

World Journal of *Ophthalmology*

World J Ophthalmol 2012 November 16; 2(1): 1-5





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**BRIEF ARTICLE**

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Kobayashi D, Takahashi O, Glasziou PP, Fukui T

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Volume 2 Number 1 November 16, 2012

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The aim of *WJO* is to report rapidly new theories, methods and techniques for prevention, diagnosis, treatment, rehabilitation and nursing in the field of ophthalmology. *WJO* covers diagnostic imaging, optometry, ocular fundus diseases, cataract, glaucoma, keratopathy, ocular trauma, strabismus, and pediatric ocular diseases, blindness prevention, traditional medicine, integrated Chinese and Western medicine, evidence-based medicine, epidemiology and nursing. The journal also publishes original articles and reviews that report the results of applied and basic research in fields related to ophthalmology, such as immunology, physiopathology, cell biology, pharmacology, medical genetics, and pharmacology of Chinese herbs.

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NAME OF JOURNAL
World Journal of Ophthalmology

ISSN
ISSN 2218-6239 (online)

LAUNCH DATE
December 30, 2011

FREQUENCY
Bimonthly

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Room 1701, 17/F, Henan Building,
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Fax: +852-31158812
Telephone: +852-58042046
E-mail: bpgooffice@wjnet.com
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PUBLICATION DATE
November 16, 2012

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Optimal screening interval for intraocular pressure measurement for Asian glaucoma patients

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Received: January 16, 2012 Revised: September 17, 2012

Accepted: October 10, 2012

Published online: November 16, 2012

Abstract

AIM: To explore the optimal interval of intraocular pressure (IOP) measurement for screening glaucoma in healthy people.

METHODS: From January to December 2005, we consecutively enrolled all participants (> 20 years old) attending the Center for Preventive Medicine at St. Luke's International Hospital in Tokyo, Japan, for the annual health check program. The program promoted the early detection of chronic diseases and their risk factors. We excluded people who had glaucoma or a high IOP (≥ 22 mmHg) at baseline. The annual health check-ups collected all demographic information and medical history with an initial evaluation, including IOP measurement. IOP was measured in both eyes with a full auto-tonometer TX-F (Canon, Tokyo, Japan). Participants with an IOP ≥ 22 mmHg in either eye were considered to require additional evaluation for glaucoma. We divided the participants into two groups based on age: under 65 years old and over 65 years old. The United States Department of Health and Human Services Cen-

ters for Medicare and Medicaid Services guideline was used as a reference.

RESULTS: From January 2005 to July 2008, 12 385 participants underwent check-ups each year. The mean \pm SD IOP in the higher eye at baseline was 13.4 (2.6) in 2005, 13.2 (2.7) in 2006, 13.3 (2.6), and 12.8 (2.6) in 2008. In addition, we analyzed the differences with an analysis of variance (ANOVA), and additional analysis was performed with Bonferroni's correction. The difference between the 4 years was significant ($P < 0.01$) with ANOVA. Bonferroni analysis revealed significant differences between 2005 and 2006 ($P < 0.01$), 2005 and 2008 ($P < 0.01$), 2006 and 2007 ($P < 0.01$), 2006 and 2008 ($P < 0.01$), and 2007 and 2008 ($P < 0.01$). Only the difference between 2005 and 2007 was not significant ($P = 0.1$). Logistic regression suggested that only age ($P < 0.01$) and baseline IOP ($P < 0.01$) were associated with high IOP; the presence of diabetes, HgbA1c level, gender, systolic blood pressure, diastolic blood pressure, low-density lipoprotein and family history were non-significant.

CONCLUSION: Annual IOP check-ups may be recommended for participants aged ≥ 65 years with baseline IOPs of 17-21 mmHg. A check-up every 3 years or more may be recommended for patients with IOPs < 17 mmHg.

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Key words: Glaucoma; Screening interval; Japan; Intraocular pressure; Asian

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Kobayashi D, Takahashi O, Glasziou PP, Fukui T. Optimal screening interval for intraocular pressure measurement for Asian glaucoma patients. *World J Ophthalmol* 2012; 2(1): 1-5 Available from: URL: <http://www.wjgnet.com/2218-6239/full/v2/i1/1.htm> DOI: <http://dx.doi.org/10.5318/wjo.v2.i1.1>

INTRODUCTION

Glaucoma is a significant public health problem: approximately 45 million patients globally have open-angle glaucoma, and approximately 8.4 million patients become blind because of glaucoma^[1-3]. Although it has been reported that the prevalence of glaucoma for Asians is less than for Caucasian and African races, approximately 4 million patients suffer from glaucoma in Japan^[4-6]. Therefore, it is necessary to treat glaucoma and prevent blindness^[7-10].

