Transpupillary thermotherapy (TTT) for the treatment of choroidal neovascularisation

R S B Newsom, J C McAlister, M Saeed, J D A McHugh

Abstract

Aim—To assess the effectiveness of transpupillary thermotherapy (TTT) for the treatment of classic and occult choroidal neovascularisation (CNV).

Method—In a retrospective, case selected, open label trial 44 eyes of 42 patients with CNV secondary to age related macular degeneration (ARMD) were studied. 44 eyes with angiographically defined CNV were treated with diode laser (810 nm) TTT. Laser beam sizes ranged between 0.8 and 3.0 mm and power settings between 250–750 mW. Treatment was given in one area for 1 minute, the end point being no visible change, or a slight greying of the retina. Outcome was assessed with Snellen visual acuity and clinical examination; in 24/44 patients angiographic follow up was available.

Results—12 predominantly classic CNV and 32 predominantly occult membranes were followed up for a mean of 6.1 months (range 2–19). Mean change in vision for classic membranes was −0.75 (SD 1.75) Snellen lines and occult membranes was −0.66 Snellen lines (2.1) (p>0.05). Predominantly classic membranes were closed in 75% (95% CI: 62.5–87.5) of eyes, remained persistent in 25% (95% CI: 12.5–37.5); no recurrences occurred. Predominantly occult membranes were closed in 78% (95% CI: 70.1–85.3) of eyes, remained persistent in 12.5% (95% CI: 6.6–18.5), and were recurrent in 5.1% (95% CI: 4.2–14.3).

Conclusions—Transpupillary thermotherapy is a potential treatment for CNV. It is able to close choroidal neovascularisation while maintaining visual function in patients with classic and occult disease. Further trials of TTT are needed to compare this intervention with the natural history and other treatment modalities.

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Age related macular degeneration (ARMD) is the leading cause of visual loss in the developed world and choroidal neovascularisation (CNV) is the main cause of severe visual loss in this group.1–3 Laser photoagulation treatment for classic CNV reduces the incidence of severe visual loss in patients with extrafoveal and juxtafoveal membranes. However, because photoagulation damages neurosensory mechanisms, severe visual loss is often experienced in patients with subfoveal lesions.4–6 The benefits of laser treatment are also reduced by recurrence in many patients.7 In addition, 85–90% of patients with CNV are not eligible for argon laser treatment because the CNV is occult, or subfoveal and large.7–9 The limitations of photoagulation have led to a proliferation of new treatments (photodynamic therapy,10–14 radiotherapy,15 interferon α2,16 macular rotation,17 18 and submacular membrane excision for occult and subfoveal lesions19).

The diode laser (emission wavelength λ 810 nm, near infrared) has a number of biophysical advantages in the treatment of macular disease. It has low absorption in xanthophyll, minimising nerve fibre layer damage, it is poorly absorbed by haemoglobin, allowing treatment through preretinal and subretinal blood but is mainly absorbed in the choroid, enabling effective treatment of choroidal lesions.20 Initial studies using photocoagulation have shown it to be effective in closing CNV21–23 and in treating macular oedema.23–25 Low irradiance, long pulse diode laser irradiation (“transpupillary thermotherapy; TTT”) has been usefully employed for the treatment of choroidal melanomas.25 26

A recent report by Reichel and colleagues27 has further demonstrated the benefit of diode laser TTT for the treatment of subfoveal CNV. In a non-randomised case selected pilot study, 94% of CNV showed clinical or angiographic improvement and vision was stabilised or improved in 75%. Optical coherence tomography demonstrated membrane resolution with a restoration of the retinal anatomy.

We aimed to assess the efficacy of transpupillary thermotherapy for the treatment of both classic and occult CNV.

Materials and methods

Forty two consecutive patients with CNV secondary to ARMD, referred to the retinal clinic at King’s College Hospital, were recruited to a non-randomised pilot trial. Baseline characteristics and eligibility criteria are shown in Table 1. The patients were over 55, had signs of ARMD, and had symptoms of visual loss or metamorphopsia within the previous 4 months. There was angiographic evidence of CNV with or without pigment epithelial detachment. The Snellen visual acuity ranged between 6/9 and counting fingers. Patients who had received previous photocoagulation or were on alternative treatments for AMD were excluded from the trial, as were patients with other serious ocular disease or medical conditions that would prevent them from completing treatment.

