service options.

At the micro level, more and more people are aware of the importance of health promotion and chronic prevention, and are becoming more active in participating in daily personal healthcare practices. They spend a lot of time and money on exercise, diet, and regular medical examinations to keep their biochemical indices as good as possible.

This trend turns out that in the US only, the healthcare domain is now growing up into a giant industrial territory worthy of about US\$2 trillion annually (MarketResearch.com. 2008). In terms of building a better healthcare environment, and as one of the initiatives in the arena of human welfare in long-term chronic treatment, we are confronting the challenges of providing effective means for vital sign monitoring technologies suitable for daily use, and large-scale data mining and a comprehensive interpretation of their physiological interconnection. These solutions are being developed across the world. Many companies are already engaged in and placing priority on, providing a total solution to these ever-increasing demands.

The "Health Data Bank" ASP service platform was released as a multifaceted aid for the health management of corporate employee medical exam results (NTT Data Corp., 2002). The service supplies healthcare personnel with a set of tools for effective employee health guidance and counselling, and takes into account factors such as an employee's current physical condition, as well as living habits and environment, and age-related changes in longitudinal management in accumulated individual data. Individual corporate employees can browse their personal data through Internet channels, and view records of their check-ups, as well as graphs detailing historical changes in their health condition, thus facilitating improved personal health management.

Companies and research institutes in the European Union have launched several multinational projects to develop wearable and portable healthcare systems for personalized care. The "MyHeart" project is a framework for personal healthcare applications led by Philips, which aims to develop on-body sensors/electronics and appropriate services to help

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fight cardiovascular disease through prevention and early diagnosis. It can monitor vital signs and physical movement via wearable textile technology, process the measured data, and provide the user with recommendations (Philips Electronics, 2004). After the completion of the "MyHeart" program, a continuation "HeartCycle" project began in March 2008. Many new sensors and key technologies, such as a cuff-less blood pressure sensor, a wearable SpO2 sensor, an inductive impedance sensor, an electronic acupuncture system, a contactless ECG, arrays of electret foils, a motion-compensation system for ECG, and a cardiac performance monitor (from bioimpedance) will be developed and built into the system. A patient's condition will be monitored using a combination of unobtrusive sensors built into the patient's clothing or bed sheets and home appliances, such as weighing scales and blood pressure meters. Data mining and decision support approaches will be developed to analyse the acquired data, to predict the short-term and long-term effects of lifestyle and medication, and to obtain an objective indicator of patient compliance (Philips Electronics, 2008).

"MobiHealth" was a mobile healthcare project funded by the European Commission from 2002 to 2004. Fourteen partners from hospitals and medical service providers, universities, mobile network operators, mobile application service providers, mobile infrastructure, and hardware suppliers across five European countries participated in the project. It allowed patients to be fully mobile while undergoing health monitoring without much discomfort in daily activities. The patients wore a lightweight unit with multiple sensors connected via a Body Area Network (BAN) for monitoring ECG, respiration, activity/movement/position, and a plethysmogram over short or long periods with no need to stay in hospital (European Commission, 2002).

The "AMON" system was designed to monitor and evaluate human vital signs, such as heart rate, two-lead ECG, blood pressure, oxygen blood saturation, skin perspiration, and body temperature using a wrist-mounted wearable device. The device gathers the data and transmits it to a remote telemedicine centre for further analysis and emergency care, using a GSM/UMTS cellular infrastructure (Anliker et al., 2004; European Commission, 2001).

"HealthVault" aims to build a universal hub of a network to connect personal health devices and other services that can be used to help store, and manage personal medical information in a single central site on the Web (Microsoft Corp., 2008). It will provide a seamless connection interface scheme for various home health and wellness monitoring devices, such as sport watches, blood glucose monitors, and blood pressure monitors marketed by medical equipment manufacturers worldwide.

On the other hand, many explorative studies on fundamental technology for vital signs monitoring have been conducted in the academic world and in research institutes. Much innovative instrumentation suitable in daily life has emerged and is gradually being commercialized.

Since the first accurate recording of an ECG reported by Willem Einthoven in 1895, and its development as a clinical tool, variants, such as Holter ECG, event ECG, and ECG mapping are now well known and have found a variety of applications in clinical practice. Measurement of ECG is now available from various scenarios. Whenever a person sits on a chair (Lim et al., 2006) or on a toilet (Togawa et al., 1989), sleeps in a bed (Kawarada et al., 2000; Ishijima, 1993), sits in a bathtub (Mizukami et al., 1989; Tamura et al., 1997), or even takes a shower (Fujii et al., 2002), his/her heart beat can be monitored, with the person unaware.

The smart dress, "Wealthy outfit", weaves electronics and fabrics together to detect the wearer's vital signs, and transmits the data wirelessly to a computer. The built-in sensors

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gather information on the wearer's posture and movement, ECG, and body temperature. Despite having nine electrodes and conductive leads woven into it, the suit looks and feels completely normal (Rossi et al., 2008; Marculescu et al., 2003).

The wellness mobile phone, "SH706iw", manufactured by the Sharp Corp. (Japan), has been released by NTT DoCoMo Corp. (Japan) in September 2008. It will have all the standard features of a mobile phone but will also act as a pedometer, a body fat meter, a pulse rate, and a breath gas monitor. Moreover, a built-in game-like application will support daily health management for fun and amusement (Sharp Corp., 2008; DoCoMo Corp., 2008).

According to an investigation report from the World Health Organization (WHO, 2002), most current healthcare systems still have some common issues that need to be addressed. (a) The difference between acute and chronic care is not sufficiently emphasized. The overall concept in system development has not shifted enough towards chronic conditions, and has not evolved to meet this changing demand. (b) Despite the importance of patients' health behaviour and adherence to improvement for chronic conditions, patients are not provided with a simple way to involve themselves in self-management and to have essential information to handle their condition to the best extent possible. (c) Patients are often followed up sporadically, and are seldom provided with a long-term management plan for chronic conditions to ensure the best outcomes.

Indeed, they are large obstacles in front of us that need to be cleared. We consider these issues a long-term difficult challenge to governments, communities, and individuals alike. We deem two main aspects should be paid primary attention. The first aspect is that vital-sign monitoring for chronic conditions requiring different philosophy and strategy tends to be ignored. Long-term chronic care is mostly oriented to untrained users in the home environment. However, many devices are far from being "plug and play", and require tedious involvement in daily operation. The second aspect is the lack of interconnection between multifarious physiological data within existing medical systems, as medication is usually decided by interpretation based on fragmented data and standards based on acute and emergent symptoms, and is often provided without the benefit of complete long-term physiological data.

To meet current needs, and to tackle the two problems above, our studies focus on developing a series of wearable/invisible vital-sign measurement technologies to facilitate data collection in the daily environment in perpetuity, and on applying data mining algorithms to conduct comprehensive interpretation of multifarious long-term data fusion, and ultimately to build a scalable healthcare integrated platform, SHIP, for various applicable domains, wherever vital signs are conducive.

2. Methods and results

Our studies included developing a series of instrumental technologies and data mining mathematical algorithms to construct finally a versatile platform, SHIP, integrated with wired and wireless network technologies. The following paragraphs describe an overall vision of SHIP and introduce three related constitutional technologies that we have been developing since 2002 (Chen et al., 2004).

2.1 SHIP

SHIP was conceived to provide three functions: (a) detection (monitoring multifarious vital signs by wearable/invisible means, (b) analysis (comprehensive interpretation of long-term

physiological data using data mining mathematics), and (c) service (providing customizable services to various human activity fields by a combination of multiple key technologies). As shown in Fig. 2, SHIP was constructed in a three-layer model which was supported by five pillars and many bricks.

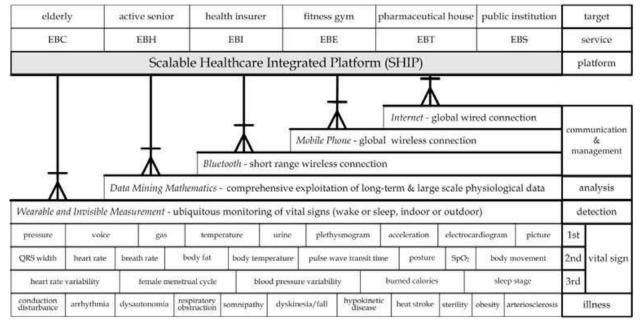


Fig. 2. Systemic architecture of the scalable healthcare integrated platform is founded on bricks, and supported by five pillars in a three layers structure. A variety of application domains can be created using the SHIP.

