

VIII. 研究成果の刊行に関する一覧表

雑誌

発表者氏名	論文タイトル名	発表誌名	巻号	ページ	出版年
小川 彰	いわて新医療モデルと遠隔医療	日本遠隔医療学会雑誌	第9巻1号	in press	2013
赤坂 俊英, 高橋 和宏	三陸沿岸部被災地域との皮膚科遠隔診療の試み	日本遠隔医療学会雑誌	第9巻1号	in press	2013
高橋 義彦, 佐藤 譲	岩手医科大学と県立宮古病院の間の糖尿病遠隔診療支援	日本遠隔医療学会雑誌	第9巻1号	in press	2013
菅井 有, 澤井 高志	岩手県における遠隔病理診断の現状と今後の展望	日本遠隔医療学会雑誌	第9巻1号	in press	2013
江原 茂	遠隔画像診断を発展させた岩手県広域ネットワークによる画像情報連携にむけて	日本遠隔医療学会雑誌	第9巻1号	in press	2013
小山耕太郎	モバイルネットワーク環境における新生児心臓病の超音波動画像遠隔診断	日本遠隔医療学会雑誌	第9巻1号	in press	2013
東福寺幾夫, 澤井高志	バーチャルスライドの利用と標準化に関する調査報告	日本遠隔医療学会雑誌	8	19-24	2012
中山育徳, 松村翼, 赤坂俊英, 澤井高志	皮膚科領域におけるvirtual slideを利用した遠隔病理診断用コンサルテーションシステムの開発	岩手医誌	64	173-182	2012
高橋信, 佐藤陽子, 中野智, 早田航, 小山耕太郎, 千田勝一	左冠動脈開口部狭窄による運動時失神	日児誌	122	in press	2013
高橋信, 中野智, 佐藤陽子, 早田航, 小山耕太郎, 千田勝一	重複僧帽弁口を合併した心室中隔欠損症の臨床経過	日超医誌	40	in press	2013
早田航, 小山耕太郎	古くて新しい心臓病-先天性心疾患-Fallot四徴症	成人病と生活習慣	43	212-217	2013
中野智, 小山耕太郎	成人期の心室中隔欠損症-短絡率だけが手術適応を決めるのか	心エコー	14	54-59	2013
小山耕太郎	修正大血管転位症の診断	日小循誌	28	73-80	2012
小山耕太郎	手術を受けた先天性心疾患に対する心エコー: Fallot四徴症修復術後	心エコー	13	296-303	2012
小山耕太郎	東日本大震災・津波と岩手県の医療情報連携・遠隔医療	Rad Fan	10	24-26	2012
高木基宏, 藤井寛, 小山耕太郎, 大平隆, 柿沼博一, 藤野雄一, 澤井高志, 猪飼秋夫	遠隔画像診断のためのSVC符号化された心臓超音波画像の主観画像評価	信学技報	111	239-244	2012

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江原茂	遠隔画像診断ガイドラインの目指す画像診断のあり方	Japanese J of Clinical Radiology	57	1141-1144	2012
Nakayama I, Matsumura T, Kamataki A, Uzuki M, Saito K, Hobbs J, Akasaka T, Sawai T	Development of a teledermatopathology consultation system using virtual slides	Diagnostic Pathology	7	177-184	2012
Seung Park, Anil Parwani, Raymond D. Aller, Lech Banach, Michael J. Becich, Stephan Borkenfel, Alexis B. Carter, Bruce A. Friedman, Marcial Garcia Rojo, Andrew Georgiou, Gian Kayser, Klaus Kayser, Michael Legg, Christopher Naugler, Takashi Sawai, Hal Weiner, Dennis Winsten, Liron Pantanowitz	The History of Pathology Informatics: A Global Perspective	Journal of Pathology Informatics		in press	2013

書籍

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小山耕太郎	修正大血管転位症	竹中克、戸 出浩之編	心エコーハ ンドブック・ 先天性心 疾患	金芳堂	京都	2013	in press
<u>小山耕太郎</u>	成人期の先天性心疾 患:修正大血管転位 症	新垣義夫、 深谷隆	新・心臓病 プラクティ ス	文光堂	東京	2012	244-249

IX. 研究成果の刊行物・別刷



RESEARCH

Open Access

Development of a teledermatopathology consultation system using virtual slides

Ikunori Nakayama^{1,2,3}, Tsubasa Matsumura², Akihisa Kamataki², Miwa Uzuki², Kenji Saito⁴, James Hobbs⁵, Toshihide Akasaka³ and Takashi Sawai^{2*}

Abstract

Background: An online consultation system using virtual slides (whole slide images; WSI) has been developed for pathological diagnosis, and could help compensate for the shortage of pathologists, especially in the field of dermatopathology and in other fields dealing with difficult cases. This study focused on the performance and future potential of the system.

Method: In our system, histological specimens on slide glasses are digitalized by a virtual slide instrument, converted into web data, and up-loaded to an open server. Using our own purpose-built online system, we then input patient details such as age, gender, affected region, clinical data, past history and other related items. We next select up to ten consultants. Finally we send an e-mail to all consultants simultaneously through a single command. The consultant receives an e-mail containing an ID and password which is used to access the open server and inspect the images and other data associated with the case. The consultant makes a diagnosis, which is sent to us along with comments.