BASELINE EXAMINATION

Initial clinical assessment included Snellen visual acuity, anterior segment examination, IOP measurement, mydriasis with 1% tropica-
mide, and funduscopy with a 78D or 60D Volk lens (Volk Optical, Mentor, OH, USA). Following this, patients were assessed with digital or photographic fundus colour (Topcon, Tokyo, Japan) and fluorescein angiography (sodium fluorescein 5 ml, 10%). Patients were counselled as to the prognosis for their condition and their treatment options. Patients not willing to undergo treatment or to complete the counselling were excluded from the trial. The patients consented to treatment and the investigation took place with the recognition of the King's College Hospital ethics committee.

**TREATMENT PROTOCOL**

Treatment was scheduled between 1 hour and 4 days of the initial angiogram. The membranes were classified in accordance with their angiographic appearance. Transpupillary thermotherapy with an 810 nm diode laser (Iris Medical OcuLight SLx, Iridex Corporation, Mountain View, CA, USA), was delivered through a dedicated slit lamp mounted delivery system. Laser settings were adjusted to give faint retinal greying following 1 minute of treatment. Beam diameters varied between 800 and 3000 µm.

The goal of TTT is to create and maintain tissue hyperthermia. It has been found that higher irradiance is needed for smaller laser spot diameters owing to the efficient heat conductance of the choroidal circulation. Hence, laser irradiance must be inversely proportional to the spot diameter, which determines the size of the thermally affected tissue volume. For a 3 mm beam the settings were between 500–700 mW (irradiance 6–9 W/cm²); for a 2 mm beam between 400–650 (irradiance 11–18 W/cm²); and for a 1.2 mm beam 300–600 mW (irradiance 24–47 W/cm²). A Volk (Mentor, OH, USA) area centralis contact lens was used in every case giving a magnification of 1.06 × of the laser spot at the retina. Patients were anaesthetised with topical benoxinate 1%. Laser targeting was achieved with a diode red beam, the lesion being covered and extending 100 µm beyond the border. If whitening of the lesion was seen the laser power was reduced by

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**Table 1 Baseline characteristics of patients**

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>No (%) of patients</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Age</strong></td>
<td></td>
</tr>
<tr>
<td>55–69</td>
<td>3 (7.2)</td>
</tr>
<tr>
<td>70–79</td>
<td>15 (35.7)</td>
</tr>
<tr>
<td>80+</td>
<td>24 (57.1)</td>
</tr>
<tr>
<td><strong>Sex</strong></td>
<td></td>
</tr>
<tr>
<td>Female</td>
<td>30 (74.8)</td>
</tr>
<tr>
<td>Male</td>
<td>12 (33.4)</td>
</tr>
<tr>
<td><strong>CNV classification</strong></td>
<td></td>
</tr>
<tr>
<td>Classic</td>
<td>12 (27.2%)</td>
</tr>
<tr>
<td>Occult</td>
<td>32 (72.8%)</td>
</tr>
<tr>
<td><strong>Position</strong></td>
<td></td>
</tr>
<tr>
<td>Extrafoveal</td>
<td>1</td>
</tr>
<tr>
<td>Juxtafoveal</td>
<td>2</td>
</tr>
<tr>
<td>Subfoveal</td>
<td>9</td>
</tr>
<tr>
<td><strong>Area on angiography/disc areas</strong></td>
<td></td>
</tr>
<tr>
<td>1–2</td>
<td>6</td>
</tr>
<tr>
<td>2–3</td>
<td>3</td>
</tr>
<tr>
<td>3+</td>
<td>1</td>
</tr>
<tr>
<td>Haemorrhage</td>
<td>7</td>
</tr>
<tr>
<td>Serous PED</td>
<td>11</td>
</tr>
</tbody>
</table>

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**Figure 1** (Top left and right) Colour and fluorescein images showing a small classic juxtafoveal membrane. (Bottom left and right) Colour and fluorescein images following TTT with resolution of the membrane; the patient's initial vision improved from 6/12 to 6/9 3 months after treatment.
Figure 2  (Top left and right) Colour and fluorescein images showing an occult subfoveal membrane. (Bottom left and right) Colour and fluorescein images following TTT, with complete closure of the CNV. The patient's vision was stabilised 3 months following treatment.