The first layer consists of a series of bricks for physiological data detection in three orders. Each brick in the first order can be considered as being a wearable/invisible measurement method, which can be used either indoors or outdoors, either awake or asleep. While each brick in the second and third orders indicates a data mining approach to derive information from the other bricks. The direct measurement signals are denoted as first-order vital signs. Second-order vital signs, such as heart rate, are derived from the first-order parameters. Third-order signs originate from the first- and second-order parameters.

Some of direct measurement objects are: pressure, voice, gas, temperature, ECG, acceleration, plethysmogram, and urine. The second-order vital signs are derived from the first-order vital signs, such as the QRS width and heart rate from ECG, the pulse rate and breathing rate from the pressure (Chen et al., 2005), posture and body movement from acceleration (Zhang et al., 2007), and pulse wave transit time from the ECG and plethysmogram. The third-order vital signs are derived from both the second-order and the first-order vital signs. For example, the variability in heart rate is derived from the heart rate profile. The female menstrual cycle is estimated from the body temperature (Chen et al., 2008a), and the variation in blood pressure is estimated from the pulse wave transit time (Chen et al., 2000). Changes in these parameters are indicators of specific ailments, such as arrhythmia from the variability in heart rate, somnipathy from body movements and sleep stage, and respiration obstruction from SpO2. We combined the directly measured and derived parameters to treat related illnesses such as cardiovascular disease, obesity, and respiratory obstruction.

Data mining mathematics for data analysis resides in the second layer. Most of statistical approaches and data warehouse technologies are applicable to conduct a comprehensive exploitation of the large volume of long-term accumulated physiological data. Innovative findings and understanding from this layer will be one of five pillars to support various application domains.

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The third layer consists of three pillars and is responsible for data communication and management through wireless and wired networking technology. Bluetooth telemetry technology is adopted as one pillar to support short-range wireless communication of data between sensor devices and home units or mobile phones. Mobile telephony and the Internet are two other pillars and used for wide-range data telecommunication and management.

SHIP has four characteristic features. (a) Its ubiquity makes it possible to detect and collect vital signs either asleep (unconscious status) or awake (conscious status), and either outdoors or indoors through wearable/invisible measurements and wired or wireless networks. (b) Its scalability allows users to customize their special package to meet individual needs, and also service providers to match different requests from medical/clinical use, industries, government agencies, and academic organizations through a variety of partnership options. (c) Its hot-line connectivity is realized by either mobile telephony or Internet (indoors or outdoors) and guarantees that any emergent event can be captured and responded to in real time. (d) Its interoperability is provided through a data warehouse that is configured in two formats. An exclusive format maintains security and enables high-speed data transmission within SHIP, and an externally accessible format ensures that SHIP is open to other allied systems through the HL7 standard (Health Level Seven Inc., 1997) to provide a seamless interface that is compatible with other existing medical information systems.

SHIP is intended to create a flexible platform for the exchange, management, and integration of long-term data collected from a wide spectrum of users, and to provide various evidence-based services to diverse domains. Subjects in target services are not only elderly and active seniors in healthcare but also subjects such as pharmaceutical houses for therapeutic effect tracing, insurance companies involved in risk assessment and claim transactions, transportation system drivers, fire fighters, and policemen involved in public security.

The layer and brick model in the SHIP architecture makes it possible to integrate many elementary achievements from ourselves and co-workers. Three different types of fundamental instrumentation (invisible/wearable/ubiquitous) and the results of data mining from our studies are introduced in the following sections.

2.2 Invisible sleep monitor

Invisible measurement means that a sensor unit can be deployed in an unoccupied area and is unobtrusive and concealable. Monitoring of vital signs can be performed in an invisible way, such that a user is unaware of its existence and does not have to take care that the device is present at all.

A schematic illustration of invisible sleep monitoring is shown in Fig. 3. There is a sensor plate and a bedside unit in the system configuration. A sensor unit is placed beneath a pillow, which is stuffed with numerous fragments of soft comfortable materials formed from synthetic resins. Two incompressible polyvinyl tubes, 30 cm in length and 4 mm in diameter, are filled with air-free water preloaded to an internal pressure of 3 kPa and set in

parallel at a distance of 11 cm from each other. A micro tactile switch (B3SN, Omron Co. Ltd) is fixed along the central line between the two parallel tubes. The two tubes above and the micro switch are sandwiched between two acrylic boards, both 3 mm thick. One end of each tube is hermetically sealed and the other end is connected to a liquid pressure sensor head (AP-12S, Keyence Co. Ltd). The inner pressure in each tube includes static and dynamic components, and changes in accordance with respiratory motion and cardiac beating. The static pressure component responds to the weight of the user's head, and acts as a load to turn on a micro tactile switch. The dynamic component reflects the weight fluctuation of the user's head due to breathing movements and pulsatile blood flow from the external carotid arteries around the head. Pressure signals beneath the near-neck and far-neck occiput regions are amplified and band-pass filtered (0.16-5 Hz), and the static component is removed from the signal. Only the dynamic component is digitized at a sampling rate of 100 Hz and transmitted to a remote database server through an Internet connection. The tactile switch is pressed to turn on a DC power supply via a delay switch (4387A-2BE, Artisan Controls Corp.) when the user lies down to sleep and places his/her head on the pillow.

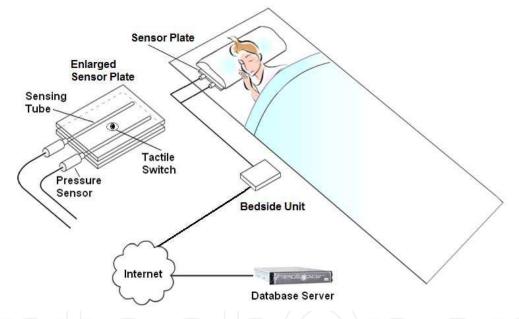


Fig. 3. Schematic illustration of the invisible monitoring of vital signs during sleep. A sensor plate is placed beneath a pillow. Signals reflecting pressure changes under the pillow are detected, digitized, and transmitted to a database server via the Internet by a bedside unit.

A 60 s fragment of raw signal measured under the near-neck occiput region during sleep is shown in Fig. 4(a). The breathing rate (BR), heart rate (HR), and body movements can be detected from the raw data measurements.

The BR and HR are detected by wavelet transformation on a dyadic grid plane using a multiresolution procedure, which is implemented by a recursive à trous algorithm. The Cohen–Daubechies–Fauraue (CDF) (9, 7) biorthogonal wavelet is the basis function used to design the decomposition and reconstruction filters (Daubechies, 1992). The raw measured signal is decomposed into an approximation and multiple detailed components through a cascade of filter banks (Mallat & Zhong, 1992; Shensa, 1992). Further mathematical theories can be found in Daubechies, 1992 and Akay, 1998. Implementation details are given in Chen et al., 2005, and Zhu et al., 2006.

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The wavelet transformation (WT) of a signal, x(t), is defined as follows:

$$W_s x(t) = \frac{1}{s} \int_{-\infty}^{+\infty} x(\tau) \psi(\frac{t-\tau}{s}) d\tau$$
 (2-2-1)

where *s* is the scale factor and $\psi(t)$ is the wavelet basis function. This is called a dyadic WT if $s = 2^{j}$ ($j \in Z$ and Z is the integral set). Two filter banks, the low-pass and high-pass decomposition filters H_0 and H_1 , and associated reconstruction filters, G_0 and G_1 , can be derived from the wavelet basis function and its scaling function, respectively. Using Mallat's algorithm, the dyadic WT of the digital signal, x(n), can be calculated as follows:

$$A_{2^{j}}x(n) = \sum_{k \in \mathbb{Z}} h_{0,2n-k} A_{2^{j-1}}x(k) , \qquad (2-2-2)$$

$$D_{2^{j}}x(n) = \sum_{k \in \mathbb{Z}} h_{1,2n-k} A_{2^{j-1}}x(k) , \qquad (2-2-3)$$

where $A_{2^{j}}x(n)$ and $D_{2^{j}}x(n)$ are the approximation and detail components, respectively, in the 2^{j} scale, and x(n) (or $A_{2^{0}}x(n)$) is the raw data signal. The terms h_{0} and h_{1} are the filter coefficients of \mathbf{H}_{0} and \mathbf{H}_{1} , respectively. Therefore, $A_{2^{j}}x(n)$ and $D_{2^{j}}x(n)$ ($j \in \mathbb{Z}$) can be extracted from x(n) (or $A_{2^{0}}x(n)$) using equations (2-2-2) and (2-2-3) recursively. The 2^{j-1} scale approximation signal can also be reconstructed from the 2^{j} scale approximation and the detail component:

$$\hat{A}_{2^{j-1}}x(n) = \sum_{k \in \mathbb{Z}} g_{0,n-2k} A_{2^j} x(k) + \sum_{k \in \mathbb{Z}} g_{1,n-2k} D_{2^j} x(k)$$
 (2-2-4)

where g_0 and g_1 are the filter coefficients of \mathbf{G}_0 and \mathbf{G}_1 , respectively. The terms $\hat{x}(n)$ (or $\hat{A}_{2^0}x(n)$) can be finally reconstructed by repeatedly using equation (2-2-4). Any noise in $D_{2^j}x(n)$ can be removed using a soft or hard threshold method before $\hat{A}_{2^{j-1}}x(n)$ is reconstructed. It should be pointed out that the sampling rate of the 2^j scale approximation and detail is $f_{2^j}(2^j)$, where f_s is the sampling rate of the raw signal.