Because this was a pilot study, we also conducted several questionnaires with consultants concerning the quality of images, operability, usability, and other issues.

Results: We solicited consultations for 36 cases, including cases of tumor, and involving one to eight consultants in the field of dermatopathology. No problems were noted concerning the images or the functioning of the system on the sender or receiver sides. The quickest diagnosis was received only 18 minutes after sending our data. This is much faster than in conventional consultation using glass slides. There were no major problems relating to the diagnosis, although there were some minor differences of opinion between consultants. The results of questionnaires answered by many consultants confirmed the usability of this system for pathological consultation. (16 out of 23 consultants.)

Conclusion: We have developed a novel teledermatopathological consultation system using virtual slides, and investigated the usefulness of the system. The results demonstrate that our system can be a useful tool for international medical work, and we anticipate its wider application in the future.

Virtual slides: The virtual slides for this article can be found here: <http://www.diagnosticpathology.diagnomx.eu/vs/1902376044831574>

Keywords: Telepathology, Teledermatology, Consultation, Virtual slide, Whole slide image

* Correspondence: sawai@wonder.ocn.ne.jp

²Division of Leading Pathophysiology, Department of Pathology, Iwate Medical University School of Medicine, Yahaba, Iwate, Japan
Full list of author information is available at the end of the article



Background

Skin diseases vary widely from conditions that can interfere with social activity because of cosmetic disorders, such as acne and alopecia, to diseases that affect patient prognosis and quality of life, such as malignant melanoma or basal cell cancer, as well as various chronic inflammatory diseases. Achieving an accurate diagnosis and deciding on an appropriate procedure for therapy often requires microscopic as well as macroscopic findings, but very few dermatologists in Japan engage in histopathological diagnosis, and very few pathologists specialize in dermatopathology [1,2]. Furthermore, given the disproportionate availability of physicians and medical facilities between densely populated urban areas and more sparsely populated areas such as mountain or coastal areas, access to dermatological care is often woefully inadequate for achieving accurate diagnoses. To compensate for this in Japan, conventional consultation based on the delivery of glass slides has been available. This is a labor-, time-, and cost-intensive system, however, because many slides must be prepared and then delivered to experts. Delays in diagnosis and slide loss or damage are also major concerns.

Recently, in an alternative consultation system, image data of digitalized pathological findings have been transmitted to remote experts via the Internet [3,4], and also via mobile phone with static images and short movies [5].

However, compared to actual microscopic observation, the visual field, magnification and focus of the static images cannot be adjusted, making this more time-consuming and stressful than routine non-telepathological diagnosis [6,7].

The use of virtual slide (VS), especially recent whole slide image (WSI) makes it possible to select the visual field and magnification, and even adjust focus, resulting in diagnostic accuracy comparable to that achieved by conventional optical microscopy [7-14]. The use of VS has contributed to a gradual increase in consultations in dermatopathology as well as in other fields. However, the consultation system has continued to rely on the traditional communication tools of e-mail and facsimile. This system does not lend itself to the systematic organization and recording of consultation case details and diagnoses, and results in delays finding pathologists in specific fields suited to particular cases. Furthermore, it is not easy to compare the diagnosis received from one consultant with those returned by others and with previous cases.

In this study we developed a consultation system combining VS with a web application offering access to many consultants in a range of professional fields. The effectiveness of the consultation system for dermatological cases was compared with that of conventional

consultations using glass slides, and with conventional telepathology using static images.

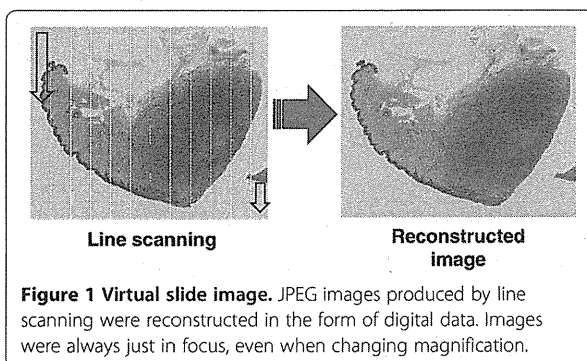
This study focused not on the accuracy of diagnoses obtained using VS, which has been fully evaluated in recent studies [7-14], but on the performance and future potential of this new consultation.

Methods

The consultation system

The system is composed of telecommunication lines, a VS system, and a public web server for consultations. Telecommunication lines included the local area network at Iwate Medical University (100BASE-TX), used for actions such as uploading VS images, as well as B Flet's® (best-effort 100 Mbps) fiber optic lines; Nippon Telegraph and Telephone Corporation EAST (NTT EAST; Tokyo, Japan) lines were used for external communication for transmission and reception via the public server. A Scan Scope CS® scanner (Aperio Technologies, Vista, CA) was used for the VS system. Resolution could be set to an objective lens magnification of 20× (0.50 μm/pixel) or 40× (0.25 μm/pixel). Diagnosis could usually be achieved at 20× magnification, but 40× magnification was used for scanning when detailed observation of cellular morphology was required for distinction of poorly differentiated or undifferentiated malignancies, or leukemia. The computer was a Workstation 4300 (Hewlett-Packard Development Company L.P., Palo Alto, CA), equipped with the Windows XP® operating system, an Intel Pentium D® 3.20 GHz CPU, and an mvBlue Fox (Matrix Vision GmbH, Oppenweiler, Germany) graphics board with 1.64 GB RAM and an 800 GB HDD. VS images were displayed by reconstructing scanned JPEG 2000 digital images (Figure 1). Images that were always in focus were thus displayed on a monitor, and the visual field and magnification could be adjusted in much the same way as with a microscope.