Figure 3  (Top left and right) Colour and fluorescein images showing an occult subfoveal membrane with extensive subretinal haemorrhage. (Bottom left and right) Colour and fluorescein images showing resolution of the CNV and absorption of subretinal haemorrhage 6 weeks following TTT treatment. The patient's vision improved from 6/60 to 6/24.
1–200 mW if further treatment was necessary. In patients with large lesions, confluent overlapping treatments were given.

**EVALUATION**

Follow up was arranged at 2, 4, 6 weeks and 3, 6, 9, 12, and 18 months. At each visit Snellen visual acuity and retinal examination were performed with fundus photography and fluorescein angiography, if clinically indicated. Retreatment was given to patients who retained active membranes for more than 4 weeks following TTT.

**STATISTICAL ANALYSIS**

Outcomes measured were clinical or angiographic resolution and visual acuity. The results are presented in terms of percentage of membranes resolved and number of lines of vision lost within each group. The amount of visual deterioration was assessed using Wilcoxon’s rank sum test on the Snellen data. Correlation between the visual acuity and the size of the CNV was performed using the Fisher’s R-Z correlation.

**Results**

**BASELINE CHARACTERISTICS:**

The patient characteristics are summarised in Table 1; 44 eyes of 42 patients were treated. The mean patient age was 77.9 years (SD 8.38 years); 12 (27%) membranes were classic and 32 (73%) were occult with a mean area of 1.7 DA (range of 0.5–4.0 DA). Of the classic membranes one was extrafoveal, two were juxtafoveal, and nine were subfoveal. There was no correlation between visual acuity and size of CNV $Z = -1.23$ $p=0.2$. Haemorrhage was present in 7/12 of the classic membranes and 22/32 of the occult membranes. Seven patients were pseudophakic and two had glaucoma treated with timolol 0.25% BD. Table 1 shows the baseline data.

**TREATMENT**

The average number of treatments sessions was 1.80 (SD 0.87) spread over 1–6 months.

The average power was 486 mW (140), with a beam diameter of 1.69 mm (0.63). Eyes received 4.25 (1.9) laser spots per treatment. Two patients were noted to have a significant post-treatment haemorrhage, no patients complained of pain, although some sensation was common. No anterior segment damage was noted or reported by the patients. Clinical examples of the treatment of occult and classic membranes are shown in Figures 1–3.

**PATIENT OUTCOME**

**Predominantly classic membranes**

Predominantly classic membranes were closed in 9/12 (75%, CI: 62.5–87.5) eyes and remained persistent in 3/12 (25%, 95% CI: 12.5–37.5) with no recurrences; with an average follow up of 6.7 months (SD 4.33, range 3–19) and 1.75 treatments. The initial visual acuity was 6/32.4 with a post-treatment acuity of 6/40.5 ($p=0.17$), an average of 0.75 (SD 1.75) Snellen lines lost (Fig 4A). Vision improved (>2 lines) in 0/12 patients, was stabilised (0 to + 1 line) in 7/12 (58.1%, 95% CI: 44.1–72.5) patients, there was mild loss (–1 line) in 1/12 patients (8.1% 95% CI: 3.5–16.3), and four patients lost vision (2–4 lines) 33.3% (95% CI: 19.7–46.9) (Table 2).
TTT for the treatment of choroidal neovascularisation

Figure 5  Kaplan–Meyer estimates of the cumulative proportion of eyes treated with TTT with moderate visual acuity loss (>3 Snellen lines).