Because the 2⁶ scale approximation waveform is close to a human breathing rhythm, while the detail waveforms of both the 2⁴ and 2⁵ scales contain peaks similar to those of human heartbeats, the 2⁶ scale approximation component, A6, is used to reconstruct the waveform for obtaining the BR, and the D4 and D5 detail components at the 2⁴ and 2⁵ scales are combined into a single synthesized waveform and then reconstructed to detect the HR. Figure 4(b) shows the reconstructed waveforms for HR detection, and Fig. 4(c) shows the reconstructed waveforms.

During a night's sleep, over a period of 4–8 h, a regular pulsation due to either the heart beating or breathing is not always detectable. Body movements may greatly distort the pressure variation signal pattern. In such a time slot, either the BR or the HR, and sometimes even both, are barely detectable. Instead, body movements are detected using a statistical method in such time slots. If a very large change, whose absolute value is four times larger than the standard deviation of the preceding detected movement-free raw

signal, is detected in the incoming signal, the preceding and succeeding 2.5 s periods from the movement detection point are treated as being body movement periods and not used to estimate the BR and HR. Detection of the BR is more sensitive to body movements than detection of the HR is.

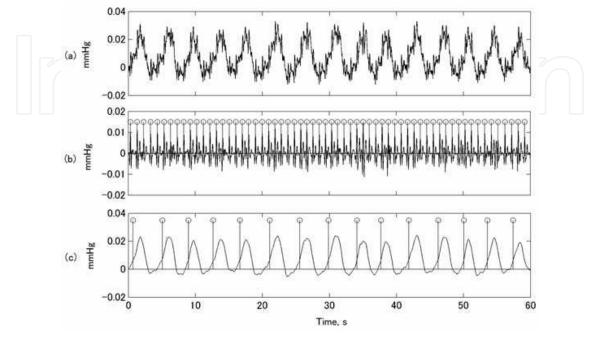


Fig. 4. A body movement-free sample and the detected BR and HR beat-by-beat. (a) The raw pressure signal data measured under the near-neck occiput region. (b) The pulse-related waveform reconstructed from the D4 and D5 components. (c) The breath-related waveform reconstructed from the A6 component. The open circles indicate the detected characteristic points for BR/HR determination.

Figure 5(a) shows a 60 s segment of a raw signal, which includes body movement during unstable sleep. When the pressure signal is distorted by a body movement, the periods detected that are from two reconstructed waveforms (HR-related and BR-related) are not always identical in both time and length. Because HR detection is usually more robust than BR detection, body movement detection from reconstructed BR-related waveforms is longer than that from HR-related waveforms. The final body movement period in terms of HR-related waveform detection is counted as 15.4 s, while that in terms of BR is 35.9 s. The final body movement outputs as 37.8 s from the OR operation of both results in the time domain.

Figure 6 shows a profile of the BR and the HR obtained from measurements over a single night. The vertical axis denotes the BR/HR in units of breaths per minute or beats per minute (bpm). The black dots and vertical bars, terminated at the upper and lower ends by short horizontal lines, show the mean values and standard deviation on a beat-by-beat basis for the HR and a breath-by-breath basis for the BR for each minute. Discontinuities in the estimation of the BR/HR are denoted by the vertical bars occurring sporadically over time, and their widths denote periods of body movement. The broader vertical bars correspond to longer body movement periods.

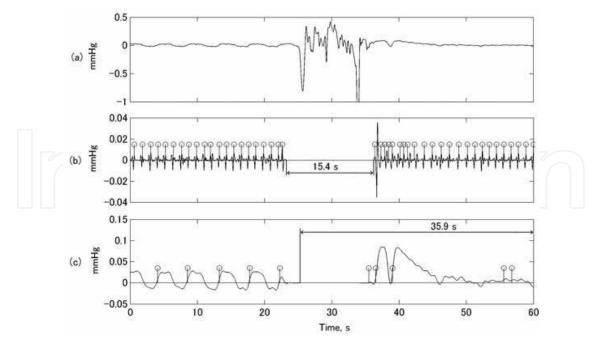


Fig. 5. An example of body movement in which both the BR and the HR are not fully detectable in a given period with different spans. The horizontal arrows indicate the body movement period in seconds. (a) Measured pressure signal distorted by body movements. (b) Reconstructed waveform and detected HR, as well as the detected body movement period. (c) Reconstructed waveform and detected BR as well as the detected body movement period.

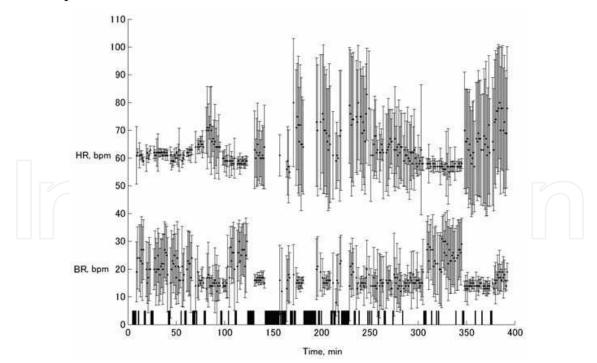


Fig. 6. Two profiles of the BR/HR obtained from measurements over a single night. The solid dots and vertical bars, terminated at the upper and lower ends by short horizontal lines, show the mean values and standard deviation within a period of one minute. The body movement periods are indicated by the variable-width vertical bars.

Figure 7 shows the complete profiles of the BR and HR during sleep over a period of 180 nights. The data were collected from a healthy female volunteer in her thirties at her own house over a period of seven months, under an informed agreement for the use of the data for research purposes. During this period, data over about 30 d were not measured. Therefore, data from a period of 180 d were recorded. The data are plotted on a day-by-day basis. The vertical axis represents the BR/HR in units of bpm. The symbols and vertical bars, terminated at the upper and lower ends by short horizontal lines, show the mean values and standard deviation of the detected HR (o) and the BR (*) in the corresponding night. The bold line is derived by filtering the mean values of the HR using a five-point Hanning window. The dashed line is an empirical estimate to indicate the possible trend of the average day base heart rate during those nights when data were not measured. Surprisingly, it is observed that the profile of the mean heart rate probably reveals a periodic property that corresponds to the female monthly menstrual cycle.

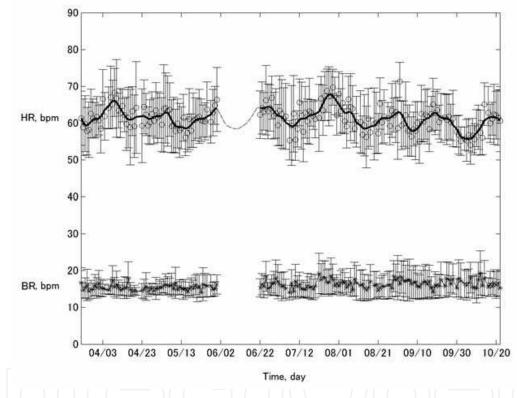


Fig. 7. Two complete profiles of the BR/HR over 180 nights. Data are plotted on a day-byday basis. The symbols and vertical bars, terminated at the upper and lower ends by short horizontal lines, show the mean values and standard deviation of the detected HR (o) and the BR (*) in the corresponding night.

