

Figure 2: Annual numbers of HIV-1-infected persons in various high-risk groups reported to Taiwan Centers for Disease Control

via southeast China, Guangxi province, and Hong Kong.⁷⁻⁹ There have been enormous increases in the amount of heroin smuggled into Taiwan and in the number of intravenous drug users since 2002, when five intravenous drug users from southern Taiwan were diagnosed as the country's first HIV-1 seropositive cases infected with CRF07_BC.⁵ Even though the Hong Kong authorities identified three cases of CRF07_BC infection in 2001, a serious outbreak in that city's population of intravenous drug users is believed to have been blocked by a methadone maintenance programme.⁹

Clearly, close monitoring of emerging HIV-1 subtypes related to intravenous drug use and implementing harm-reduction programmes are vital to preventing similar outbreaks in other populations of intravenous drug users in neighbouring countries. In 2005, Alex Wodak, Jerry Stimson, and other harm-reduction experts were invited to Taiwan to share their experiences with government officials, medical field-workers, and public-health professionals. After careful study of harm-reduction programmes in place in Hong Kong and Australia, a pilot programme was started in four of Taiwan's 23 administrative areas in September, 2005. This programme has since been expanded nationally, and consists of 427 service sites for syringe exchange plus centres for methadone maintenance therapy, Free methadone is provided to HIV-1-infected intravenous drug users while HIV-1 seronegative intravenous drug

users have to pay about US\$1600 a year. The Taiwan Centers for Disease Control plans to provide methadone maintenance to intravenous drug users in prisons, and the country's Bureau of Controlled Drugs will start producing methadone to assist in the government's commitment to providing methadone maintenance to 30 000 intravenous drug users by 2009.

All parts of Asia are reporting rising numbers of HIV-positive and AIDS patients in male homosexuals and bisexuals. In Taiwan, HIV-1 infection rates in men who have sex with men in gay saunas in different cities currently range from 5.2% to 15.8%.10,11 The same population has high rates of syphilis, 8-1-13-8%, depending on the city. 10,11 Taiwanese male homosexual and bisexual HIV-1/AIDS patients have also been diagnosed with significantly higher rates of syphilis than have heterosexual patients.2 Furthermore, the percentage of homosexual or bisexual HIV-1/AIDS patients under the age of 20 years is significantly higher than that of heterosexual patients, 3.0% versus 1.7%.32 In addition to the stigmatisation of homosexuality in Taiwanese society, the lack of accurate information on homosexuality in sex education and on risk factors in AIDS education increases the risk of contracting HIV and other sexually transmitted infections within the country's population of men who have sex with men. Whilst a community-based prevention programme for such men has been developed by a group of academic and grass-roots non-governmental organisations, a current challenge is the implementation of this programme into a national programme, and making it a priority.

Taiwan's clinical spectrum of AIDS patients is similar to those reported in other developed countries, but significant differences have been noted in incidences of opportunistic infections. For example, the incidence of tuberculosis in patients with advanced illness is high in Taiwan (24-6%) and the rate of endemic fungal (*Penicillium marneffei*) infections is increasing.¹²⁻¹⁴ On the positive side, the effort by the Taiwanese Government since April, 1997, to distribute highly-active antiretroviral therapy for free¹⁵ has resulted in dramatic decreases in morbidity and mortality from HIV-1 infection.¹⁶

Because of their high background prevalence, HBV and HCV coinfections with HIV are particularly important in Asian countries in terms of HIV transmission via injecting drug use.^{12,18} In a survey of

www.thelancet.com Vol 369 February 24, 2007

459 intravenous drug users infected with HIV-1, one of us (Y-MAC) found that 456 (99-6%) also had anti-HCV antibodies and 77 (16-8%) were seropositive for HBsAg. The long-term impact of hepatitis coinfections on HIV and on morbidity and mortality from liver disease requires monitoring.

By the end of 2006, 19 confirmed cases of vertical HIV-1 transmission have been reported to the Taiwan Centers for Disease Control. In January, 2005, the agency started a national programme focused on prevention of mother-to-child transmission, and five cases of vertical transmission were reported in 2005. By June, 2006, the screening rate had reached 97-4%, and 47 of 338 452 pregnant women (13-9 per 100 000) tested in Taiwan have been identified as having HIV-1 infections and have received antiretroviral therapy to prevent mother-to-child transmission. To increase the participation rate, there is discussion of changing the voluntary counselling and testing strategy from opt in to opt out.