Predominantly occult membranes
Predominantly occult membranes were closed in 25/32 (78%, CI: 70.1–85.3) eyes, remained persistent in 4/32 (12.5%, CI: 6.6–18.5), and were recurrent in 3/32 (5.1%, CI: 4.2–14.3) eyes; at an average follow up of 7.2 months (SD 4.33 range 3–18) and 1.8 treatments. The initial visual acuity was 6/24 with a post-treatment acuity of 6/26.4 (p=0.68), an average of 0.66 (SD 2.1) Snellen lines lost (Fig 4B). The vision improved (2+ lines) in 4/32 (12.5%, CI: 6.6–18.3) patients, 14/32 (43.8%; CI: 34.9–52.5) were stable (0 to +1 line), there was mild loss (−1 line) in 5/32 patient (15.6%, CI: 9.2–25.6), 6/32 (19.2% CI: 11.8–25.6) had moderate loss (2–4 lines), and 3/32 eyes (9.5% CI: 4.2–14.5) eyes had severe visual loss (Table 3). A Kaplan–Meyer estimate of a three line change in vision is shown in Figure 5.

Discussion
The results from this study show that TTT is effective in closing membranes in patients with CNV. Seventy seven per cent of membranes were closed at 6 months and only 7.1% developed recurrent CNV. This high closure rate was associated with a stabilisation or improvement in vision in 71% of patients with occult disease and 67% of patients with classic disease. These results compare well with the natural history of occult CNV where 63% of disease and 67% of patients with classic membranes were closed at 6 months and only 7.1% developed recurrent CNV. This high closure rate was associated with visual loss for subfoveal CNV. Seventy seven per cent of membranes

Table 4  Chart of closure rate and visual acuity results at 6 months for TTT, PDT and argon laser photocoagulation of subfoveal membranes

<table>
<thead>
<tr>
<th>Treatment</th>
<th>Closure rate</th>
<th>Visual loss /lines</th>
</tr>
</thead>
<tbody>
<tr>
<td>Argon laser</td>
<td>72%</td>
<td>−3.25</td>
</tr>
<tr>
<td>PDT</td>
<td>11.6%</td>
<td>−1.0 (3 months)</td>
</tr>
<tr>
<td>TTT</td>
<td>74%</td>
<td>−0.78</td>
</tr>
</tbody>
</table>

Their results show that, at 12 months, PDT stabilised vision in 61.2% (<3 lines lost) of the treatment group compared with 46% of the control group. The visual results of our study are comparable; 75% of patients, with predominantly classic membranes, having <3 lines of visual loss. However, TTT closed a higher proportion of CNV at 6 months (75%) compared with PDT (19%), and the control group (9%).

The MPS demonstrated that argon laser photocoagulation was effective in closing class membranes (72% at 6 months) but that this was associated with visual loss for subfoveal membranes. TTT may therefore combine the high rate of closure found with argon photocoagulation and the low recurrence rate with PDT, by closing CNV without damaging the neuroretina. Table 4 shows a comparison of the results between argon laser photocoagulation, photodynamic therapy, and transpupillary thermotherapy.

Transpupillary thermotherapy uses hyperthermia to disrupt the neovascular process and has been widely used for the treatment of small choroidal melanoma. The laser is set to a lower power with long pulse duration, the aim being to produce a moderate intraretinal temperature rise (4–9°C), compared with 40°C for conventional laser photocoagulation. The lower temperature rise confines the thermal damage profile in the target tissue, thereby reducing damage to the middle and inner retina. Conventional photocoagulation may cause dense scotoma and recurrence following argon laser photocoagulation, particularly when treating subfoveal CNV. A large spot (0.8–3 mm) size, and a long low energy pulse enables complete treatment of the lesion and allows dose titration. The mechanism of action with TTT is still unclear. However, histological sections of treated human melanoma have demonstrated intravascular thrombosis, which may account for the small amount of leakage, or haemorrhage following CNV treatment. Vascular closure may be precipitated by release of cytotoxic free radicals released from irradiated tissue. Certainly the relatively rapid involution of the CNV following TTT prevents further subretinal scarring and permanent photoreceptor damage, thereby limiting central scotomata in patients with subfoveal membranes.

The treatment variables have not been completely elucidated. In contrast with Reichel et al, we used several spots during one treatment session (average 4.25). It is clear that the end point is clinically difficult to determine (as there is little retinal reaction) and that treatment parameters may need to be adjusted for the degree of neurosensory retinal detach-
ment and the amount of subretinal haemorrhage, as well as for the location of the CNV.