The devised system is completely invisible to the user during measurements. "Plug is all" is one of its significant characteristics in securing perpetuity in data collection. All a user has to do is just to plug in an AC power cable and a LAN cable. A user can even forget the existence of the device and perform no other operation once it is installed beneath a pillow on a bed. This property will substantially enhance its feasibility and usability in the home environment. A subtle variation in pressure under the pillow is detected as a first-order signal. The BR/HR and body movements are derived as second-order parameters. Sleep stage estimation, assessment of sleep quality, and biphasic menstrual cycle properties are thirdorder parameters. In the end, a comprehensive interpretation of the multiple parameters obtained and a fully automatic operation property would increase its applicability in sleep lab studies, and also for screening patients who do not need a full sleep diagnosis at an early stage.

2.3 Wearable monitor for body temperature

Body temperature is one of the most important barometers indicating human health status. Moreover, the basal body temperature (BBT) is usually used for women to help estimate ovulation and to manage menstruation. However, a reliable evaluation of the menstrual cycle based on the BBT requires measurement of a woman's temperature under constant conditions for long periods. It is indeed a tedious task for a woman to measure her oral or armpit temperature under similar conditions when she wakes up every morning over a long period, because it usually takes an average of 5 min to measure temperature orally, or 10 min under the armpit. Moreover, the traditional method for evaluating ovulation or menstrual cycle dynamics in clinical practice is often based on a physician's empirical observations on serial measurements of BBT. It has been pointed out that the BBT failed to demonstrate ovulation in approximately 20% of ovulation cycles among 30 normally menstruating women (Moghissi, 1980). To improve user accessibility and the accuracy of the application of the BBT, we have developed a tiny wearable device for cutaneous temperature measurements and a Hidden Markov Model (HMM) based a statistical approach to estimate the biphasic properties of body temperature during the menstrual cycle using a series of cutaneous temperature data measured during sleep.

The wearable monitor can be attached to a woman's underwear or brassiere when asleep to measure the cutaneous temperature around the abdominal area or between the breasts, as shown in Fig. 8.

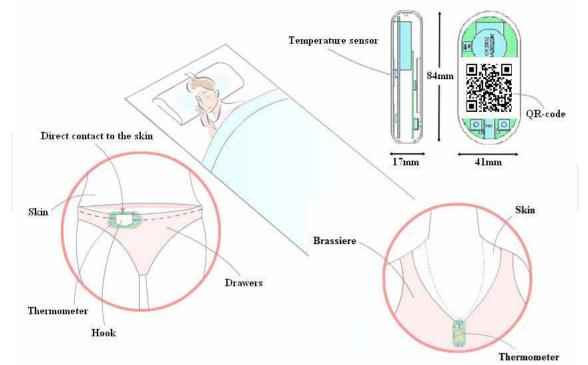


Fig. 8. A small, light wearable device (size = $41 \times 84 \times 17$ mm³, weight = 59 g) for cutaneous temperature measurements during sleep (QOL Co. Ltd., 2008).

The device is programmed to measure temperature over 10 min intervals from midnight to 6 am. At most, 37 data points can be collected during the six hours. Outliers above 40 °C or below 32 °C are ignored. The collected temperature data are encoded in a two-dimensional bar code, known as "Quick Response" code (QR code) (Denso Wave Inc., 2000) and depicted on an LCD display. As shown in Fig. 9, the user uses the camera built into a mobile phone to capture the QR code image (a) on the device display (shown in the circle on the left-hand side of Fig. 9). Once the QR code is captured into the mobile phone (b), the original temperature data (c) are recovered from the captured image and transmitted to a database server via the mobile network for data storage and physiological interpretation through data mining.



Fig. 9. A procedure for temperature data collection using a wearable sensor and a mobile phone. (a) A QR code image. (b) A QR code captured by a camera built into a mobile phone. (c) Original data recovered from the image captured by a mobile phone (QOL Co. Ltd., 2008).

The temperature data measured during sleep over a six-month period are shown in Fig. 10(a). The nightly data are plotted in the vertical direction and have a range of 32 to 40 °C. The purpose of data mining in this study was to estimate the biphasic properties in the temperature profile during the menstrual cycle from cutaneous temperature measurements. As shown in Fig. 11, the biphasic properties of the menstrual cycle can be modelled as a discrete Hidden Markov Model (HMM) with two hidden phases. The measured temperature data are considered to be observations being generated by the Markov process from an unknown phase: either a low-temperature (LT) phase or a high-temperature (HT) phase, according to the probability distribution. The probability $b_{\rm H}(k)$ is indicative that the value *k* is generated from the hidden HT phase. The probability $a_{\rm ii}$ is indicative of a hidden phase transition between LT and HT phases.

Figure 10(b) shows the results after pre-processing to removing outliers from the raw data and eliminating any discontinuities from non-data-collection days. Figure 10(c) shows the HMM estimation output using the pre-processed data from Fig. 10(b) as the input. Figure 10(d) shows the estimation of the biphasic properties after post-processing. The superimposed black symbols "*" denote the menstrual periods recorded by the user. A transition from the HT phase to the LT phase denotes a menstrual period, while the reverse transition denotes ovulation.

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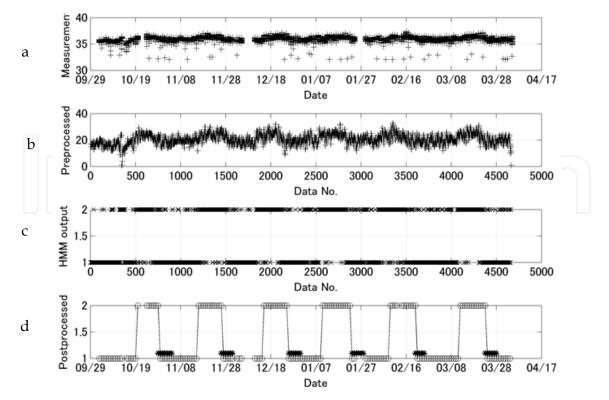


Fig. 10. Estimation procedure of a biphasic temperature profile. (a) Raw temperature data measured over a period of six months. (b) Pre-processed results. (c) Biphasic estimation based on an HMM approach. (d) Post-processed results. The symbol "*" denotes a menstrual period recorded by the user.

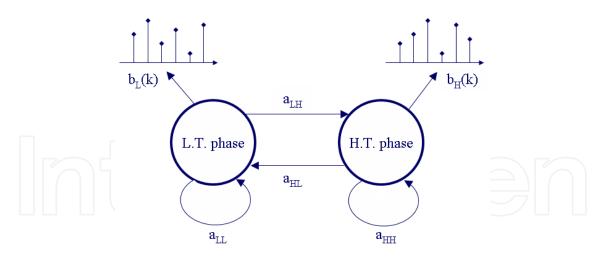


Fig. 11. A discrete hidden Markov model with two hidden phases for estimating biphasic property in a menstrual cycle from cutaneous temperature measurements.

The biphasic properties shown in Fig. 10(c) were estimated by finding an optimal HMM parameter set that determines the hidden phase from which each datum arises. This is based on a given series of measured temperature data, as shown in Fig. 10(b). The parameter set $\lambda(A,B,\pi)$ is assigned randomly in the initial condition and optimized through the forward-backward iterative procedure until P(O | λ) converges to a stable maximum value or until the absolute logarithm of the previous and current difference in P(O | λ) is not greater than δ .

The algorithm for calculating the forward variable, α , the backward variable, β , and the forward–backward variable, γ , are shown in equations (2-3-1) to (2-3-3).

The forward variable, $\alpha_t(i)$, denotes the probability of phase, q_i , at time, t, based on a partial observation sequence, $O_1, O_2, ..., O_t$, until time t, and can be calculated using the following steps for a given set of $\lambda(A, B, \pi)$.

$$\alpha_{i}(i) = P_{r}(O_{1}, O_{2}, ..., O_{t}, i_{t} = q_{i} \mid \lambda)$$

$$\alpha_{1}(i) = \pi_{i}b_{i}(O_{1}), \quad 1 \le i \le N, t = 1$$

$$\alpha_{t+1}(j) = \left[\sum_{i=1}^{N} \alpha_{i}(i)a_{ij}\right]b_{j}(O_{t+1}), \quad 1 \le j \le N, t = 1, 2, ..., T - 1 \quad (2-3-1)$$

The backward variable, $\beta_t(i)$, denotes the probability of phase, q_i , at time, t, based on a partial observation sequence, $O_{t+1}, O_{t+2}, \dots, O_T$, from time t+1 to T, and can be calculated using the following steps for a given set of $\lambda(A, B, \pi)$.

$$\beta_{t}(i) = P_{r}(O_{t+1}, O_{t+2}, ..., O_{T} | i_{t} = q_{i}, \lambda)$$

$$\beta_{T}(i) = 1, \quad 1 \le i \le N, t = T$$

$$\beta_{t}(i) = \sum_{j=1}^{N} a_{ij} b_{j}(O_{t+1}) \beta_{t+1}(j), \quad 1 \le i \le N, t = T - 1, T - 2, ..., 1 \quad (2-3-2)$$

To find the optimal sequence of hidden phases for a given observation sequence, O, and a given model, $\lambda(A,B,\pi)$, there are multiple possible optimality criteria.