Intraocular pressure (IOP) measurement is one method to evaluate glaucoma^[3,11], and it is sufficient as a screening procedure^[12]. An IOP of 22 mmHg is considered a screening cut-off level^[13,14], and IOP is measured with a contact or non-contact IOP tonometer. A non-contact tonometer is as reliable as Goldmann applanation^[15]. Patients with IOP values greater than 22 mmHg require additional work-up for glaucoma treatments.

There are various recommendations for glaucoma screening^[16-18]. The United States Preventive Services Task Force found insufficient evidence to recommend either for or against screening adults for glaucoma^[19]. Routine population-based mass screening for glaucoma may not be cost-effective^[20]. However, some guidelines suggest that screening high-risk participants, for example, the elderly, African-Americans, those who have a family history or those who have high IOP at baseline, may be effective^[21] and cost-effective^[20]. Therefore, we aimed to evaluate the optimal screening interval for glaucoma, especially among high-risk groups in Japan.

MATERIALS AND METHODS

Study participants

From January to December 2005, we consecutively enrolled all participants (> 20 years old) attending the Center for Preventive Medicine at St. Luke's International Hospital in Tokyo, Japan, for the annual health check program. The program promoted the early detection of chronic diseases and their risk factors. The data collected contained current and past medical history, including diabetes^[22], and family history, including glaucoma^[23-26]. We excluded people who had glaucoma or a high IOP (≥ 22 mmHg) at baseline.

Measurements

The annual health check-ups collected all demographic

information and medical history with an initial evaluation, including IOP measurement. IOP was measured in both eyes with a full auto-tonometer TX-F (Canon, Tokyo, Japan). Participants with an intraocular pressure ≥ 22 mmHg in either eye were considered to require additional evaluation for glaucoma. The diagnosis of glaucoma was reported by individual participants. We divided participants into two groups based on age: under 65 years old and over 65 years old. The United States Department of Health and Human Services Centers for Medicare and Medicaid Services guideline was used as a reference. A total of 12 385 participants were enrolled in our study. The number of participants under 65 years old was 10 600, and the number of participants aged 65 years or older was 1785. Among those who were under 65 years old, 1331 had an IOP of 17-21 mmHg at baseline, 6781 had an IOP of 12-16 mmHg, and 1219 had an IOP of 11 mmHg or lower. Among those who were aged 65 or older, 184 had an IOP of 17-21 mmHg at baseline, 1048 had an IOP of 12-16 mmHg, and 553 had an IOP of 11 mmHg or lower.

Statistical analysis

All data analyses were performed with an exact binominal using SPSS software 15.0J (IBM Japan, Tokyo, Japan) and Stata version 10 (STATA Corp, College Station, TX).

RESULTS

From January 2005 to July 2008, 12 385 participants underwent check-ups every year. The mean \pm SD age of participants was 50 (12) years, and 7617 (53%) were male. A total of 1852 (15%) were current smokers, and 436 (3.7%) had diabetes. A total of 133 (1.1%) had a family history of glaucoma. The mean \pm SD body mass index was 22.5 (3.2) kg/m², the height was 163.7 (8.7) cm, the systolic blood pressure (SBP) was 119.3 (17.7) mmHg, the diastolic blood pressure (DBP) was 74.2 (11.3) mmHg, the fasting plasma glucose was 100.4 (15.6) mg/dL, and the hemoglobin A1c (HbA1c) at baseline was 5.1 (0.6%). The total cholesterol, low-density lipoprotein (LDL) cholesterol, high-density lipoprotein (HDL) cholesterol, and triglyceride levels at baseline were 206.1 (33.5) mg/dL, 117.9 (29.6) mg/dL, 62.2 (15.7) mg/dL, and 102.4 (75.5) mg/dL, respectively (Table 1).