Several complications have been reported following TTT for uveal tumours, including branch retinal vein and artery occlusions and retinal and choroidal haemorrhage. There also have been reports of ciliary nerve damage and corneal ulceration following panretinal photocoagulation. However, low power settings used in TTT for CNV have meant that complications are rare and patient tolerance is good.

The precise role of TTT in the treatment of CNV needs to be further defined. However, the clinical improvement found in several of our patients and the high closure rate of occult CNV membranes, with little recurrence, suggest that this treatment will have a role in the management of some patient groups. Prospective, randomised controlled trials will further clarify the role of TTT in the treatment of CNV.

We thank Dr Lyn Jenkins for coordination of angiography and image processing; Iris Medical for the loan of an OcuLight 810 nm diode laser; and Colin Clements, medical photographer, King's College Hospital, London.


Acute posterior vitreous detachment

EDITOR,—We read Tanner et al’s paper on the predictive value of vitreous pigment (Schafer’s sign) for retinal breaks in posterior vitreous detachment with great interest. Based on their figures, patients who have a negative Schafer’s sign had a 1% chance of having a retinal tear and a 0.8% chance of having a lesion where prophylaxis was thought to be appropriate. Thus, Schafer’s sign has a negative predictive value of 99% in their series. They go on to recommend that if vitreous pigment is present then the patient should be referred for urgent vitreoretinal opinion while those with no pigment should be referred on a less urgent basis.

We would like to put these findings in perspective. The incidence of retinal breaks in patients aged 10 years or more who do not have any history of ocular disease is 6–14%.² Retinal breaks have been found in 37/250 (14.8%) of necropsy eyes with posterior vitreous detachment by Foos.³ The incidence of retinal detachment is approximately 12/100 000 of the general population per year.² This suggests that less than 0.2% people with a retinal break eventually have a detachment of the retina. This value may be higher in patients with a symptomatic posterior vitreous detachment; however, it is reasonable to conclude that only a minority of retinal breaks will go on to cause a retinal detachment. Prophylactic treatment of retinal breaks by laser or cryotherapy is not without complications; also detachments can occur in eyes that have had prophylactic treatment.¹ Byer has reported that retinal breaks in unoperated eyes with posterior vitreous detachment that need treatment can be followed up without treatment, with only a minority progressing to retinal detachments.⁴ We have a test that has a negative predictive value of 99%. We know that only a minority of patients who have a retinal tear or hole actually benefit from prophylactic treatment. Can we still justify referring all patients with a retinal tear or hole and a 0.5% chance of having a lesion where prophylaxis was thought to be appropriate? Thus, the management of recent onset, symptomatic posterior vitreous detachment (PVD) and associated retinal breaks in patients who have a retinal tear or hole and a 0.5% chance of having a lesion where prophylaxis was thought to be appropriate is still necessary. In our series only one patient represented a retinal break which had not been identified during initial indentation ophthalmoscopy. The presence of vitreous pigment in patients presenting with acute PVD is indeed highly predictive of the presence of a retinal break, but a thorough retinal examination is still necessary. In our series only one patient represented a retinal break which had not been identified during initial indentation ophthalmoscopy. The presence of vitreous pigment in patients presenting with acute PVD is indeed highly predictive of the presence of a retinal break, but a thorough retinal examination is still necessary.

The rationale for treating fresh, symptomatic retinal breaks has been reviewed by numerous contributors, Drs De Kozak, Forrester, Caspers-Velu, LeHoang, McCluskey, Okada, and Whitcup provide a comprehensive yet concise summary of this field in their recently released text, which was prepared under the auspices of the International Ocular Inflammation Society. Each of the eight chapters contained in this book are clearly written, well referenced, and accompanied by a number of very nice tables and illustrations, many of which are in full colour. Although by no means encyclopaedic, this manual includes excellent overviews of the clinical characteristics of all of the important causes of anterior uveitis, as well as systematic approaches to diagnostic, treatment, and surgery in patients with anterior uveitis.