Choosing the phases, q_t , that are individually most likely at each time, t, i.e., maximizing $P(q_t = i | O, \lambda)$, is equivalent to finding the single best phase sequence (path), i.e., maximizing $P(Q | O, \lambda)$ or $P(Q, O | \lambda)$. The forward-backward algorithm is applied to find the optimal sequence of phases, q_t , at each time, t, i.e., to maximize $\gamma_t(i) = P(q_t = i | O, \lambda)$ for a given observation sequence, O, and a given set of $\lambda(A, B, \pi)$.

$$\begin{split} \gamma_{t}(i) &= P(q_{t} = i \mid 0, \lambda) \\ &= \frac{P(O, q_{t} = i \mid \lambda)}{P(O \mid \lambda)} = \frac{P(O, q_{t} = i \mid \lambda)}{\sum_{i=1}^{N} P(O, q_{i} = i \mid \lambda)} \\ &= \frac{P(o_{1}o_{2}...o_{t}, q_{t} = i \mid \lambda) P(o_{t+1}o_{t+2}...o_{t} \mid q_{t} = i, \lambda)}{\sum_{i=1}^{N} P(o_{1}o_{2}...o_{t}, q_{t} = i \mid \lambda) P(o_{t+1}o_{t+2}...o_{t} \mid q_{t} = i, \lambda)} \\ &= \frac{\alpha_{t}(i)\beta_{t}(i)}{\sum_{i=1}^{N} \alpha_{t}(i)\beta_{t}(i)} \end{split}$$
(2-3-3)

The most likely phase, q_t^* at time *t* can be found as:

$$q_{t}^{*} = \arg\max_{1 \le i \le N} \left[\gamma_{t}\left(i\right) \right], \quad 1 \le t \le T .$$

$$(2-3-4)$$

As there are no existing analytical methods for optimizing $\lambda(A,B,\pi)$, $P(O|\lambda)$ or $P(O,I|\lambda)$ is usually maximized (i.e., $\lambda^* = \arg \max_{\lambda} [P(O|\lambda)]$ or $\lambda^* = \arg \max_{\lambda} [P(O,Q|\lambda)]$) using gradient

techniques and an expectation-maximization method. In this study, the Baum–Welch method was used because of its numerical stability and linear convergence (Rabiner, 1989). To update $\lambda(A,B,\pi)$ using the Baum–Welch re-estimation algorithm, we defined a variable, $\xi_t(i,j)$, to express the probability of a datum being in phase *i* at time *t* and phase *j* at time *t*+1, given the model and the observation sequence:

$$\xi_{t}(i,j) = P(q_{t} = i, q_{t+1} = j \mid O, \lambda) = \frac{P(q_{t} = i, q_{t+1} = j, O \mid \lambda)}{P(O \mid \lambda)}.$$
(2-3-5)

From the definitions of the forward and backward variables, $\xi_t(i,j)$ and $\gamma_t(i)$, can be related as:

$$\xi_{t}(i,j) = \frac{\alpha_{t}(i)a_{ij}b_{j}(o_{t+1})\beta_{t+1}(j)}{P(O|\lambda)} = \frac{\alpha_{t}(i)a_{ij}b_{j}(o_{t+1})\beta_{t+1}(j)}{\sum_{i=1}^{N}\sum_{j=1}^{N}\alpha_{t}(i)a_{ij}b_{j}(o_{t+1})\beta_{t+1}(j)},$$
(2-3-6)

$$\gamma_{t}(i) = P(q_{t} = i \mid O, \lambda) = \sum_{j=1}^{N} P(q_{t} = i, q_{t+1} = j \mid O, \lambda) = \sum_{j=1}^{N} \xi_{t}(i, j), \quad (2-3-7)$$

where $\sum_{i=1}^{T-1} \gamma_i(i)$ denotes the expected number of transitions from phase *i* in *O*. The term

 $\sum_{t=1}^{T-1} \xi_t(i, j)$ denotes the expected number of transitions from phase *i* to phase *j* in *O*.

Therefore, $\lambda(A,B,\pi)$ can be updated using equations (2-3-8) to (2-3-10) as follows.

As π_i is the initial probability and denotes the expected frequency (number of times) in phase *i* at time t = 1 as $\pi_i = \gamma_1(i)$, it can be calculated using the forward and backward variables.

$$\pi_{i} = \frac{\alpha_{i}(i)\beta_{i}(i)}{\sum_{i=1}^{N}\alpha_{i}(i)\beta_{i}(i)} = \frac{\alpha_{i}(i)\beta_{i}(i)}{\sum_{i=1}^{N}\alpha_{T}(i)}, \qquad (2-3-8)$$

The transition probability from phase *i* to phase *j*, a_{ij} , can be calculated from the expected number of transitions from phase *i* to phase *j* divided by the expected number of transitions from phase *i*.

$$a_{ij} = \frac{\sum_{t=1}^{T-1} \xi_t(i,j)}{\sum_{t=1}^{T-1} \gamma_t(i)} = \frac{\sum_{t=1}^{T-1} \alpha_t(i) a_{ij} b_j(o_{t+1}) \beta_{t+1}(j)}{\sum_{t=1}^{T-1} \alpha_t(i) \beta_t(i)},$$
(2-3-9)

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The term $b_j(k)$ is the expected number of times in phase *j* and the observed symbol o_k divided by the expected number of times in phase *j*, and it can be calculated using:

$$b_{j}(k) = \frac{\sum_{\substack{i=1\\s_{i},o_{i}=o_{k}}}^{T} \gamma_{i}(j)}{\sum_{i}^{T} \gamma_{i}(j)}$$
(2-3-10)

The initial input quantities are the known data N, M, T, and O and the randomly initialized $\lambda(A,B,\pi)$. Once α , β , and γ are calculated using equations (2-3-1) to (2-3-3), then $\lambda(A,B,\pi)$ is updated using equations (2-3-8) to (2-3-10), based on the newly obtained values of α , β , and γ . The search for the optimal parameter set, λ_{opt} , is terminated when P(O | λ) converges to a stable maximum value or when the absolute logarithm of the previous and current difference in P(O | λ) is equal to or smaller than δ . Thus, the most likely phase from which a datum is observed can be estimated using equation (2-3-4). A sample result estimated using the HMM algorithm is shown in Fig. 10(c).

The algorithmic performance is evaluated by comparing the user's own record of their menstrual periods with the algorithmically estimated result. When a transition line coincides with a self-declared menstrual period, then it is counted as a "true positive". If a transition line does not coincide with a self-declared menstrual period, then it is counted as either a "false negative" or a "false positive".

Figure 12 shows a poor sample with two different estimation errors. One is a false negative estimation occurring around November 14, where there was a menstrual period but it was not detected. Another is a false positive error, occurring around July 17, where an HT to LT transition was detected but there was no actual menstrual period.

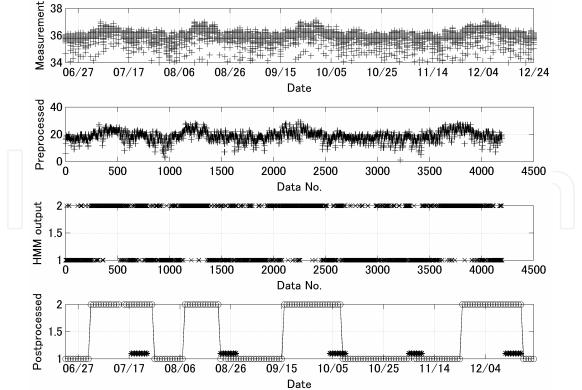


Fig. 12. A poor sample of biphasic profile estimation, where two menstrual periods occurring around July 17 and November 14 were not detected correctly by the algorithm.