The mean \pm SD IOP in the higher eye at baseline was 13.4 (2.6) in 2005, 13.2 (2.7) in 2006, 13.3 (2.6) in 2007, and 12.8 (2.6) in 2008. In addition, we analyzed the differences with an analysis of variance (ANOVA), and additional analysis was performed with Bonferroni's correction. The difference between the 4 years was significant ($P < 0.01$) with the ANOVA. Bonferroni analysis revealed significant differences between 2005 and 2006 ($P < 0.01$), 2005 and 2008 ($P < 0.01$), 2006 and 2007 ($P < 0.01$), 2006 and 2008 ($P < 0.01$), and 2007 and 2008 ($P < 0.01$). Only the difference between 2005 and 2007 was not significant ($P = 0.1$) (Figure 1).

After three years, for participants with an IOP of 11

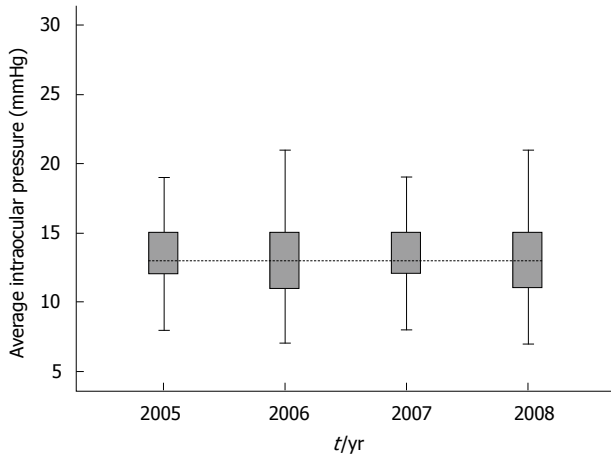


Figure 1 Average of intraocular pressure in each year.

Table 1 Participants' characteristics at baseline $n = 12\,385$
 n (%)

Characteristics	mean \pm SD
Age (yr)	50 \pm 12
Male	7617 (53)
Current smoker	1852 (15.0)
Height (cm)	163.7 \pm 8.7
Body mass index (kg/m ²)	22.5 \pm 3.2
Diabetes	463 (3.7)
Family history of glaucoma	133 (1.1)
Higher intraocular pressure in each eye (mmHg)	13.3 \pm 2.6
Systolic blood pressure (mmHg)	119.3 \pm 17.7
Diastolic blood pressure (mmHg)	74.2 \pm 11.3
Glucose (mg/dL)	100.4 \pm 15.6
Hemoglobin A1c (%)	5.1 \pm 0.6
Total-cholesterol (mg/dL)	206.1 \pm 33.5
Low-density lipoprotein-cholesterol (mg/dL)	117.9 \pm 29.6
High-density lipoprotein-cholesterol (mg/dL)	62.2 \pm 15.7
Triglyceride (mg/dL)	102.4 \pm 75.5

or less than 11 mmHg, 12-16 mmHg and 17-21 mmHg at baseline, the cumulative incidence [95% confidence interval (CI) was 1.1% (0.4%-2.4%), 2.8% (1.9%-4.1%) and 9.5% (5.5%-15.0%)], respectively, in the group over 65 years old and 0.5% (0.3%-0.9%), 0.8% (0.6%-1.0%) and 2.6% (1.8%-3.6%), respectively, in the group under 65 years old (Figure 2). We analyzed these data with a log-rank test. The result revealed that the group over 65 years old with a baseline IOP of 17-21 mmHg had significant differences compared to all of the other groups ($P < 0.01$) (Figure 2).

Logistic regression suggested that only age ($P < 0.01$) and baseline IOP ($P < 0.01$) were associated with high IOP; the presence of diabetes, HgbA1c level, gender, SBP, DBP, LDL and family history were non-significant.