EMMETT T CUNNINGHAM, JR

BOOK REVIEW


Anterior uveitis is an important and common cause of ocular morbidity. Drs BenEzra, Ohno, Secchi, and Ali are well known for their contributions. Drs De Kozak, Forrester, and Whitcup provide a comprehensive yet concise summary of this field in their recently released text, which was prepared under the auspices of the International Ocular Inflammation Society. Each of the eight chapters contained in this book are clearly written, well referenced, and accompanied by a number of very nice tables and illustrations, many of which are in full colour. Although by no means encyclopaedic, this manual includes excellent overviews of the clinical characteristics of all of the important causes of anterior uveitis, as well as systematic approaches to diagnostic, treatment, and surgery in patients with anterior uveitis.

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NOTICES

Vision 2020: cataract outcomes

The latest issue of Community Eye Health (35) discusses cataract surgery outcomes. For further information please contact Community Eye Health, International Centre for Eye Health, Institute of Ophthalmology, 11–43 Bath Street, London EC1V 9EL. Tel: (+44) (0) 20-7608 6909/6910/6923; fax: (+44) (0) 7250 3207; email: eyeresource@ucl.ac.uk)

Annual subscription £25. Free to workers in developing countries.

Second Sight

Second Sight, a UK based charity whose aims are to eliminate the backlog of cataract blind in India by the year 2020 and to establish strong links between Indian and British ophthalmologists, will be sending volunteer surgeons to India early in 2001. Details can be found at the charity website at www.second-sight.org.uk or by contacting Dr Lucy Mathen (email address lucymathen@yahoo.com).

Office of Continuing Medical Education

A symposium “Randomised trials in ophthalmology: past, present, future” will be held 2–3 April 2001 at the Thomas B Turner Building, Johns Hopkins University School of Medicine, Baltimore, Maryland, USA. Further details: Johns Hopkins University School of Medicine, Office of Continuing Medical Education, Turner 20, 720 Rutland Avenue, Baltimore, MD 21205-2195, USA (tel: (410) 955-2959; fax: (410) 955-0807; email: cmene9@jhmi.edu).

XXV Detachment Course

The XXV Detachment course, retinal and vitreous surgery, will be held in Poznan, Poland on 5–6 April 2001. Further details: Professor Krystyna Pecold, Katedra I Klinika Oku- listyki, ul Dluga 1/2, 61–848 Poznan, Poland (tel/fax: 004861-8527619) or Professor Ingrid Kreissig, Univ-Augenklinik, Schleichstrasse 12, D-72076 Tuebingen, Germany (fax: 49-7071-293746; email: ingrid.kreissig@uni-tuebingen.de).

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MAILBOX

Correspondence to: Mr Mohit Gupta

Optometry 01
Optometry 01 will take place on 21–23 April 2001 with more than 100 events—lectures and workshops—at the Atrium Gallery, NEC, Birmingham, UK. Further details: tel: 0207 261 9661; email: info@optometry01.co.uk; website: www.optometry01.co.uk.

14th Annual Meeting of German Ophthalmic Surgeons
The 14th Annual Meeting of German Ophthalmic Surgeons will be held in the Meistersingerhalle, Nuremberg, Germany on 17–20 May 2001. Further details: MCN Medizinische Congress-organisation Nuremberg, AG, Zentralbüro, Hauptstrasse 29, 90478 Nuremberg, Germany (tel: ++49-911-3931621; fax: ++49-911-3931620; email: doerflinger@mcn-nuernberg.de).

European Association for the Study of Diabetic Eye Complications (EASDEC)
The next meeting of the European Association for the Study of Diabetic Eye Complications (EASDEC) will be held in Paris, France, on 19–20 May 2001. Further details: Colloquium, 12 Rue de la Croix Faubin, 75 557 Paris Cedex 11, France (tel: +33-1-44 64 15 15; fax +33-1-44 64 15 10; email: s.mundler@colloquium.fr).

2nd Interdisciplinary Symposium on the Treatment of Autoimmune Disorders 2001
The 2nd Interdisciplinary Symposium on the Treatment of Autoimmune Disorders 2001 will take place on 7–9 June 2001 at the University, Zentralbüro, University of Kiel, Kiel, Germany. Further details: Prof Dr Med Michael Sticherling, Department of Dermatology, University of Kiel, Schittenhelmstrasse 7, D-24105 Kiel, Germany (tel: +49-431 597 1512; fax: +49-431 597 1611; email: msticherling@dermatology.uni-kiel.de).