To evaluate the algorithmic performance in finding the biphasic properties of the temperature data during menstrual cycles, both sensitivity and positive predictability are introduced. The sensitivity (Sens) denotes an algorithm's ability to estimate correctly the biphasic properties that coincide with the user's records of menstrual periods. This value is calculated using equation (2-3-11). When Sens has a value close to 1.0, then it means that there is less underestimation.

$$Sens = \frac{TP}{TP + FN},$$
(2-3-11)

The positive predictability (PP) denotes the confidence of the positive estimation. This is calculated using equation (2-3-12), and when PP has a value that is close to 1.0, then it means that there is less overestimation:

$$PP = \frac{TP}{TP + FP},$$
(2-3-12)

where *TP* denotes a true positive where the algorithmic estimation and the user's records coincide. *FN* is a false negative, and it counts the number of undetected menstrual periods. *FP* is a false positive, and it counts the number of detected HT to LT transitions where no actual menstrual period occurred.

In this study, a tiny wearable thermometer and an HMM-based data mining approach were developed and validated using data collected over a period of six months from 30 female volunteers. For a total of 190 self-declared menstrual cycles, the number of estimated cycles was TP = 169, the number of undetected cycles was FN = 15, and the number of falsely detected cycles was FP = 6. Therefore, by applying equations (2-3-11) and (2-3-12), the sensitivity was 91.8% and the positive predictability was 96.6%.

The device automatically collected the cutaneous temperature around the abdominal area or between the breasts without much discomfort during sleep. This met the requirement for perpetuity in body temperature measurements due to a low disturbance to daily life. The algorithm estimated the biphasic cyclic properties of the temperature profiles during menstrual cycles based on a series of long-term temperature data, with no need for any experimental or subjective involvement. This provides a promising approach for managing female premenstrual syndromes and birth control.

2.4 Ubiquitous monitoring based on a mobile phone

The voice is the sound made by the vibration of the vocal cords, caused by air passing through the larynx and bringing the cords closer together. Voice production is a complex process that starts with muscle movement and involves: phonation (voice), respiration (breathing process), and articulation (throat, palate, tongue, lips, and teeth). These muscle movements are initiated, coordinated, and controlled by the brain, and monitored through hearing and touch. A variety of changes in voice, such as pitch, intensity, and fundamental frequency, can take place when any of the above factors change, and will be one of the first and most important symptoms of mental depression, or some physical diseases, such as cancer, Parkinson's disease, epilepsy, stroke, and Alzheimer's disease, which means that

early detection significantly increases the effectiveness of any treatment (France & Kramer, 2001; O'Shaughnessy, 1999; Mathieson, 2001).

Recently, many studies have been conducted to identify pathological voices by applying various classifiers to acoustic features, such as cepstral coefficients and pitch dynamics using Gaussian mixtures in a Hidden Markov Model (HMM) classifier. These have obtained a 99.44% correct classification rate for discrimination of normal and pathological speech using the sustained phoneme "a" from over 700 subjects (Dibazar et al., 2002). In contrast to sustained vowel research, the effectiveness of acoustic features extracted from continuous speech has also been evaluated. A joint time-frequency approach for classifying pathological voices using continuous speech signals was proposed to obviate the need for segmentation of voiced, unvoiced, and silence periods. The speech signals were decomposed using an adaptive time-frequency transform algorithm, and several features, such as the octave max, octave mean, energy ratio, length ratio, and frequency ratio were extracted from the decomposition parameters and analysed using statistical pattern classification techniques. Experiments with a database consisting of continuous speech samples from 51 normal and 161 pathological talkers yielded a classification accuracy of 93.4% (Umapathy et al., 2005).

On the other hand, mobile phones are a pervasive tool, and not only are they able to be used in voice communication but also they are an effective voice collector in daily conversation. This study aims to develop a ubiquitous tool for identifying various pathological voices using mobile phones and support vector machine (SVM) mathematics through the classification of voice features.

A. Voice data and acoustic features

The voice data used in this study were excerpted from the Disordered Voice Database Model 4337 v2.7.0 (Key Elemetrics Corp., 2004). Because parts of the recordings and clinical information were incomplete, we only chose 214 subjects aged from 13 to 82 years (mean \pm STD = 45 \pm 16 years) whose records were fully contained. The selected dataset included 33 healthy subjects (14 males and 19 females) and 181 patients (86 males and 95 females) who suffered from various voice disorders, such as polypoid degeneration, adductor spasmodic dysphonia, vocal fold anomalies, hyperfunction, and erythema, totalling more than 40 types of abnormality. The 25 acoustic features are listed in Table 1. They were calculated using the voice data, which were sampled from the vowel "a" pronunciation by each subject. The calculation method for each feature can be found in detail in the CDROM (Key Elemetrics Corp., 2004).

B. SVM principle

As a related supervised learning method used for classification and regression, SVM is a generalized linear classifier, but it differs from other methods because of its largest margin and its simplest form among several others. The optimal separation hyperplane was determined to maximize the generalization ability of the SVM. However, because most real-world problems are not linearly separable, the SVM introduced kernel tricks to deal with the linearly inseparable problems. Therefore, in theory, a linearly inseparable problem in the original data space can be completely transformed into a linearly separable problem in high-dimensional feature space.

	т. (NT	T (
No	Feature	Meaning	No	Feature	Meaning
1	APQ	Amplitude perturbation quotient	14	PPQ	Pitch period perturbation quotient
2	ATRI	Amplitude tremor intensity index	15	RAP	Relative average perturbation
3	DSH	Degree of subharmonic components	16	sAPQ	Smoothed amplitude perturbation quotient
4	Fatr	Amplitude-tremor frequency	17	Shdb	Shimmer in dB
5	Fftr	Fo-tremor frequency	18	Shim	Shimmer per cent
6	Fhi	Highest fundamental frequency	19	SPI	Soft phonation index
7	Flo	Lowest fundamental frequency	20	sPPQ	Smoothed pitch period perturbation quotient
8	Fo	Average fundamental frequency	21	STD	STD of fundamental frequency
9	FTRI	Frequency tremor intensity index	22	То	Average pitch period
10	Jita	Absolute jitter	23	vAm	Coeff. of amplitude variation
11	Jitt	Jitter per cent	24	vFo	Coeff. of fundamental frequency variation
12	NHR	Noise-to-harmonic ratio	25	VTI	Voice turbulence index
13	PFR	Phonatory fundamental freq. range			

A Scalable Healthcare Integrated Platform (SHIP) and Key Technologies for Daily Application

Table 1. Abbreviations of acoustic features and their physical meanings.

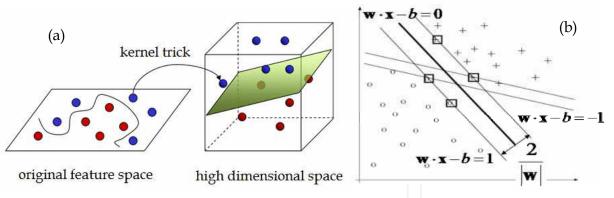


Fig. 13. Principle of the SVM. A separating hyperplane in high-dimensional feature space and a maximum margin boundary between two data sets. (a) A non-linear classification problem in an original lower-dimensional feature space is transformed into a linear classification problem in a higher-dimensional feature space through a non-linear vector function, or a kernel trick. (b) An optimal hyperplane to separate two classes of data in highdimensional space with the largest geometric margin.

As shown in Fig. 13, suppose we have an original dataset, $\{x_i, y_i\}$, $x_i \in \mathbb{R}^d$, $y_i \in \{-1, 1\}$, i = 1, ..., M. A separating hyperplane separates the positive from the negative subdataset. The points, x_i , which lie on the hyperplane satisfy $\mathbf{w} \cdot \mathbf{x}_i + b = 0$, where \mathbf{w} is normal to the hyperplane,

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 $|b|/||\mathbf{w}||$ is the perpendicular distance from the origin to the hyperplane, and $||\mathbf{w}||$ is the Euclidean norm of w. Let d_+ and d_- be the shortest distances from the separating hyperplane to the closest positive data point and the closest negative data point, respectively. The margin of a separating hyperplane is defined as d_++d_- . In the linearly separable case, all the data points satisfy the following constraints.

$$\mathbf{x}_{i} \cdot \mathbf{w} + b \ge +1 \text{ for } y_{i} = +1$$
(2-4-1)
$$\mathbf{x}_{i} \cdot \mathbf{w} + b \le -1 \text{ for } y_{i} = -1$$
(2-4-2)
ese can be combined into one set of inequalities.