DISCUSSION

Our study shows that the likelihood of IOP increasing to approximately 22 mmHg is strongly predicted by baseline IOP level and age. The screening interval for glaucoma

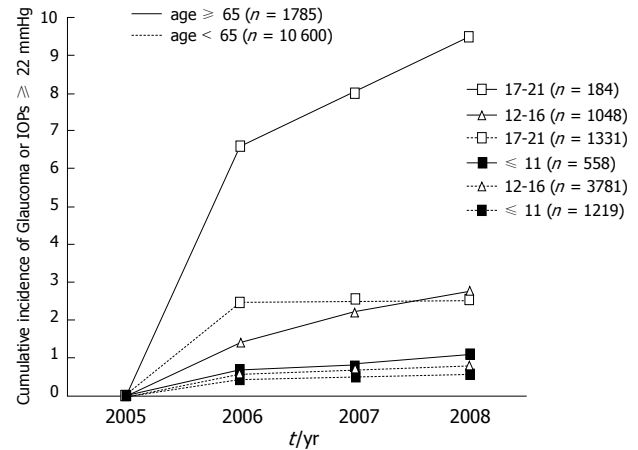


Figure 2 The cumulative incidence of glaucoma or high intraocular pressure over 4 years in each group.

by IOP measurement may be determined from a participant's age and baseline IOP.

For participants over 65 years old, an IOP of 17 mmHg at baseline may be considerable. In the group with IOPs of 17-21 mmHg at baseline, the cumulative incidence of glaucoma or high IOP was approximately 7% in the following year. In contrast, in the groups with IOPs ≤ 11 mmHg and 12-16 mmHg at baseline, the cumulative incidence was below 3% in 3 years. Therefore, annual IOP check-ups may be appropriate for individuals with IOPs of 17-21 mmHg at baseline, and check-ups every 3 years or more may be appropriate for individuals with IOPs below 17 mmHg.

The cumulative incidence of glaucoma or high IOP was low for participants less than 65 years old. In groups whose IOP was ≤ 11 mmHg and 12-16 mmHg at baseline, the cumulative incidence was below 1% in 3 years. In the group that had IOPs of 17-21 mmHg at baseline, the cumulative incidence was below 3% in 3 years. Therefore, check-ups every 3 years or more may be appropriate for people under 65 years old in the Asian population.

The American Optometric Association recommends annual eye examinations for people at risk for glaucoma. Our results demonstrated that elderly patients with high baseline IOP meet the criteria of being at high risk for glaucoma.

In our study, participants whose baseline IOPs were 17-22 mmHg had a high incidence of high IOPs in the first year. In our opinion, this result was due to measurement error of IOPs^[27]. Because IOP measurement using the non-contact method is prone to error, participants who had borderline IOPs at baseline and a wide range of measurement error tended to have high IOPs the following year. Because we analyzed cumulative incidence, the incidence of high IOPs tended to be high in the first year for those with IOPs of 17-22 mmHg.

There are some limitations to our study. First, our data lack possible risk factors for glaucoma, such as cataracts^[28,29], steroid use^[30] and myopia^[31-33]. More frequent examination may be recommended for individuals who

have these risk factors. Second, there are some missing data in our study because not all participants returned every year. Although 34 234 participants came to the health check-up in 2005, only 12 385 (36.2%) continued to come for 4 years. Because our health check-up was not mandatory, some participants did not return. Third, our data did not have the results of other optic nerve measurements to diagnose glaucoma^[34-36]. Because Asians are more likely to have normal tension glaucoma compared to other races, further evaluations are required in a future study. Finally, our data lack evaluation for glaucoma. Although we could identify participants with high IOP using non-contact measurements from our data, we could not identify glaucoma patients. Additional studies that include glaucoma patient information are necessary to decide the optimal screening interval for glaucoma.

In conclusion, for the high-risk group (age ≥ 65 years and baseline IOP 17-21 mmHg), careful IOP check-up might be recommended. For all others, check-ups every 3 years or more appear to be reasonable.

COMMENTS

Background

Glaucoma is one of the most serious causes of blindness. Early detection and treatment are required. However, the screening interval for glaucoma is still controversial. This study aims to evaluate the optimal screening interval for glaucoma with non-contact intraocular pressure measurement.

Research frontiers

The United States Preventive Services Task Force and National Institute for Health and Clinical Excellence address this type of screening guideline.

Innovations and breakthroughs

This study is innovative because of the evaluation of a screening interval for glaucoma, which is still controversial.

Applications

This study may be useful for glaucoma screening with non-contact intraocular pressure measurement. However, additional evaluations are needed to evaluate high-risk populations.