Several positive responses to the HIV/AIDS epidemic in Taiwan should be mentioned. In 1990 an AIDS Prevention and Control Law was passed to protect the rights of people with HIV/AIDS for treatment, education, and employment. Since 1992, 16 non-governmental organisations registered or established in Taiwan have provided shelter, care, counselling, anonymous testing, and AIDS education. One in particular, the People Living with HIV/AIDS Rights' Advocacy Association, has been addressing human rights issues related to HIV/AIDS since 1997. However, most such organisations have their headquarters and facilities in northern Taiwan, and two-thirds of the country's intravenous drug users live in central and southern parts. In addition, many social workers employed by non-governmental organisations are still unfamiliar with issues related to drug abuse and inexperienced in interacting with intravenous drug users. There is a clear and immediate need for counselling workshops for medical staff and social workers.

As the HIV-1 infection threat increases, there are many signs of persistent denial and resurgent discrimination in Taiwan. Several important issues need to be addressed: sentinel surveillance of female sex workers, social welfare institutions and housing for homeless people with HIV/AIDS, financial support for non-governmental organisations, training and re-education programmes aimed at changing the attitudes of medical staff toward

people with HIV/AIDS, and more funding for AIDS research, especially vaccine development.

*Yi-Ming Arthur Chen, Steve Hsu-Sung Kuo
AIDS Prevention and Research Center and Institute of Public
Health, National Yang-Ming University, 112 Taipei, Taiwan,
(Y-MAC); and Centers for Disease Control, Taipei, Taiwan (SH-SK)
Arthur@ym.edu.tw

We declare that we have no conflict of interest.

- Centers for Disease Control, R.O.C. (Taiwan). HIV/AIDS data. 2006: http://www.cdc.gov.tw/website_en/health%20topics/Communicable%20Diseases%208%20Prevention/issues%206%20HIV-AIDS/Statistics%20of%20HIV-AIDS/Ownload%20HIV-AIDS/S0Data.files/2006.files/9512Monthly%20Resport.pdf (accessed Jan 30, 2007).
- 2 Ministry of Health, People's Republic of China, Joint United Nations Programme on HIV/AIDS, World Health Organization. 2005 update on the HIV/AIDS epidemic and response in China. Jan 24, 2005: http://data.unaids. org/Publications/External-Documents/RP_2005ChinaEstimation_25Jan06_ en.pdf (accessed Jan 23, 2007).
- 3 Chen YM. Molecular epidemiology of HIV-1 Infection among Injecting drug users in Taiwan. 2005 Taipei International Conference on Drug Control and Addition Treatment, Taipei, Taiwan, Nov 22–24, 2005.
- Chen YM, Lan YC, Lai SF, Yang JY, Tsai SF, Kuo SH. HIV-1 CRF07_BC infections, injecting drug users, Taiwan. Emerg Infect Dis 2006; 12: 703-05.
- 5 Lin YT, Lan YC, Chen YJ, et al. Molecular epidemiology of HIV-1 infection and full-length genomic analysis of HIV-1 circulating recombinant form 07_BC strains from injecting drug users in Taiwan. J Infect Dis (in press).
- 6 Suit, Graf M, Zhang Y, et al. Characterization of a virtually full-length human immunodeficiency virus type 1 genome of a prevalent intersubtype (C/B') recombinant strain in China. J Virol 2000; 74: 11367-76.
- Piyasirisilp S, McCutchan FE, Carr JK, et al. A recent outbreak of human immunodeficiency virus type 1 infection in southern China was initiated by two highly homogeneous, geographically separated strains, circulating recombinant form AE and a novel BC recombinant. J Virol 2000; 74: 11286–95.
- 8 Cohen J. Asia and Africa; on different trajectories? Science 2004; 304: 1932-38.
- 9 Lim WL, Xing H, Wong KH, et al. The lack of an epidemiological link between HIV type 1 Infections in Hong Kong and Mainland China. AIDS Res Hum Retroviruses 2004; 20: 259–62.
- 10 Lai SF, Hong CP, Lan YC, et al. Molecular epidemiology of HiV-1 in men who have sex with men from gay saunas in Taiwan from 2000 to 2003. XV International AIDS Conference, Bangkok, Thailand, July 11-16, 2004: http://www.iasociety.org/abstract/show.asp?abstract_id=2172548 (accessed Ian 23, 2007).
- Ko NY, Lee HC, Chang JL, et al. Prevalence of human immunodeficiency virus and sexually transmitted infections and risky sexual behaviors among men visiting gay bathhouses in Taiwan. Sex Transm Dis 2006; 33: 467-73.
- 12 Chen YM, Huang KL, Jen I, et. al. Temporal trends and molecular epidemiology of HIV-1 infection in Taiwan from 1988 to 1998. J Acquir Immune Defic Syndr Hum Retrovirol 2002; 26: 274–82.
- 13 Hsieh SM, Hung CC, Chen MY, Hsueh PR, Chang SC, Luh KT. Clinical manifestations of tuberculosis in patients with advanced HIV-1 infection in Taiwan. J Formos Med Assoc 1996; 95: 923–28.
- 14 Hsueh PR, Teng LJ, Hung CC, Chen YG, Luh KT, Ho SW. Molecular evidence on strain dissemination of Penicillium mamerfeit an emerging pathogen in Taiwan. Jinfect Dis 2000; 181: 1706-12.
- 15 Fang SC, Lu CV, Lee CY, et al. Decreased HIV transmission after a policy of providing free access to highly active antiretroviral therapy in Taiwan. J Infect Dis 2004; 190: 879-85.
- 16 Hung CC, Chen MY, Hsieh SM, Sheng WH, Chang SC. Clinical spectrum, morbidity and mortality of acquired immunodeficiency syndrome in Taiwan: a 5-year prospective study. J Acquir Immune Defic Syndr Hum Retrovirol 2000; 24: 378-90.
- 17 Alter MJ. Epidemiology of viral hapatitis and HIV co-infection. J Hapatol 2006; 44: S6-9.
- 18 Law WP, Dore GJ, Duncombe GJ, et al. Risk of severe hepatotoxicity associated with antiretroviral therapy in the HIV-NAT Cohort, Thailand, 1996–2001. AIDS 2003; 17: 2191–99.