These can be combined into one set of inequalities.

$$y_i(\mathbf{x}_i \cdot \mathbf{w} + b) - 1 \ge 0 \quad \forall I \tag{2-4-3}$$

Clearly, the data points, which lie in the hyperplane H_{1} , $\mathbf{w} \cdot \mathbf{x}_i + b = 1$, lie at a perpendicular distance $|1-b|/||\mathbf{w}||$ from the origin. Similarly, the data points that lie in the hyperplane H_2 , $\mathbf{w} \cdot \mathbf{x}_i + b = -1$, lie at a perpendicular distance $|-1-b|/||\mathbf{w}||$ from the origin. Hence, the margin is simply 2/||w||. Therefore, we can find an optimal separating hyperplane that has the largest margin by minimizing $\|\mathbf{w}\|^2$, subject to the constraints in equation (2-4-3).

The data points that lie on hyperplanes H_1 or H_2 are called support vectors. Their removal would change the solution found. This problem can now be solved through a Lagrangian formulation by introducing non-negative Lagrangian multipliers, α_i , i = 1,...,M, for each of the inequality constraints (2-4-3). The Lagrangian takes the form:

$$L_{p} \equiv \frac{1}{2} \|\mathbf{w}\|^{2} - \sum_{i=1}^{M} \alpha_{i} y_{i} (\mathbf{x}_{i} \cdot \mathbf{w} + b) + \sum_{i=1}^{M} \alpha_{i}.$$
(2-4-4)

By taking the gradient of L_P with respect to **w**, *b* vanishes, giving the outcome:

$$w = \sum_{i=1}^{M} \alpha_{i} y_{i} x_{i}, \qquad (2-4-5)$$

$$\sum_{i=1}^{M} \alpha_{i} y_{i} = 0, \ \alpha_{i} \ge 0, \ i = 1, \dots, M. \qquad (2-4-6)$$

Substituting equations (2-4-5) and (2-4-6) into equation (2-4-4) gives:

$$L_{p} = \sum_{i=1}^{M} \alpha_{i} - \frac{1}{2} \sum_{i=1,j=1}^{M} \alpha_{i} \alpha_{j} y_{i} y_{j} x_{i} \cdot x_{j}.$$
(2-4-7)

Maximizing equation (2-4-7) under the constraints in equation (2-4-6) is a concave quadratic programming problem. If the dataset is linearly separable, then the global optimal solution, α_i , *i* = 1,...,*M*, can be found. Then, the boundary decision function is given by:

$$D(\mathbf{x}) = \sum_{i \in i_{S}} \alpha_{i} y_{i} \mathbf{x}_{i} \cdot \mathbf{x} + b , \qquad (2-4-8)$$

where, *i*^{*S*} is the set of support vector indexes, and *b* is given by:

$$b = y_i - \mathbf{w} \cdot \mathbf{x}_i, \, i \in i_S. \tag{2-4-9}$$

To tackle linearly inseparable problems, the SVM is extended by incorporating slack variables and kernel tricks. The non-negative slack variables determine the trade-off between maximization of the margin and minimization of the classification error. The kernel tricks use a non-linear vector function, $g(\mathbf{x})$, to map the original dataset into a higher-dimensional space. Therefore, the linear decision function in the new feature space can be given by:

$$D(\mathbf{x}) = \mathbf{w} \cdot \mathbf{g}(\mathbf{x}) + b$$

= $\sum_{i \in i_{s}} \alpha_{i} y_{i} \mathbf{g}(\mathbf{x}_{i}) \cdot \mathbf{g}(\mathbf{x}) + b = \sum_{i \in i_{s}} \alpha_{i} y_{i} H(\mathbf{x}_{i}, \mathbf{x}) + b'$ (2-4-10)

where $H(\mathbf{x}_i, \mathbf{x})$ is the Mercer kernel. The advantage of using the kernel trick is that the highdimensional feature space can be treated implicitly by calculating $H(\mathbf{x}_i, \mathbf{x})$ instead of $g(\mathbf{x}_i) \cdot g(\mathbf{x})$.

Twenty-five acoustic features were calculated from each subject's voice data, and 214 data segments from 214 individuals were labelled as "-1" for a healthy voice and "1" for a pathological voice. There are two major steps in voice classification. Firstly, a principal component analysis (PCA) was used to reduce the original feature dimensions as much as possible by ignoring correlated features while retaining the most significant information in the new dataset. Secondly, a transformed subdataset in the new coordinate system was applied to the SVM to elucidate the identification boundary for healthy and pathological voices. Two kernels, Gaussian and polynomial, with different parameter combinations, were evaluated, as shown in Figs 14 and 15. The black contours refer to the identification boundaries, the white contours to the margins, and the symbols on the white contours to the support vectors.

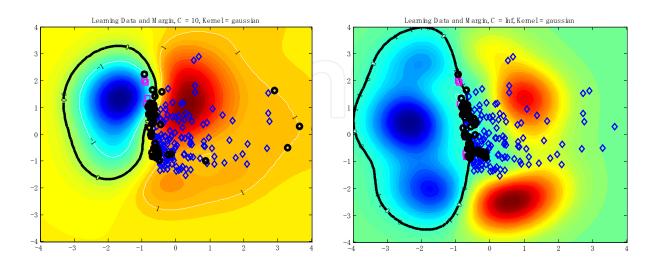


Fig. 14. Identification boundary, margin, and support vectors obtained using the Gaussian kernel with different slack variables ξ (left = 10 and right = + ∞).

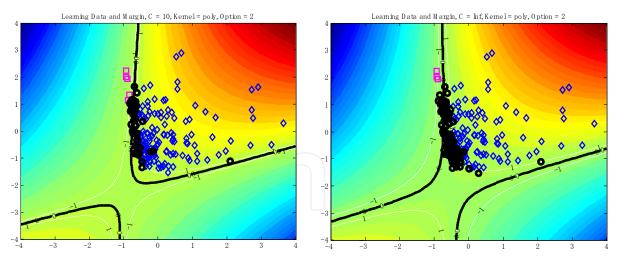


Fig. 15. Identification boundary, margin, and support vectors obtained using the polynomial kernel with d = 2 and different slack variables ξ (left = 10 and right = + ∞).

In this study, 25 acoustic features derived from a single vowel sound "a" were used to identify pathological and healthy voices. The accuracy under different combinations of features was tested using a fivefold cross-validation method. The results show that the STD, Fatr, and NHR were the most sensitive features in detecting pathological changes. Moreover, the length of the voice sample, the number of voice segments, and the total number of detected pitch periods affected the identification accuracy. The highest accuracy in detecting pathological voices from healthy voices was 97.0% (Peng et al., 2007a; Peng et al., 2007b).

Not only its mathematical elegance but also the practical advantage of simplicity means the SVM approach is promising for implementing in any portable device. A mobile phone's built-in microphone works perfectly as a "one-stone-kills-two-birds" solution for identifying subtle distinctions in voices ubiquitously without any inexpediency during daily conversation.

3. Discussion

Life is not just a biochemical process but rather is a symphony of many rhythms on the micro and macro levels from the milliseconds of single-neuron activity to monthly procreation, and yearly developmental aging. However, intrinsic biorhythms are gradually stupefying in modern lifestyles because of artificial lighting and controlled environments. A healthy life is considered a state or process of complete physical, mental, and social well-being, and not merely the absence of disease or infirmity (WHO, 1948). Even though research that may unravel the interplay between depression and other diseases has barely begun, there is a strong statistical link between the incidence of depression and several other diseases, including cancer, Parkinson's disease, epilepsy, stroke, and Alzheimer's disease. More and more doctors and patients recognize that mental state and physical well-being are intimately connected (Wisneski & Anderson 2005).

Because many currently used vital signs, such as blood pressure, ECG, and metabolized chemicals, are so important in life processes, the body must exert itself through the feedback regulatory mechanisms of the autonomic nervous system and the immune system to avoid disorders. In the case where these parameters are detected out of range, it is actually a sign

that the body has become intolerant, and this is indicated by its exhaustion in maintaining proper biological functions. By then, it is already too late for us to take action. We should look for more sensitive symptoms that can reflect any small changes in the early stages of the development of ailments. Extrinsic extremities, such as capillary vessels and peripheral tissues, are less important in body functioning, and these areas are the first candidates to be deserted to guarantee that vital organs and tissues do not malfunction. Traditional Chinese Medicine (TCM) diagnoses by observing the colour of skin and the tones of sound by skilful means of inquiry, smell, and touch to acquire insight into pathological changes in a patient (Qu, 2007).

Much effort is being conducted towards chronic illness prevention and early warning. It has been found that the correlation between the blood sugar level and the acetone in a patient's breath is significant in the diagnosis of diabetes (Zhang et al., 2000). More barometers, such as the colour and texture of the skin, nails, palms, and tongue, are worthy of further investigation.