A ProLiant Server DL120 G6 (Hewlett-Packard) with a FreeBSD operating system and an Intel Xeon® X3430 2.4 GHz CPU, with 4 GB RAM and a 640 GB HDD, was



used as the public web server for consultations. This included a database function for managing digital images and patient data (such as case number and medical history), and consultation-related web application functions, such as order input, digital image display, e-mail notifications from the sender site, response input from the receiver site (consultant), and response verification from the sender. This allowed the series of actions required for carrying out consultations to be implemented through a web browser not dependent on a particular operating system or dedicated software. As an additional function, VS images could be displayed up to five at a time, and static images (macroscopic photographs, X-rays, medical charts, and other documents needed for diagnosis) could be displayed up to three at a time. In addition, up to 10 pre-registered diagnosticians could be selected, and a consultation request could be sent to all in a single action (Figure 2).

To protect patient identity and personal details, information was limited to age, region, and clinical diagnosis.

Questionnaire survey about our system

Forty-two pathologists and dermatologists were surveyed by questionnaire on items such as the content, operability, and image quality of the system. Individual e-mail requests to fill out the questionnaire were sent, and responses were returned by either e-mail or fax. The following items were surveyed:

- 1) The percentage of consultation cases among routinely diagnostic cases
- 2) Operability of VS displayed on the web browser;
- 3) Image quality, especially for diagnosis; and
- 4) Convenience and usability of this system for supporting their work.

Results

The consultation system

Our system was used to consult 10 pathologists or dermatologists about skin diseases. Diagnostic consultations were requested in 36 cases (13 men, 23 women) (Table 1). Generally the data volume of VS was

Figure 2 Ordering page. a) Click the thumbnail image of a VS to display the image. b) Input patient information (sex, age, region, clinical diagnosis, comments). c) Select VS to send (up to 5 slides). d) Add static images such as previous histological figures, X-ray films or macroscopic photographs necessary for patient diagnosis (up to 3). e) Select consultants from the list (up to 10).

proportional to the scanning area and resolution, usually at about 150–300 MB with an area of about 15×15 mm² and a resolution corresponding to an objective lens magnification of 20 power. Consultations covered neoplastic diseases that were identified or determined to be benign or malignant in 20 cases, and non-neoplastic diseases not amenable to definitive diagnosis involving rare inflammatory, degenerative, or congenital disease in 16 cases. In 24 cases 1 VS image was referred for diagnosis, in 2 cases 2 slides, and in 10 cases 3 or more slides. Macroscopic photographs were attached in 2 cases. Other laboratory data were also sent in 1 case. Most consultations were with dermatologists in Iwate Medical University Hospital and in other universities or medical facilities throughout Japan. Thirty cases were sent to 1 consultant, 1 case to 3, 1 case to 4, 2 cases to 5, and 2 cases to 8 consultants. The fastest response time was 18 minutes. Some cases were not answered by all pathologists. All pathological diagnoses from consultants were relayed with their comments to clinicians. However, after considering all diagnoses and comments, the final responsibility for critical pathological diagnosis relating to patients' therapy usually rested with the sender pathologist. Cases in which diagnoses differed between multiple consultants were reviewed and left to the discretion of the requesting side.

Questionnaire survey

The questionnaire was answered by 54.8% of participants (23/42). Among all routinely diagnostic cases, 0.6% cases were transferred to experts inside Japan. Of those, 20% were of dermatological disease (Figure 3a). Concerning evaluation of VS, 78% (18/23) answered that there was no difficulty with operability, and no experts noted poor operability (Figure 3b), while 74% (17/23) considered that the image quality of VS was suitable/for diagnosis (Figure 3c). Seventy percent of all participants (16/23) answered that they were interested in using this system themselves for work requiring consultation (Figure 3d). Many pathologists/dermatopathologists answered that this system is convenient as it eliminates the need to package and send prepared slides, several consultants can be involved at the same time, and prompt responses can be obtained.

Discussion

Telemedical applications in the field of dermatology have been attempted in various ways. In 1995, Perednia et al. proposed teledermatology, in which dermatological macroscopic findings were captured by digital camera, and sent remotely via the Internet from personal computer for consultation [15]. The usefulness and some associated problems of this system have been reported [16-18]. The subsequent widespread use of digital

devices and personal computers, together with the development of telecommunications, has led to the development [19] and gradual popularization [20-22] of teledermatopathology, which involves the digitization and remote transmission of microscopic dermatopathological findings. The use of dermatological consultation based on VS has been reported by Massore et al. [8,12], followed by Mooney et al. [9].

The application of teliagnosis in the field of pathology is relatively advanced, and significant contributions to the development of telepathology have been made by Weinstein et al. in the USA beginning in the early 1980s [23-25], Kaiser et al. in Europe [26,27], and by Sawai et al. in Japan from the early 1990s [28]. Along with the shift from static to robotic images and from analog to digital lines, recent developments in IT led to the emergence of the worldwide use of VS, especially in the fields of diagnosis and education [29,30], followed by diagnostic developments such as automated diagnosis of histological screening via the Internet [31]. Furthermore, efforts have been made to reduce image data volume as much as possible [32].