Peer review

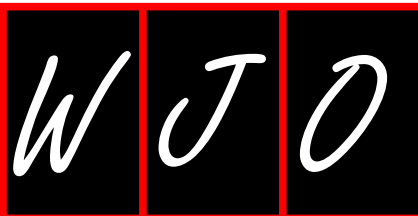
In this study, a large population undergoing general medical screening had their intraocular pressure evaluated at enrolment and annually thereafter. The authors attempt to define the optimal interval between intraocular pressure measurements.

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ACKNOWLEDGMENTS

Acknowledgments to reviewers of *World Journal of Ophthalmology*

Many reviewers have contributed their expertise and time to the peer review, a critical process to ensure the quality of *World Journal of Ophthalmology*. The editors and authors of the articles submitted to the journal are grateful to the following reviewers for evaluating the articles (including those published in this issue and those rejected for this issue) during the last editing time period.

Stephen A Vernon, Professor, Department of Ophthalmology, University Hospital, Nottingham, NG7 2UH, United Kingdom

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Events Calendar 2012

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Athens, Greece

January 13-15, 2012

7th Pan-Hellenic Vitreo-Retinal
Meeting
Athens, Greece

January 14-15, 2012

7th Pan-Hellenic Vitreo-Retinal
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January 20-21, 2012

2nd EURETINA Winter Meeting
Madrid, Spain

January 28, 2012

16th ESCRS Winter Meeting
Rome, Italy

February 3-5, 2012

ASCRS 2012 - Winter update
Prague, Czech Republic

February 16-20, 2012

World Ophthalmology Congress
2012
Abu Dhabi, United Arab Emirates

February 16-20, 2012

World Ophthalmology Congress
2012
Play del Carmen, Mexico

February 16-20, 2012

2nd EUROLAM Macula and Retina
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March 5-7, 2012

2nd International Conference
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Ophthalmology
Omaha Marriott, NE, United States

March 16-17, 2012

3rd COPHY - Controversies in
Ophthalmology
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March 22-25, 2012

27th Asia Pacific Academy of
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Istanbul, Turkey

March 22-25, 2012

The 3rd World Congress on
Controversies in Ophthalmology
Istanbul, Turkey

March 29-April 1, 2012

International Congress of
Ophthalmology and Optometry
Hangzhou, China

April 12, 2012

Conference for Ophthalmic
Educators in Busan
Busan, South Korea

April 13-16, 2012

The 27th Asia Pacific Academy of
Ophthalmology Congress
BEXCO, Busan, South Korea

April 13-16, 2012

ARVO 2012
Busan, South Korea

April 20, 2012

The America Conference on Pediatric
Cerebral Visual Impairment
Children's Hospital and Medical
Center
Omaha, Nebraska

May 5, 2012

ARVO/ISIE Imaging Conference
Fort Lauderdale, FL, United States

May 6-10, 2012

10th International Congress of
Società Oftalmologica Italiana
Fort Lauderdale, FL, United States

May 23-26, 2012

16th Afro-Asian Congress of
Ophthalmology - 5th Mediterranean
Retina Meeting
Milan, Italy

June 13-16, 2012

25th International Congress of
German Ophthalmic Surgeons
Istanbul, Turkey

June 14-17, 2012

10th European Glaucoma Society
Congress
Nuernberg, Germany

June 15-16, 2012

Drug and Gene Delivery to the Back
of the Eye: From Bench to Bedside
Aurora, Colorado

June 17-22, 2012

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Copenhagen, Denmark

June 29-July 1, 2012

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Milan, Italy

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Milan, Italy

September 19-20, 2012

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on Ophthalmology
Berlin, Germany

September 21-22, 2012

92nd National Congress of Società
Oftalmologica Italiana
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Milan, Italy

November 28-December 1st, 2012

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Rome, Italy

December 26-28, 2012

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Ophthalmology and Optometry
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Name of journal

World Journal of Ophthalmology

ISSN

ISSN 2218-6239 (online)

Editor-in-chief

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World Journal of Ophthalmology
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Indexed and abstracted in

Digital Object Identifier

Published by

Baishideng Publishing Group Co., Limited

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Acknowledgments

Brief acknowledgments of persons who have made genuine con-

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- 2 **Lin GZ**, Wang XZ, Wang P, Lin J, Yang FD. Immunologic effect of Jianpi Yishen decoction in treatment of Pixu-diarhoea. *Shijie Huaren Xiaohua Zazhi* 1999; **7**: 285-287