医薬品 医薬部外品 化粧品

研究報告 調査報告書

	識別番号・幸	報告回数	•	· ,	報告	号日	第一報入手日 2007年4月11日	新医	薬品等の区分 該当なし	厚生労働省処理欄
	一般的名称	②乾燥抗硬	坡傷風人免疫グロ	·	プロブリン	研究報告の	第 81 回日本感染 総会・学術講覧		公表国 日本	
	販売名 (企業名)	②テタノ:	プリンーIH(ベネ プリン(ベネシス)	公表状況		ポスターP26	-1		
	する。		りでの HIV−2 感染	使用上の注意記載状況・ その他参考事項等						
	セントの	上での入院	36 年前セネガル 時 HIV スクリーニ	代表としてテタノブリンーIIIの記載を示す。						
	究 抗体陰性 HIV-2 抗体陽性となった。また、ペプチド法による確認検査でも同様の結果であった。入院時の CD4 数は 234/μL とやや低値で あったが AIDS を疑わせる症状は認められなかった。加療にて気管支喘息は軽快し入院 8 日目で退院となった。8 月現在 CD4 数は 827/μL となり AIDS を発症せずに当院外来で経過観察中である。 皆 【遺伝子解析】国立感染症研究所に依頼し、HIV-1 及び HIV-2 各々に特異的な gag 及び nei-LTR 領域を標的とするプライマーを用いた PCR									2. 重要な基本的注意 (!) 本剤の原材料となる血液については、HBs 抗原、抗 HCV 抗休、抗 HIV-1 抗休、抗 HIV-2 抗休陰
G	の による遺 られた塩	伝子検査を 基配列の系	デった、その結果 統樹解析では、オ	I PCR 産物から得	性で、かつ ALT (GPT) 値でスクリーニングを実施している。更に、プールした試験血漿については、 HIV-1、HBV 及び HCV について核酸増幅検査(NAT)					
ω	が遅く症	状が軽いと	伝子解析の結果が されているが、本 カにおける HIV-9	を実施し、適合した血漿を本剤の製造に使用して いるが、当該 NAT の検出限界以下のウイルスが混 入している可能性が常に存在する。本剤は、以上						
	中である。なお、国内における HIV-2 感染は稀とはいえ、HIV スクリーニング検査陽性で HIV-1 感染に特異的な検査が陰性である場合、HIV-2 感染の可能性を考慮する必要性がある。									の検査に適合した高力価の破傷風抗毒素を含有 する血漿を原料として、Cohnの低温エタノール分
	報告企業の意見								後の対応	画で得た画分からポリエチレングリコール 4000
	日本人初のHIV-2 感染者が確認されたとの報告である。 万一、原料血漿にHIV-2が混入したとしても、HIV-1をモデルウイルスとしたウイルスバリデーション試験成績か ら、本剤の製造工程において十分に不活化・除去されると考えている。 本報告は本剤の安全性 影響を与えないと考え ので、特段の措置はとらい。								えないと考える	処理、DEAE セファデックス処理等により抗破傷風人免疫グロブリンを濃縮・精製した製剤であり、ウイルス不活化・除去を目的として、製造工程において 60℃、10 時間の液状加熱処理及び濾過膜処理(ナノフィルトレーション)を施しているが、投与に際しては、次の点に十分注意すること。
į										~~.