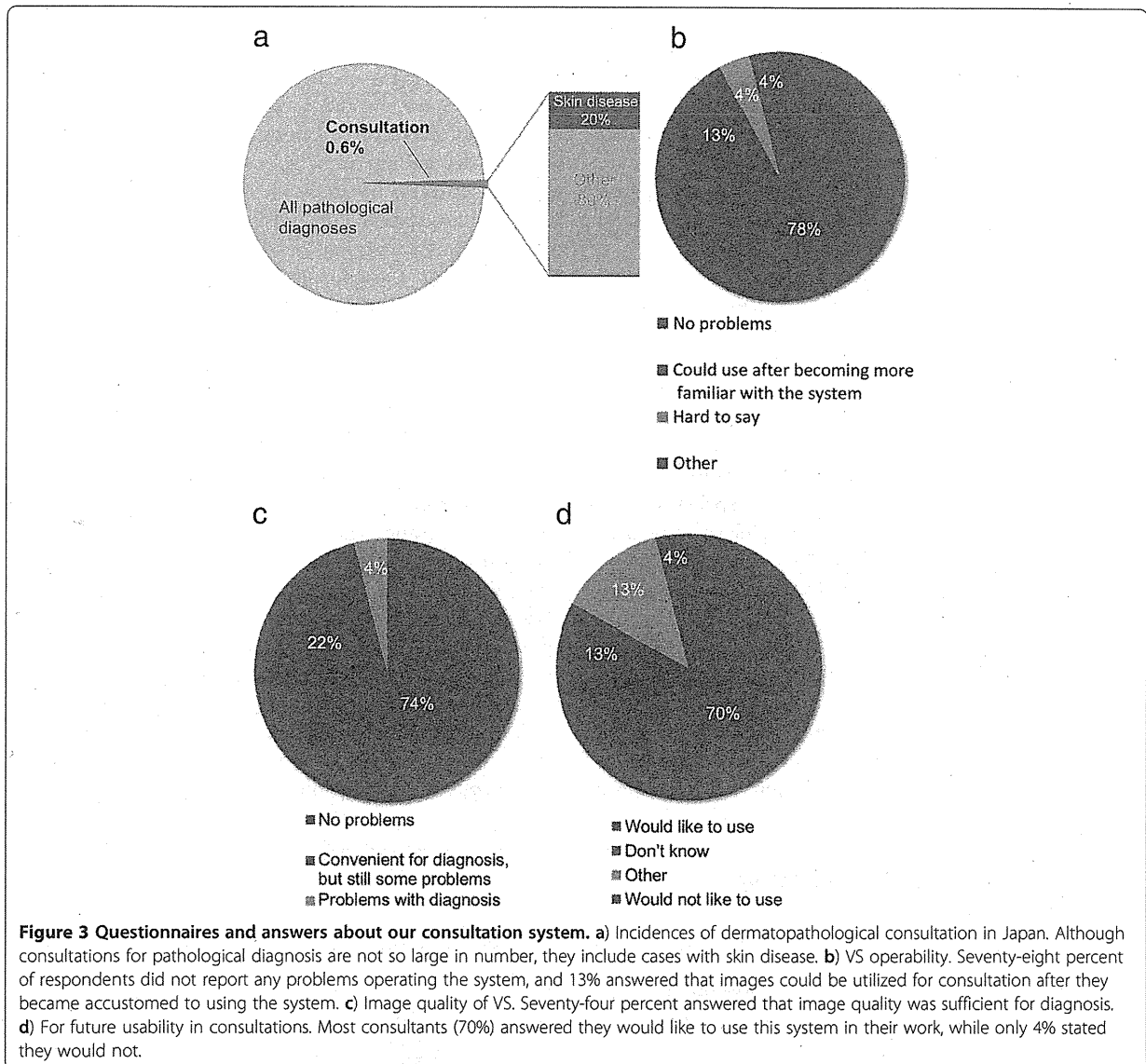
Thirty or more companies around the world now deal with VS systems, and several recent devices have been rated as having rapid scanning capabilities and high image quality [33].

With the increased availability of VS, many institutions have begun applying VS for telepathology. Furthermore, modern high-volume and high-speed communication lines facilitate the use of high-resolution WSI. However, in terms of magnification, focus adjustment, and selection of specific points on a slide, diagnosis using WSI still cannot match the performance achieved with a microscope and glass slides [34-36]. However, our system is far more convenient than conventional microscopes and slides in terms of the time and manpower required, and shipping costs are avoided, although of course initial set-up costs are incurred.

Many medical universities and laboratory institutes now offer consultation services using VS data [37]. The Medical Electronic Consultation Expert System (MECES) based on an open platform has performed in Europe [26]. This is an Internet communication service based on grid technology [38].

Although there may be local consultation systems using static and robotic images, until now a consultation system using VS has not been available in Japan [39]. This is the first multi-function VS system equipped to handle both domestic and international consultations.