In press

- 3 **Tian D**, Araki H, Stahl E, Bergelson J, Kreitman M. Signature of balancing selection in Arabidopsis. *Proc Natl Acad Sci USA* 2006; In press

Organization as author

- 4 **Diabetes Prevention Program Research Group**. Hypertension, insulin, and proinsulin in participants with impaired glucose tolerance. *Hypertension* 2002; **40**: 679-686 [PMID: 12411462 PMCID:2516377 DOI:10.1161/01.HYP.0000035706.28494.09]

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- 5 **Vallancien G**, Emberton M, Harving N, van Moorselaar RJ;

Alf-One Study Group. Sexual dysfunction in 1, 274 European men suffering from lower urinary tract symptoms. *J Urol* 2003; **169**: 2257-2261 [PMID: 12771764 DOI:10.1097/01.ju.0000067940.76090.73]

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- 6 21st century heart solution may have a sting in the tail. *BMJ* 2002; **325**: 184 [PMID: 12142303 DOI:10.1136/bmj.325.7357.184]

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- 7 **Geraud G**, Spierings EL, Keywood C. Tolerability and safety of frovatriptan with short- and long-term use for treatment of migraine and in comparison with sumatriptan. *Headache* 2002; **42** Suppl 2: S93-99 [PMID: 12028325 DOI:10.1046/j.1526-4610.42.s2.7.x]

Issue with no volume

- 8 **Banit DM**, Kaufer H, Hartford JM. Intraoperative frozen section analysis in revision total joint arthroplasty. *Clin Orthop Relat Res* 2002; **(401)**: 230-238 [PMID: 12151900 DOI:10.1097/00003086-200208000-00026]

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Personal author(s)

- 10 **Sherlock S**, Dooley J. Diseases of the liver and biliary system. 9th ed. Oxford: Blackwell Sci Pub, 1993: 258-296

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Patent (list all authors)

- 16 **Pagedas AC**, inventor; Ancel Surgical R&D Inc., assignee. Flexible endoscopic grasping and cutting device and positioning tool assembly. United States patent US 20020103498. 2002 Aug 1

Statistical data

Write as mean \pm SD or mean \pm SE.

Statistical expression

Express *t* test as *t* (in italics), *F* test as *F* (in italics), chi square test as χ^2 (in Greek), related coefficient as *r* (in italics), degree of freedom as *v* (in Greek), sample number as *n* (in italics), and probability as *P* (in italics).

Units

Use SI units. For example: body mass, *m* (B) = 78 kg; blood pressure, *p* (B) = 16.2/12.3 kPa; incubation time, *t* (incubation) = 96 h,

blood glucose concentration, *c* (glucose) 6.4 ± 2.1 mmol/L; blood CEA mass concentration, *p* (CEA) = 8.6 ± 24.5 μ g/L; CO₂ volume fraction, 50 mL/L CO₂, not 5% CO₂; likewise for 40 g/L formaldehyde, not 10% formalin; and mass fraction, 8 ng/g, *etc.* Arabic numerals such as 23, 243, 641 should be read 23 243 641.

The format for how to accurately write common units and quantum numbers can be found at: http://www.wjgnet.com/2218-6239/g_info_20100724174652.htm.

Abbreviations

Standard abbreviations should be defined in the abstract and on first mention in the text. In general, terms should not be abbreviated unless they are used repeatedly and the abbreviation is helpful to the reader. Permissible abbreviations are listed in Units, Symbols and Abbreviations: A Guide for Biological and Medical Editors and Authors (Ed. Baron DN, 1988) published by The Royal Society of Medicine, London. Certain commonly used abbreviations, such as DNA, RNA, HIV, LD50, PCR, HBV, ECG, WBC, RBC, CT, ESR, CSF, IgG, ELISA, PBS, ATP, EDTA, mAb, can be used directly without further explanation.

Italics

Quantities: *t* time or temperature, *c* concentration, *A* area, *l* length, *m* mass, *V* volume.

Genotypes: *gyrA*, *arg 1*, *c myc*, *c fos*, *etc.*

Restriction enzymes: *EcoRI*, *HindI*, *BamHI*, *Kho I*, *Kpn I*, *etc.*

Biology: *H. pylori*, *E. coli*, *etc.*

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