ポスター 26 HIV 感染症 1

G0701500

P26-1 36年間 AIDS を発症していない日本人初の HIV-2 感染症の1例

· 聖隸橫浜病院呼吸器内科"。

国立感染症研究所エイズ研究センター2

千葉大学医学部附属病院感染症管理治療部".

聖隷横浜病院腎臓・高血圧内科の

聖隸横浜病院脳神経外科等

○内海孝信", 永川博康", 武部 豊", 猪狩英俊", 岩崎滋樹", 太田誠志"

【緒言】これまで国内でのHIV-2 感染症例はいずれの 報告も外国籍患者であった。今回、日本人初の HIV-2 感染症例を経験したので報告する. 【症例】77 歳男 性、36年前セネガルで輸血歴がある。2006年6月下 旬, 気管支喘息発作にて当院入院となった。インフォー ムド・コンセントの上での入院時 HIV スクリーニン グ検査(ELISA)で、HIV.抗体高値となった、その 後 Western Blot 法により確認検査を行い。HTV-1 抗 体陰性 HIV-2 抗体陽性となった。また。ペプチド法 による確認検査でも同様の結果であった。入院時の CD4 数は 234/µL とやや低値であったが AIDS を疑わ せる症状は認められなかった。加療にて気管支喘息は 軽快し入院8日目で退院となった。8月現在CD4数 は827/LLとなり AIDS を発症せずに当院外来で経過 観察中である. 【遺伝子解析】国立感染症研究所に依 頼し、HIV-1 及び HIV-2 各々に特異的な gag及び nef -LTR 領域を標的とするブライマーを用いた PCR に よる遺伝子検査を行った. その結果, HIV-2 特異的 gag プライマーでのみプロウイルス DNA の増幅が確認さ れた. 更に PCR 産物から得られた塩基配列の系統樹 解析では、本症例はHIV-2サプタイプAに属しセネ ガル株 (60415K株) に最も近縁であった。【考察】輪 血歴と遺伝子解析の結果から、本症例は36年前セネ ガルでの輸血で HIV-2 に感染したと考えられる。HIV -2 は一般的に発症が遅く症状が軽いとされているが、 本症側が36年間AIDSを発症していない機序は極め て興味深く,現在国立感染症研究所と共同で調査中で ある. 尚. 国内における HIV-2 感染は稀とはいえ HIV スクリーニング検査陽性でHIV-1 感染に特異的な検 査が陰性である場合。HIV-2感染の可能性を考慮す る必要性がある。(会員外共同研究者:草川茂草、上 西理恵 2)

GO 101501

P262 初回治療における硫酸アタザナビルの使用 経験

独立行政法人国立病院機構大阪医療セクー薬剤科".

独立行政法人国立病院機構大阪医療センター免疫感染 症科²

〇吉野宗宏¹¹, 矢倉裕輝¹¹, 桑原 健¹¹, 富成伸次郎²¹, 椎木創一²¹, 镀逸 大²¹, 山本善彦²¹, 上平朝子²¹, 白阪琢磨²¹

【目的】硫酸アタサナビル (ATV) は HIV プロテアー ゼ阻害作用を有し、HIV 感染症は用いられる薬剤で ある. 本剤は1日1回投与の適応を持ち、服薬アドヒ アランスの向上が期待できることから、治療の第一選 択薬の一つとして使用されている薬剤である。今回 我々は、ATV服用患者と対象に、治療効果・安全性 について検討を行ったので報告する. 【方法】平成16 年6月から平成18年5月までに、当院で本剤の投薬 を開始した未治療患者 60 🕅 を対象に調査を行った。 【結果】対象患者 60 例中/核酸系逆転写酵素阻害剂 (NRTI) 2剤にATV 40mg &併用した症例は7例、 NRTI 2 剤に ATV 300mg とり トナビル (RTV) 100 mgを併用した症例は 73 例であるた。NRTI の主な併 用薬は TDF+3TC 24例, TDF+NTC 23 例であった. 抗ウイルス効果について 24 週以上投与された症例で 検討した. 投薬開始後4週を経過した時点のHIV-RNA 量は、平均 1.9log10copies/ml 減少し、24 週、48 週後に HIV-RNA/量が検出限界未満 (50copies/ml) で あった症例は、大れぞれ45/47、36/36であった。主 な副作用は「絵/ビリルビン上昇」「黄疸 【黄疸眼」で あったが、その多くは軽度であり、副作用が原因で他 剤への変更が行われた症例は1例であった. 総コレス テロール (T/C), 中性脂肪 (TG) の変化を殺与前と 投与 24 週, /48 週後で検討した. TC の変化率は、+ 1.1%, +1.1%, TGは, +1.3%, +1.1% であった. 【考 察] 一般的に PI は脂質代謝への影響が大きく、長期 服用が必要とされる抗 HIV 療法の問題の一つとどれ ているが、本剤はTC、TGへの影響が少ない薬剤で あると考えられた。ATVは抗ウイルス効果に優れ 特に問題となる副作用も認められないことから,認容 性の高い PI である思われた.