The system that we have developed not only shows VS images, but also facilitates within a single system the communication of opinions involving clinical and pathological data necessary for diagnosis. The most characteristic point of our system is that data for diagnosis



involving scanned images and other related data can be sent out at the same time via the Internet to multiple consultants. One of the biggest advantages of consulting many pathologists is that we can obtain various opinions reflecting the professional field and interests of each pathologist. Using our system we consulted more than one consultant in six cases, because of uncertainty about our diagnosis, or concern over difficult cases outside our professional field. For general pathologists, diagnosis in the very specific field of dermatopathology is rather difficult compared with other fields such as diseases of the gastrointestinal tract and respiratory tract. So far we have not changed our diagnosis on the basis of replies from consultants, but when consultants returned replies consistent with our own diagnosis, this gave us added

confidence to communicate the diagnosis to the clinician. For this reason, this system could be particularly helpful for pathologists working alone in a single institute. Even if our diagnosis could not consistent with the diagnoses of consultants, we could learn much about the process of how to determine the diagnosis, and could obtain new knowledge from experienced pathologists. In the case of differences in diagnosis, we can consider the opinions of other consultants, but we must take final responsibility for making a diagnosis and informing the clinicians.

This system is much faster, more economical, and more convenient than the conventional method of directly mailing glass slides. Until now it required from 5 to 10 days to receive a diagnosis because of the delays

associated with preparing, packaging and sending slide glasses by mail. In our VS system the fastest diagnosis was returned only 18 minutes after sending. Quick responses are obviously helpful for patients, as well as for pathologists and for clinicians who must administer therapy. The waiting time until therapeutic decisions are made is a big source of stress for patients. Quick diagnosis using telepathology relieves this stress and so gives satisfaction to patients as well as to pathologists.

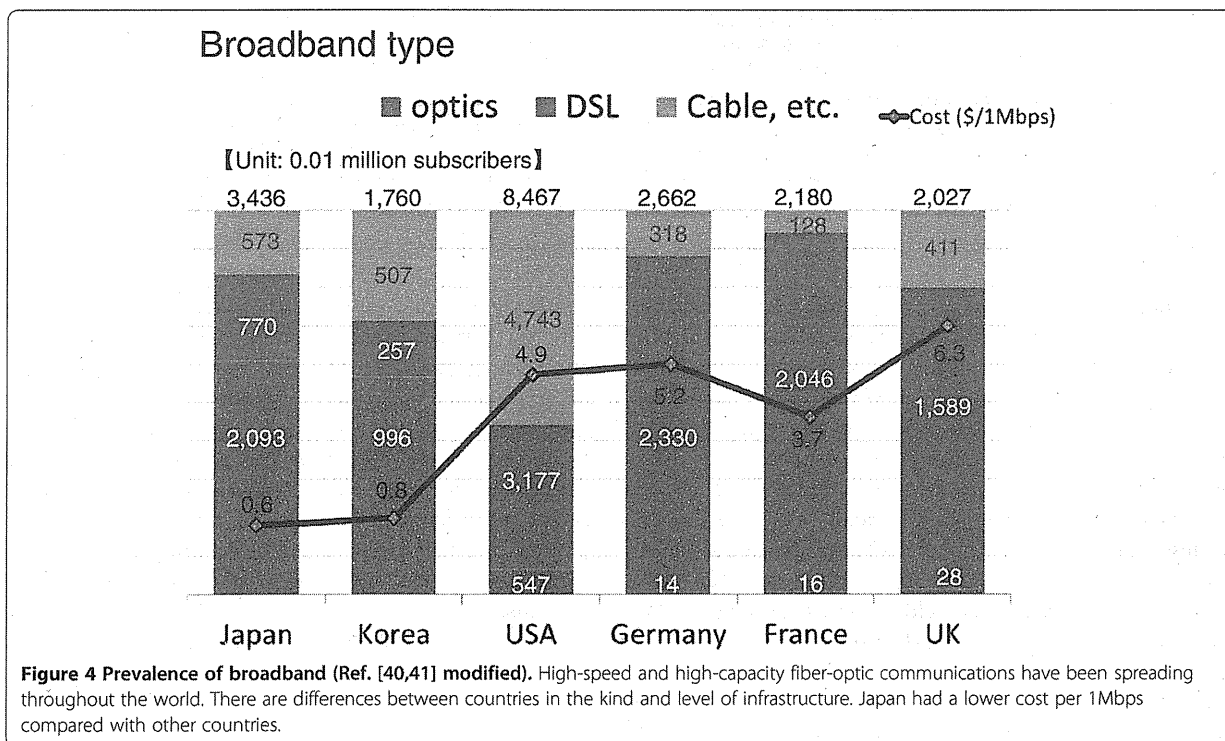
This system is now in development for practical application overseas. Despite time differences, the answer from Yanbian University in China was returned only 16 hours after sending the request. Although it sometimes took several days to receive a response, this delay mainly resulted from consultant dermatologists and pathologists being too busy to log on to their personal computers (personal communication), or from cases for consultation being fairly difficult even for specialists and thus requiring time for investigation. Delays were thus probably not due to the poor quality of the consultation system, but to factors related to rare cases or to the consultants' busy schedules.

The VS data ranged in size from 150 MB to 300 MB, far greater than that in the case of static images. Therefore, a high-performance personal computer and high-speed telecommunication lines are required for rapid monitor display. The infrastructure in Japan is characterized by the wide availability of inexpensive, high-speed

broadband (Figure 4) [40,41] and offers an environment well-suited to mass data transmission and reception.