14th World Congress of the International Society for Laser Surgery and Medicine
The 14th World Congress of the International Society for Laser Surgery and Medicine is to be held on the 27–30 August 2001 at Sri Ramachandra Medical College and University Hospital, Chennai, India. The American Society of Lasers in Medicine and Surgery has indicated that it will designate the 14th World Congress of ISLSM as its society’s co-sponsoring meeting. A pre-conference course and separate sessions in ophthalmology will be held as a part of this international meeting.

Further details: Dr B Krishna Rau, President, 14th World Congress of the International Society for Laser Surgery and Medicine, Department of Surgery, D2 Ward, Sri Ramachandra Medical College and Research Institute, Porur, Chennai - 600 116, India (tel: 91-44-47655756; fax: 4768027-28, 8527776; 95749804; fax: 91-44-8594578, 4767008; email: krishnar@giasm01.vsnl.net.in and website: www.medindia.net/islsm2001).

American Institute of Ultrasound in Medicine—Millennium Ultrasound Course Series
A course entitled “Obstetrical and Gynecological Ultrasound” will be held in New York City, NY, on 24–26 August 2001. Further details: Stacey Blessing, Public Relations Coordinator, AIUM, 14750 Sweitzer Lane, Suite 100, Laurel, MD 20707-5906, USA (tel: 301-498-4100; email: sblessing@aium.org).

31st Cambridge Ophthalmological Symposium
The 31st Cambridge Ophthalmological Symposium will be held 3–5 September 2001 at St John’s College Cambridge. The subject is Retinal Detachment. Further details: COS Secretariat, Cambridge Conferences, The Lawn, 33 Church Street, Great Shelford, Cambridge CB2 5EL, UK (tel: 01223 847464; fax: 01223 847465; email: b.ashworth@easynet.co.uk).

1st Asia Pacific Forum on Quality Improvement in Health Care
The 1st Asia Pacific Forum on Quality Improvement in Health Care will be held from 19–21 September 2001 in Sydney, Australia. Presented by the BMJ Publishing Group (London, UK) and Institute for Healthcare Improvement (Boston, USA), with the support of the Commonwealth Department of Health and Aged Care (Australia), Safety and Quality Council (Australia), NSW Health (Australia) and Ministry of Health (New Zealand). Further details: quality@bma.org.uk; fax +44 (0) 7383 6360.

41st St Andrew’s Day Festival Symposium on Therapeutics
The 41st St Andrew’s Day Festival Symposium on Therapeutics will be held on 6–7 December 2001 at the Royal College of Physicians of Edinburgh. Further details: Ms Eileen Strawn, Symposium Co-ordinator (tel: 0131 225 7324; fax: 0131 220 4393; email: e.strawn@rcpe.ac.uk; website: www.rcpe.ac.uk).

4th International Conference on the Adjuvant Therapy of Malignant Melanoma
The 4th International Conference on the adjuvant therapy of malignant melanoma will be held at The Royal College of Physicians, London on 15–16 March 2002. Further details: Conference Secretariat, CCI Ltd, 2 Palmerston Court, Palmerston Way, London SW8 4AJ, UK (tel: + 44 (0) 20 7720 0600; fax: + 44 (0) 20 7720 7177; email: melanoma@confcomm.co.uk; website: www.confcomm.co.uk/Melanoma).

International Society for Behçet’s Disease
The International Society for Behçet’s Disease was inaugurated at the 9th International Congress on Behçet’s Disease. Professor Shigeaki Ohno represents the ophthalmology division (Department of Ophthalmology and Visual Sciences, Hokkaido University Graduate School of Medicine, Sapporo, Japan: tel: +81-11-716-1161 (ext 5944); fax: +81-11-736-0952; email: sohno@med.hokudai.ac.jp). The 10th International Congress on Behçet’s Disease will be held in Berlin 27–29 June 2002. Further details: Professor Ch Zouboulis (email: zoubbere@zedat.fu-berlin.de).

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