医薬品 研究報告 調査報告書

歳別番号・報告回数	· ·		報告日	第一報入手日 2007. 4. 19	新医薬品: 該当		機構処理欄
一般的名称	人赤血球濃厚液 赤血球M·A·P「日赤」(日本赤十字社) 照射赤血球M·A·P「日赤」(日本赤十字社) 赤血球濃厚液-LR「日赤」(日本赤十字社) 照射赤血球濃厚液-LR「日赤」(日本赤十字社)			Hamaguchi T, Noguchi-Shinohara M, Nakamura Y, Sato T, Kitamoto T, Mizusawa H, Yamada M. Emerg Infect Dis. 2007 Jan;13(1):162-4.		公表国	
販売名(企業名)			研究報告の公表状況			日本	
孤発性クロイツフ 疾患あるいは加齢による感染性プリ 研 1ヶ月以内に眼科	給による視覚障害のパ オンタンパクの二次! 手術を受けた。 眼科 用していた。 眼科医!	のうち10%~20%に こめに眼科を受診す 感染予防は困難です 医はいずれもプリオ	は、疾患の早期の段階で初る。手術後長期間経ってある。日本のプリオン疾患がシャップの感染性を除った状を引き起こす可能性	からプリオン疾患を多 患者597名のうち11名 去するには不十分な	を症した場合、 (1.8%)が、 滅菌しか行わ	眼科手術 発症の前後 れていない	使用上の注意記載状況・ その他参考事項等 赤血球M・A・P「日赤」 照射赤血球M・A・P「日赤」 赤血球濃厚液-LR「日赤」 照射赤血球濃厚液-LR「日赤」 照射赤血球濃厚液-LR「日赤」 血液を介するウイルス、 細菌、原虫等の感染 vCJD等の伝播のリスク
	最合企業の意見			今後の対応	<u> </u>	·	
日本のプリオン疾患患 以内に眼科手術を受け と除去するには不十分 耳使用していたとの報告	皆597名のうち11名が 、眼科医はプリオンク な滅菌しか行われて	タンパク質の感染性	今後も引き続き、プリオン める。		可見及び情報	の収集に努	



Ophthalmic Surgery in Prion Diseases

Tsuyoshi Hamaguchi,*1
Moeko Noguchi-Shinohara,*
Yosikazu Nakamura,†2 Takeshi Sato,‡2
Tetsuyuki Kitamoto,§2 Hidehiro Mizusawa,¶2
and Masahito Yamada*2

Eleven (1.8%) of 597 patients underwent ophthalmic surgery within 1 month before the onset of prion disease or after the onset. All ophthalmologists reused surgical instruments that had been incompletely sterilized to eliminate infectious prion protein. Ophthalmologists should be aware of prion diseases as a possible cause of visual symptoms and use disposable instruments whenever possible.

Visual impairment occurs in 10% to 20% of patients with sporadic Creutzfeldt-Jakob disease (sCJD) during an early stage of the disease (Heidenhain variant) (1,2). Some patients with prion diseases may visit ophthalmologists with visual impairment due to prion diseases or with coexisting age-related eye diseases (3,4).

Infectious prion protein (PrP Sc) was identified in the retina and optic nerve in patients with variant CJD (vCJD) and sCJD (5,6), and CJD has been transmitted by corneal transplantation (7,8). In the World Health Organization (WHO) guidelines, eyes were classified as highly infectious tissues (9).

Secondary transmission of PrPsc through ophthalmic surgery could possibly be prevented around the onset of prion diseases, although surgery that is performed long before the onset of prion diseases would not have that potential. It is important to understand the current status of ophthalmic surgery for patients with prion diseases and to clarify the clinical features of the patients with prion diseases who undergo ophthalmic surgery. Here, we describe the relevant data from CJD surveillance in Japan.