Major advantages of this system are that the integration of a VS system and a consultation web application allows diagnoses to be requested via only a web browser, without depending on a specific operating system or specific software; up to 5 VS images can be sent at a time; and requests for diagnosis can be sent to up to 10 diagnosticians within and outside Japan at the same time (Figure 2), and the consultants can do everything from accessing the data to making the diagnosis and providing a response, anywhere and at anytime, using only a standard personal computer and public networks. Thus, the system is advanced in that the entire procedure can be managed on a web browser through a personal computer, from the preparation of the digital images and patient information and transmission of image data, to the retrieval of data and the transmission of answers by the receiver side, and again in the verification of the response on the sender side. The results of the questionnaire showed that more than 70% of consultants felt there were no problems with the quality of VS images and operability (Figure 3b, c), and that 70% were interested in using the system for their own work (Figure 3d).

Making our consultation system more widely available in the future will require more powerful personal computers, lower costs, and provision of financial resources for payment for diagnosticians, as well as public



insurance coverage, standardization of formats of VS data, and more widespread use of high-speed telecommunication lines.

The ramifications of crossing national boundaries also remain to be considered. Our study is still a pilot study aimed at improving usability and overcoming any other problems with using this system. If the system is well received, we will then have to consider problems such as licensure, quality control, billing and medico-legal liability across national boundaries.

We nevertheless hope to be able to implement the system on a wider basis both to facilitate conventional consultation about difficult and uncertain cases, and to better facilitate the establishment of international diagnostic criteria by increasing international collaboration between many specialists without the need for the time and cost associated with participation in international conferences and other gatherings. However, the wider implementation of this system depends crucially on the level of IT infrastructure and the specifications of personal computers. With continued progress in IT infrastructure, we believe that many pathologists throughout the world will be able to apply this system for diagnosis.

Conclusion

We have developed a novel teledermatopathological consultation system using VS, and investigated the operability, convenience, and image quality when attaching the necessary data related to diagnosis and transferring them via fiber-optic lines to consultants. The results demonstrated that our consultation system is a useful tool not only for dermatopathology but also for clinical dermatology in the future. However, the current study had its limitations, and in future we hope to conduct a more rigorous study into the effectiveness of our system compared with conventional glass slides, and with other VS systems.

Competing interests

The authors declare that they have no competing interests.

Authors' contributions

Performed the experiments and wrote the paper: TM and IN. Revised the paper: AK and MU. Supervised the developer of the system: KS. Assisted with writing the manuscript: JH. Participated the experiments: TA. Participated in the conception of this study, designed the experiments and final approval of the article: TS. All authors read and approved the final manuscript.

Authors' information

First author: Ikunori Nakayama.
Co-first author: Tsubasa Matsumura.

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Author details

¹Department of Dermatology, Course of Integrated Medicine, Graduate School of Medicine, Osaka University, Suita, Osaka, Japan. ²Division of

Leading Pathophysiology, Department of Pathology, Iwate Medical University School of Medicine, Yahaba, Iwate, Japan. ³Department of Dermatology, Iwate Medical University School of Medicine, Morioka, Iwate, Japan. ⁴Information Center for Iwate Medical University, Morioka, Iwate, Japan. ⁵Department of English, Center for Liberal Arts and Sciences, Iwate Medical University, Yahaba, Iwate, Japan.

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The History of Pathology Informatics: A Global Perspective

Seung Park, MD¹; Anil V. Parwani, MD, PhD¹; Raymond D. Aller, MD²; Lech Banach, MD³; Michael J. Becich, MD, PhD⁴; Stephan Borkenfeld⁵; Alexis B. Carter, MD⁶; Bruce A. Friedman, MD⁷; Marcial Garcia Rojo, MD⁸; Andrew Georgiou, PhD⁹; Gian Kayser, MD¹⁰; Klaus Kayser, MD, PhD¹¹; Michael Legg, PhD¹²; Christopher Naugler, MD¹³; Takashi Sawai, MD¹⁴; Hal Weiner, MBA¹⁵; Dennis Winsten, MS¹⁶; Liron Pantanowitz, MD¹

¹Department of Pathology, University of Pittsburgh Medical Center, Pittsburgh, PA, USA

²Department of Pathology, Keck School of Medicine, University of Southern California, Los Angeles, CA, USA

⁴Department of Biomedical Informatics, University of Pittsburgh School of Medicine, Pittsburgh, PA, USA

³NHLS and Department of Anatomical Pathology, Walter Sisulu University, Mthatha, South Africa

⁵International Academy of Telepathology, Heidelberg, Germany

⁶Department of Pathology and Laboratory Medicine, Department of Biomedical Informatics, Emory University School of Medicine, Atlanta, GA, USA

⁷Department of Pathology, University of Michigan Medical School, Ann Arbor, MI, USA

⁸Pathology Department, Hospital General de Ciudad Real, Ciudad Real, Spain

⁹Centre for Health Systems and Safety Research, Australian Institute of Health Innovation, Faculty of Medicine, University of New South Wales, NSW, Australia

¹⁰Institute of Pathology, University Freiburg, Freiburg, Germany

¹¹Institute of Pathology, Charite, Berlin, Germany

¹²Centre for Health Informatics and e-Health Research, University of Wollongong, Wollongong, NSW, Australia