Application of these new approaches in treating chronic conditions requires advanced sensory technology feasible for daily monitoring, and a flexible platform for users, physicians, and allied health workers to manage the long-term data, as well as data mining mathematics suitable for the comprehensive interpretation of multiple data fusion.

Innovative sensory instrumentation technologies are indispensable for monitoring a wide range of physiological data in the environment of everyday living, because most users are not trained professionally, and as perpetual monitoring is preferable in chronic conditions, the sensory device should have some key advantages, such as zero administration, easy manipulation, automatic fault recovery, and the absence of unpleasantness or disturbance to everyday living. Fortunately, changes in long-term physiological data are much more important than the absolute values in long-term application, measurement resolution is prior to the accuracy, and this partly relieves the requirements to sensory instrumentation. Studies in this direction in several of our projects have been conducted using three main models of sensory technologies and real data validation in their field tests.

Invisible instrumentation – e.g., the sleep monitor – requires no user intervention at all during monitoring. This "plug is all" property makes it possible for a user to plug in several cables, such as a power supply and LAN, and the system then works automatically. Users do not have to remember how to use the machine and notice what it is doing behind them.

A wearable device – e.g., the women's temperature monitor – is worn as if a part of the underwear or brassiere with little discomfort. The feedback from the participants in our feasibility study indicated that taking a magic-like QR-code picture is much more enjoyable than taking an oral or armpit BBT measurement after morning arousal from sleep. Many of the participants found "play and fun" in this routine activity.

Ubiquitous monitoring technology based on a mobile phone has been developed for carryon usage, at any time or place, without any need for attention. Indeed, a mobile phone is a marvellously powerful portable machine that has many built-in sensors, such as a microphone for sound, a camera for pictures, and a touch panel for pressure measurement, applicable for detecting vital signs. An additional built-in gas sensor would make it possible for a mobile phone to monitor breath gases.

As more and more innovative vital signs are detected in the environment of everyday living from a large number of the population and are accumulated over a longer period, a flexible data warehouse for facilitating the large volume of data is necessary. SHIP was devised as a scalable and customizable platform to make it possible to integrate a broad spectrum of

physiological data, and also to perform multiple data fusion through comprehensive data mining to provide versatile services in diversified application domains.

By mining the long-term physiological data accumulated from the large-scale population, more physiological insights related to physical and mental ailments will be discovered from changes in the biorhythms in the long-term profiles, which are important for chronic conditions but is unable to be derived from fragmentary data collected over the short term.

Biological rhythms are found in a wide range of frequencies from seconds to years, and in different biological levels from the entire body to individual cells. Much research has been conducted on different scales, such as circadian, diurnal, monthly, and seasonal rhythms. Many rhythmic events are already well known. Sudden death due to cardiovascular disease happens mostly in the early morning. A high risk of stroke and silent cerebral infarct are related to the morning surge of blood pressure that occurs on waking. One of the most prominent biorhythms is the heartbeat, which is currently used to obtain insight into many cardiovascular-related diseases (Malik et al., 1996). There is growing evidence indicating that a marked diurnal variation exists in the onset of cardiovascular events, with a peak incidence of myocardial infarction, sudden cardiac death, and ischemic and hemorrhagic stroke occurring in the morning (from 6 am to noon), after a nadir in these events during the night (Kario et al., 2003).

On the other hand, the "Medical Classic of Emperor Huang" (Wikipedia, 2008), one of the TCM representative masterpieces published more than 2,000 years ago, divided a day into 12 time slots. Each time slot is two hours long and is responsible for alternating metabolic oscillations in different visceral organs. Each visceral organ has its optimal on-duty and offduty time slot. During the period 5–7 am, the cardiovascular system is on duty and is most active. This leads to an increase in the burden on the heart and induces a morning surge in blood pressure. Even worse, patients suffering from cardiovascular disease are subject to sudden death. Such rhythmic concepts are close to current understanding (Qu, 2007).

In contrast, a dualism that considered mind and body to be separate prevailed until René Déscartes in the 17th century. Oriental medicine treats the body and mind as a whole and aims to enhance holistic balance and visceral organic harmony through enhancing self-healing functions by offering a series of therapies, such as herbal medicine, meditation, acupuncture, and osteopathy in the early stage of illness development, or "un-illness status".

The term "un-illness status" is used in the "Medical Classic of Emperor Huang". The proposed prescriptions, such as shadow boxing and rhythmic gymnastics, are still effective for the present management of most lifestyle-related chronic diseases, such as hypertension, hyperlipidaemia, diabetes, obesity, hyperuricaemia, arteriosclerosis, osteoporosis, hepatitis, asymptomatic stroke, potential heart attack, and fatty liver. The Japan Mibyou System Association (JMSA) was founded in 2005, and it defined "mibyou" (un-illness) as a situation between health and disease (JMSA, 2006). JMSA's mission is committed to the better control of un-illness situations and to improving human wellness. JMSA established an official accreditation system for health promoters that aim to provide wholesome and secure care to all citizens.

A "challenge to 100 years of age" project in the county community of Nishi-Aizu in central Japan has been mobilized since 1994 with governmental financial support of 2 billion Japanese yen (Nishi-Aizu, 2003). The fundamental goal is to increase healthy longevity by providing a total solution package to villagers. They built an ICT infrastructure, improved the soil, enhanced the nutritional balance, and initiated a health promotion campaign.

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A Scalable Healthcare Integrated Platform (SHIP) and Key Technologies for Daily Application

Homecare devices were distributed to 687 families among a total of 2,819 families. These devices can measure blood pressure, ECG, photoelectric plethysmograms, body temperature and weight, and can receive answers to queries. These endeavours increased the villagers' longevity from 73.1 years (80.0 years for females) in 1985 to 77.6 years (84.1 years for females) in 2000 and decreased the mortality from stomach cancer from 138.9% (125.4% for females) in 1988 to 91.7% (66.7% for females) in 2002.

In their "China study", Campbell et al. described a monumental survey of diet and death rates from cancer in more than 2,400 Chinese counties, and its significance and implications for nutrition and health. They showed that by changing their behaviour based on nutritional balance and lifestyle, patients can dramatically reduce their risk of cancer, diabetes, heart disease, and obesity (Campbell & Campbell, 2005).

Both oriental and occidental professional societies have now converged to a common understanding that many chronic problems can be avoided through practicing a total solution package, including nutritional balance, rhythmic lifestyle, duly exercise, and mental wellness.

With an increasing global penetration of mobile telephony and a mature ICT infrastructure, as illustrated in Fig. 16, a future SHIP must be built by integrating modern approaches, such as network technology, wearable/invisible sensory instrumentation, and data mining mathematics, with ancient wisdom from both oriental and occidental philosophy. An overall strategy and tactics for healthcare and allied applications can then be provided. Many new areas and industries, such as public transportation security, fire fighting and police, senior health insurance, professional athlete training, home-based care, and the tracking of the effects of pharmaceuticals will blossom.



Fig. 16. Conceptual illustration of a future SHIP showing its constitutional fundamentals and application domains. This will be built by merging oriental holistic philosophy and occidental accurate treatment and the latest achievements in network technology, wearable/invisible measurements, and data mining mathematics. The SHIP will be applied in the chronic illness prevention domain, and also in a variety of applications, wherever vital signs are helpful.

4. Conclusions

In a series of R&D projects, our initiative endeavour was focused on SHIP architecture design and system integration. We have developed several sensory technologies for vitalsign monitoring in environments of everyday living, and data mining algorithms for the interpretation of comprehensive large-scale data. However, much new knowledge remains to be discovered from long-term data mining. The open architecture of SHIP makes it possible to integrate diversified vital signs and data mining algorithms, either from ours or from cooperative partners.

We also realize that radical societal change demands an active paradigm shift from acute disease treatment based in hospitals towards preventative care in lifelong activities, and in addition, individuals and also organizations and communities must be involved in daily efforts. We are optimistic in our belief that the global epidemic of chronic diseases can be relieved or controlled by using the latest research achievements, and by advocating healthy living behaviour, such as a positive attitude, sound nutritional balance, and regular physical exercise.

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Data Mining in Medical and Biological Research

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This book intends to bring together the most recent advances and applications of data mining research in the promising areas of medicine and biology from around the world. It consists of seventeen chapters, twelve related to medical research and five focused on the biological domain, which describe interesting applications, motivating progress and worthwhile results. We hope that the readers will benefit from this book and consider it as an excellent way to keep pace with the vast and diverse advances of new research efforts.

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