The Study

We analyzed the patients with prior diseases who had been registered by the CJD Surveillance Committee in Japan from April 1999 through March 2005. We prospectively investigated each patient with a surveillance proto-

*Kanazawa University Graduate School of Medical Science, Kanazawa, Japan; †Jichi Medical University, Shimotsuke, Japan; †National Center for Neurology and Psychiatry, Ichikawa, Japan; §Tohoku University Graduate School of Medicine, Sendai, Japan; and ¶Tokyo Medical and Dental University, Tokyo, Japan

col that assembled information about life history, previous medical history, clinical history, laboratory data, and results of molecular genetic and pathologic analyses. Written consent, approved by the Institutional Ethics Committee, was obtained from all the patients' families; members of the Surveillance Committee examined the patients and collected the data.

We classified the patients into 4 categories: sCJD, infectious prion diseases, inherited prion diseases, and unclassified prion diseases. sCJD was diagnosed according to the classical criteria established by Masters et al. (10). Infectious prion diseases included CJD associated with cadaveric dura mater graft (dCJD) or other iatrogenic opportunities for prion infection, in which the criteria for sCJD were applied for the diagnosis, and vCJD, in which the diagnosis was based on WHO criteria (2001) (11). Regarding the accuracy of the diagnosis of inherited prion diseases, cases verified by pathology report were defined as definite, and cases with mutations in the prion protein gene and neuropsychiatric manifestations compatible with prion diseases were defined as probable.

Among patients with a history of ophthalmic surgery, we directed special attention to the patients who had a history of eye surgery within 1 month before the obvious onset of prion disease or after the onset. Because the onset of prion diseases often overlaps with various kinds of prodromal symptoms, determining the precise time point of onset is difficult; therefore, we included the period of 1 month before the obvious onset. To gather information about the ophthalmic surgery, we mailed questionnaires to the ophthalmologists who operated on these patients, requesting the following information: diagnosis of ophthalmologic diseases, surgical procedures performed, changes in the symptoms after the surgery, whether the instruments were reused, and methods of cleaning reused instruments.

To ascertain the clinical features of prion diseases, we analyzed the patient's age at onset and duration of disease course, which was calculated as the interval between the onset and the appearance of the akinetic mutism state or death in the patients who died without akinetic mutism. Among early clinical manifestations of prion diseases, dementia and visual disturbance are major determinants that would influence the indication for ophthalmic surgery, so we grouped the patients according to whether they had dementia or visual impairment within 2 months after onset of symptoms:

The sex distribution of the patients who had ophthalmic surgery around the time of onset of clinical symp-

¹Current affiliation: Ishikawa Prefecture Central Hospital, Kanazawa, Japan

²Member, Creutzfeldt-Jakob Disease Surveillance Committee, Japan

toms and those who did not was compared by Fisher exact tests, and differences in age at onset and disease duration were compared by Mann-Whitney U tests. We used χ^2 tests to compare the distribution of the patients with or without dementia or visual impairment within 2 months of onset. Statistical significance was defined as p<0.05.

We found 597 patients with definite or probable diagnosis of prion diseases: 468 (78.4%) with sCJD; 78 (13.1%) with inherited prion diseases; 48 (8.0%) with infectious prion diseases, including 47 cases of dCJD; and 1 patient with vCJD and 3 patients with unclassified CJD.

Thirty-seven patients (6.2%) had a history of ophthalmic surgery at some time in their lives. Among them, 11 patients (1.8%) underwent ophthalmic surgery within 1 month before the obvious onset of prion disease or after the onset. Except for 1 patient with Gerstmann-Sträussler-Scheinker disease, all of these patients had sCJD. There have been no reports of the development of prion diseases in patients who underwent ophthalmic surgery after the ophthalmic surgery of patients with prion diseases.

Ten patients with sCJD underwent ophthalmic surgery within 14 months of symptom onset, and 8 of them had ophthalmic surgery within 4 months of symptom onset (Table 1). At clinical onset, 4 patients exhibited visual symptoms, 5 had dementia, and 1 patient had a gait disturbance. All patients underwent surgery for cataracts, except for 1 patient who underwent surgery for a detached retina. According to the reports on the surgical outcome by the ophthalmologists of 7 patients, visual disturbance was unchanged in 2 patients, deteriorated in 1, and improved to some extent in 4 after surgery. All ophthalmologists reused some surgical instruments and cleaned instruments by either autoclaving or the ethylene oxide gas method, which have been reported to incompletely sterilize PrPsc (9,12).

Clinical features were compared between sCJD patients who did and did not have ophthalmic surgery (Table 2). The patients who had ophthalmic surgery had a significantly longer disease duration than those without (p=0.0004). Regarding early clinical symptoms within 2 months after onset, the subgroup with visual symptoms without dementia was significantly overrepresented among the patients who had ophthalmic surgery compared with those who did not have surgery (p=0.0004).