¹³ Department of Pathology and Laboratory Medicine, University of Calgary, and Calgary Laboratory Services, Calgary, Alberta, Canada

¹⁴Department of Pathology, School of Medicine, Iwate Medical University, Iwate Prefecture, Japan

¹⁵Weiner Consulting Services, Florence, OR, USA

¹⁶Dennis Winsten & Associates, Inc., Healthcare Systems Consultants, Tucson, AZ, USA

Department of Pathology of University of Sao Paulo in Brazil, the Hospital Británico de Buenos Aires in Argentina, and Centro Consulenze Anatomia Patologica in Milano, Italy.^[160]

3.2.3.2 Teaching and Continuing Medical Education

In 2000, a website containing a comprehensive collection of histopathologic images with a special focus on oral pathology was published by the Fundação Odontológica de Ribeirão Preto (FUNORP), Universidade de São Paulo.^[161] Also in 2000, the University of Cauca served as mirror site for the 6th Internet World Congress for Biomedical Sciences, organized by the Pathology Department of Hospital de Ciudad Real in Spain.^[162] Since June 2002, autopsies have been broadcast online on a weekly basis, with the participation of 12 Brazilian medical schools.^[163]

In 2004 in Uruguay, the Pathology Department of the Medical School of the Hospital de Clínicas "Dr. Manuel Quintela" in Montevideo, Uruguay started publishing online study material for medical students.^[164] In 2005, the Virtual Hispano-American Congress of Pathology began utilizing WSIs instead of static images in its presentations. Finally, at present, iPath hosts the Telemedicina Sur telemedicine network, active in South-American countries for medical discussions, including pathology continuing medical education (CME) and consultations.

4. Asia

The progress of pathology informatics in Asia has been much like the phenomenon of watching ripples spread across the surface of a once-placid pond after a pebble has been thrown into it. The pond in this metaphor is Asia; the pebble represents progress in telepathology and WSI from the West. Although this historical review of informatics in Asia is focused largely on advances in digital imaging, much progress has been achieved in these countries utilizing computers to establish laboratory information systems. In general, development of digitized telepathology was supported by the development of

computers, and improvements in the performance of digital cameras. In the modern era, WSI have been a primary focus of pathology informatics activity across the world. Asia is no different in this respect. In some Asian countries (e.g. Japan), where network infrastructure and high-speed Internet-based telemedicine are well developed, WSI systems are in heavy use. In other Asian countries (e.g. China), there are significant bottlenecks to further penetration of telepathology, including (i) low levels of understanding in society in general about the importance of pathological diagnosis, (ii) physical constraints, including infrastructure development not keeping up in large geographic areas, (iii) high prices of WSI systems, (iv) lack of mutual trust between pathologists in different areas, and (v) regulatory issues.

The story of digital imaging in pathology was, in its earliest years, confined largely to the USA and Europe. With the development of the Internet came the possibility of sending and receiving digital images across the world; most historians of our still-nascent field trace the lion's share of the evolution of the current state of telepathology – and indeed pathology at large – to this singularly disruptive event. Many organizations – such as the AITN – sprang up in the so-called “Web 1.0” era, providing platforms for diagnoses and consultations based on international telepathology involving not only the USA, but also many other nations, including China and Japan.^[165] While these early efforts uniformly used static telepathology as their primary diagnostic modality, in the modern era we have seen a shift to the usage of WSI instead.^[166] In Asia, the story of true digital pathology has just begun; it currently lags far behind the more developed state of digital pathology among the Western nations. However, Asian nations – particularly those with advanced network infrastructures like Japan and South Korea – are making more and more use of digital pathology as broadband saturation in these countries have reached (and indeed by now have exceeded) 100%. More recently, fast-growing economies like China and India have been pushing forward with digitization. Iran and Uzbekistan are also promoting digital pathology.^{[167][168]}

Telepathology options differ from country to country: offerings run the gamut from relatively slow transfer of static images taken by digital cameras via digital subscriber line (DSL) to nearly-instantaneous transfer of WSIs via fiber optic networks.^{[169][170][171]} Governmental support for telepathology and digital pathology is also quite variable – some countries have embraced these new technologies as quickly as they are introduced, whereas others have applied heavy regulation that has effectively stifled the growth of digital pathology in those nations. A case in point is the comparison between Japan and South Korea: although both countries have impressive network infrastructures (South Korea’s broadband penetration approached 100% as of 2012), the uptake of digital pathology in South Korea has been relatively slow due to an onerous regulatory environment. Compare this to the governmental policies of Japan, which openly promote a “standardization of cancer medical services” based on WSIs as well as other medical advances. It should therefore come as no surprise that Japan’s growth in telepathology and WSI adoption is outstanding as compared to that of South Korea – a nation that not only has a smaller landmass, but also an arguably better-developed network infrastructure (Figure 9).^[172]

4.1 Japan

Japan’s network infrastructure is among the best-developed in the world. Population coverage and network speeds also rank among the highest in the world - it is worth noting that fiber optics is a common connectivity option even among general households! High-speed network-based telemedicine has been developed to such a level that intraoperative rapid diagnosis and consultation take place actively in the field of pathological diagnosis. The first reports of digital pathology in Japan date from the first half of the 1990s. At first, static images were the major telepathology modality; now, real-time remote control of robotic microscopes and access to WSI is the norm. The essential driver of this change is widely accepted to have been the government’s policymaking.