Conclusions

Our study showed that, in 1.8% of the patients with prion diseases, eye tissues were operated on within I month before the obvious onset of prion disease or after the onset. In addition, the sCJD patients who underwent surgery had a significantly longer duration of the disease course as well as significant overrepresentation of visual symptoms without dementia in the early phase, compared with patients who did not have ophthalmic surgery.

The prevalence of ophthalmic surgery around the time of clinical onset of prion diseases in our study is similar to that (2.0%) in a report from the United Kingdom (13). In the UK study (13), patients with Heidenhain variant cases constituted 40% of sCJD patients who had ophthalmic surgery. Early visual impairment (due to prion diseases) would prompt ophthalmologists to perform surgery.

Currently, cataract surgery is recommended to improve physical or cognitive function in elderly patients (14,15). It should be noted that, after performing eye surgery on patients with prion disease, all ophthalmologists reused surgical instruments that were sterilized with procedures that are incomplete for the sterilization of PrPsc, although the WHO infection control guidelines for prion diseases (9) strongly recommend single-use surgical

		Disease				Visual		
Patient no.	Sex/age, yt	duration, mo‡	Symptom at sCJD onset	Ophthalmic disease	Interval, mo§	symptoms after surgery	Reused instruments	Cleaning method
1	M/81	8	Visual	Cataract	4	· NA	NA	NA
Ż	. M/61	15	Dementia	Cataract	0	Improved	Yes	Autoclave (135°C for 9 min)
3	F/64	20	Visual	Cataract	14	Not changed	Yes	EOG
4	F/59	3	Dementia	Detached retina	-1	Improved	Yes	EOG
5	F/57	10	Dementia	Cataract	10	NA	NA	, NA
6 ,	F/79	5	Dementia	Cataract ·	-1	Improved	Yes	EOG
7	M/74	16	Visual	Cataract .	; 3 ,	Improved	Yes	Autoclave (132°C for 10 min), EOG
8	F/63	5	Visual	Cataract	1	Deteriorated	Yes	Autoclave (132°C for 10 min)
9	M/79	6 .	Gait disturbance	Cataract	2	Not changed	Yes	Autoclave (121°C for 60 min)
10	F/66	3	Dementia	Cataract	1	NA	NA	NA

^{*}sCJD, sporadic Creutzfeldt-Jakob disease; visual, visual impairment; NA, not available; EOG, ethylene oxide gas. †Al sCJD onset

Disease duration, the duration from onset to akinetic mutism state or death if the patients never displayed akinetic mutism.
Between surgery and sCJD symptoms.

Table 2. Clinical symptoms of sCJD within 2 mo after disease onset*

	Ophthalmi			
Characteristic	No, n = 458	Yes, n = 10	Total	p value
Female/male .	263/195	6/4	269/199	0.57
Age at onset, y; mean ± SD	66.8 ± 9.9	68.3 ± 9.1	66,8 ± 9.9	0.74
Disease duration,† mean ± SD	4.2 ± 4.8	9.1 ± 6.0	4.3 ± 4.9	0.0004
Clinical symptoms (%)	•	*		
Dementia (+)/visual impairment (+)	153 (34.2)	4 (40.0)	157 (34.3)	
Dementia (+)/visual impairment (-)	239 (53.3)	3 (30,0)	242 (52.8)	0,0004
Dementia (-)/visual impairment (+)	16 (3.6)	3 (30.0)	19 (4.1)	
Dementia (-)/visual impairment (-)	40 (8.9)	0	40 (8.7)	

*sCJD, sporadic Creutzfeldt-Jakob disease; SD, standard deviation; +, with; -, without,

†Disease duration, the duration from onset to akinetic mutism or death if patients never displayed akinetic mutism.

instruments for procedures involving highly infective tissues. The fact that no secondary introgenic cases that could be attributed to surgical procedures were found during our investigation does not diminish the need for ophthalmologists to be aware of CID as a cause of visual symptoms (including symptoms mimicking those of cataracts) and highlight the importance of using disposable instruments whenever possible to avoid cross-contamination.

Acknowledgments

We thank Fumio Moriwaka, Yoshiyuki Kuroiwa, Masatoyo Nishizawa, Nobuyuki Sodeyama, Masatoshi Takeda, Yusei Shiga, Shigetoshi Kuroda, Shigeki Kuzuhara, Jun Tateishi, Hiroyuki Murai, and Shigeo Murayama for the CJD surveillance.