4.1.1 Infrastructure

Telepathology in Japan was first conducted on an analog system. It started shifting to digital modalities in approximately 1996 by using the ISDN protocol, which was the first step toward full implementation. In 2001 and 2002, asymmetric digital subscriber line (ADSL) and fiber optics, respectively, were implemented in telepathology. The advances in transmission technology combined with wide spread digitization made it possible to transfer still images and videos of tissues for pathological diagnosis. With the more recent addition of Hi-Vision (HDTV) technology, intraoperative rapid diagnosis is performed utilizing dynamic methods with full remote control of a robotic microscope.^{[173][10]} WSI is also utilized for consultation and second opinions, while their application in medical education is expanding. In 2009 and 2010, the high-speed satellite “Kizuna” was used for the first-ever Japanese fully dynamic/WSI telepathology study via satellite; this study allowed for simultaneous live telepresence across three sites (Iwate, Tokyo, and Okinawa).^[174]

4.1.2 Digitization

In Japan, the static, dynamic (live video feed without control of the microscope), fully dynamic (live video feed with direct control of robotic microscope), and WSI methods of digital pathology are all in use. As of today, two Japanese providers offer fully dynamic and/or WSI methods. The Ministry of Health, Labour and Welfare has provided (and continues to provide) half of the funds necessary to procure WSI scanners and other such equipment at institutes and hospitals across the nation. The total number of WSI systems deployed in Japan is approximately 400, most of which are provided by Hamamatsu Photonics and Olympus. The use of WSI has taken root not only for pathological diagnosis, but also for education. The usage rate in medical faculties of Japanese universities for teaching histology and pathology is 46%.^[175] Most of the universities utilize WSI in combination with existing microscopes; however, depending on the content of the lectures, some have fully shifted to WSI.^[176] Although a complete shift from microscopes to WSI still requires validation of their educational effectiveness, WSI

has been highly praised by students and researchers alike, as they allow more than one user to look at a specimen simultaneously and to conduct discussions among themselves. WSI is also more flexible compared with traditional microscopes.

4.2 China

Expectations for telemedicine including telepathology are very high in China, which is a country with an extensive national territory. Telepathology, however, is not currently actively practiced because (i) the infrastructure has not developed fast enough to cover all areas, (ii) hardware cost is still high, (iii) digital imagery is not fully trusted, (iv) people have a strong attachment to traditional optical microscopic diagnoses, (v) not enough physicians engage in telepathology, and (vi) state regulations concerning remote diagnosis are inadequate. In terms of infrastructure, DSL is still the dominant technology, but more recently some cases of telepathology are reported as using fiber optics and WSI.

4.2.1 Infrastructure and Equipment

Along with its recent outstanding economic growth, China has been rapidly expanding its infrastructure. The speed of development can be exemplified by the number of Internet users reaching 400 million in 2011 and the number of cell phone users reaching 900 million.^{[177][178]} Nevertheless, China's overall network infrastructure remains less developed than that of the USA, Japan, and South Korea. Although the absolute number of people who have access to the Internet is the highest in the world, if divided by China's large population, the penetration rate remains as low as 36.3%, as opposed to 100% in South Korea, 78.3% in North America, and 78.4% in Japan.^[179] Moreover, the digital divide in terms of Internet use between urban and rural areas is significant.

4.2.2 Digitization and Telepathology

Pathologists are scarce in China, particularly in the southwest region. To ascertain the telepathology situation in the country, we performed a PubMed search with the keywords, "Telepathology" and

“China.” There were five hits, three of which were related to consultation using the AITN, as reported by Weinstein. These telepathology cases utilized static telemicroscopy over the Internet. Telepathology efforts indigenous to China, however, began in the first half of the 1990s.^[165] Since the early 2000s, telepathology studies have been conducted based mainly on employing digitized still images and live video feeds without direct microscope control. These two diagnostic modalities appear to be the current mainstream in China. More recently, however, telepathology using fiber optics and WSI has tentatively begun between Peking University and its First Hospital. The most popular way of currently conducting telepathology in China is to either share WSI on a server with a trusted partner or to send an e-mail with a WSI as an attached file in the file transaction hub (FTH) format via DSL. In this format, the sender observes an image enlarged by a factor of 4, extracts his/her area of interest into a 40-times-larger WSI, attaches the image as an FTH file to an e-mail, and sends it to the consulting pathologist. The advantage of this process is that the WSI files are of relatively small size, between 2 and 30 MB. This represents a middle ground between WSI-based and static image-based approaches. The attempt started with validating the result of this mode of telepathology by comparing it to conventional optical microscopy using biopsy cases; the diagnoses were reported to show good agreement for all cases.^{[170][171]} Nevertheless, the use of static telepathology still remains more prevalent than dynamic methods. In addition, the issue of disparity between urban and rural areas remains unsolved in terms of limited infrastructure development and utilization of IT in hospital facilities. As such, the practice of telepathology in China is currently limited to certain institutes only.

4.3 India

Telepathology in India is generally limited to static telemicroscopy utilizing the Internet.^[169] Similar to China, constraints include the size of the country, the gap between urban and rural areas, startup cost, a power grid electrical supply system that is subject to occasional blackouts, and also the complex human relations among several groups.