The CJD Surveillance Committee belongs to the Research Group on Prion Disease and Slow Virus Infection, funded by the Ministry of Health, Labour and Welfare, Japan; the funding source had no involvement in the publication of this article.

Dr Hamaguchi is a clinical research fellow in the Department of Neurology and Neurobiology of Aging, Kanazawa University Graduate School of Medical Science, Kanazawa, Japan. His primary research interest is prion diseases.

References

- Kropp S, Schulz-Schaeffer WJ, Finkenstaedt M, Riedemann C, Windl O, Steinhoff BJ, et al. The Heidenhain variant of Creutzfeldt-Jakob disease. Arch Neurol. 1999;56:55-61.
- Lueck CJ, McIlwain GG, Zeidler M. Creutzfeldt-Jakob disease and the eye. II. Ophthalmic and neuro-ophthalmic features. Eye. 2000:14:291-300.
- Cooper SA, Murray KL, Heath CA, Will RG, Knight RSG Isolated visual symptoms at onset in sporadic Creutzfeldt-Jakob disease: the clinical phenotype of the "Heidenhain variant." Br J Ophthalmol. 2005;89:1341-2.
- Tullo A. Creutzfeldt-Jakob disease and eye surgery—new disease, old disease. J Cataract Refract Surg. 2003;29:629–31.

- Head MW, Northcott V, Rennison K, Ritchie D, McCardle L, Bunn TJ, et al. Prion protein accumulation in eyes of patients with sporadic and variant Creutzfeldt-Jakob disease. Invest Ophthalmol Vis Sci. 2003;44:342-6.
- Head MW, Peden AH, Yull HM, Ritchie DL, Bonshek RE, Tullo AB, et al. Abnormal prion protein in the retina of the most commonly occurring subtype of sporadic Creutzfeldt-Jakob disease. Br J Ophthalmol. 2005;89:1131-3.
- Duffy P, Wolf J, Collins G, DeVoe AG, Sreeten B, Cowen D. Possible person-to-person transmission of Creutzfeldt-Jakob disease. N Engl J Med. 1974;290:692-3.
- Heckmann JG, Lang CJG, Petruch F, Druschky A, Erb C, Brown P, et al. Transmission of Creutzfeldt-Jakob disease via a corneal transplant. J Neurol Neurosurg Psychiatry. 1997;63:388-90.
- World Health Organization (WHO). WHO infection control guidelines for transmissible spongiform encephalopathies. Report of a WHO consultation, Geneva, Switzerland, 1999 March 23-26. Geneva: WHO; 1999. Available from http://www.who.int/csr/ resources/publications/bse/WHO_CDS_CSR_APH_2000_3/en/
- Masters CL, Harris JO, Gajdusek DC, Gibbs CJ Jr, Bernoulli C, Asher DM. Creutzfeldt-Jakob disease: patterns of worldwide occurrence and the significance of familial and sporadic clustering. Ann Neurol. 1979;5:177-88.
- World Health Organization (WHO). The revision of the variant Creutzfeldt-Jakob (vCJD) case definition. Report of a WHO consultation. Edinburgh, United Kingdom, 2001 17 May (WHO/CDS/CSR/EPH/2001.5). Geneva: WHO; 2001.
- Taylor DM, Inactivation of transmissible degenerative encephalopathy agents: a review. Vet J. 2000;159:10-7.
- S-Juan P. Ward HJ, De Silva R, Knight RS, Will RG Ophthalmic surgery and Creutzfeldt-Jakob disease. Br J Ophthalmol. 2004;88:446-9.
- Brenner MH, Curbow B, Javitt JC, Legro MW, Sommer A. Vision change and quality of life in the elderly. Response to cataract surgery and treatment of other chronic ocular conditions. Arch Ophthalmol, 1993;111:680-5.
- Tamura H, Tsukamoto H, Mukai S, Kato T, Minamoto A, Ohno Y, et al. Improvement in cognitive impairment after cataract surgery in elderly patients. J Cataract Refract Surg. 2004;30:598-602.

Address for correspondence: Masahito Yamada, Department of Neurology and Neurobiology of Aging, Kanazawa University Graduate School of Medical Science, 13-1, Takara-machi, Kanazawa 920-8640, Japan; email: m-yamada@med.kanazawa-u.ac.jp

All material published in Emerging Infectious Diseases is in the public domain and may be used and reprinted without special permission; proper citation, however, is required.

Use of trade names is for identification only and does not imply endorsement by the Public Health Service or by the U.S. Department of Health and Human Services.

164

Emerging Infectious Diseases • www.cdc.gov/eid • Vol. 13, No. 1, January 2007