· Original article ·

# Research progress of treatment for amblyopia

Hua Li<sup>1</sup>, Qing Ling Xiang<sup>1</sup>, Meng Liao<sup>2</sup>, Long Qian Liu<sup>2</sup>

<sup>1</sup>Department of Ophthalmology, Yongchuan Hospital, Chongqing Medical University, Chongqing 402160, China <sup>2</sup>Department of Ophthalmology, West China Hospital, Sichuan University, Chengdu 610041, Sichuan

**Correspondence to:** Long-Qian Liu. Department of Ophthalmology, West China Hospital, Sichuan University, Chengdu 610041, Sichuan Province, China. bq15651 @ hotmail.com

Received: 2010-12-13 Accepted: 2011-02-13

#### **Abstract**

Province, China

• The most common forms of treatment for amblyopia are occlusion therapy and pharmacologic penalization. But these methods can not recover all the visual deficits of amblyopes and there are some disadvantages of the treatment which need to be considered, including adverse effects, compliance and social stigma. Recently, some novel approaches are investigated for improving the effectiveness and compliance of treatment for amblyopia.

• KEYWORDS: amblyopia; perceptual learning; virtual reality; effectiveness; compliance

DOI: 10.3969/j. issn. 1672-5123.2011.03.002

Li H, Xiang QL, Liao Meng, Liu LQ. Research progress of treatment for amblyopia. *Guoji Yanke Zazhi( Int J Ophthalmol)* 2011; 11(3):385-387

#### INTRODUCTION

A mblyopia is a developmental disorder of spatial vision caused by abnormal visual stimulation during sensitive period  $^{[1]}$ , which is characterized by reduced visual functions in the absence of any detectable structural abnormalities. Most of the cases are due to squint or refractive error  $^{[2]}$ . It is the most common ocular disorder in children, affecting approximately 3% of the population worldwide  $^{[3,4]}$ . So it is critical for us to treat amblyopia effectively and safely.

### CONVENTIONAL TREATMENT FOR AMBLYOPIA

Occlusion therapy and pharmacologic penalization are the most common forms of treatment for amblyopia. But these methods can not recover all the deficits that caused by amblyopia even long after the visual acuity had become normal, and there is no convincing evidence that they are clinically effective for patients > 12 years old up to now<sup>[5, 6]</sup>. Furthermore there are some disadvantages which need to be considered before they are prescribed.

Except for the side-effects, several studies indicated that both occlusion and penalization in amblyopia treatment might have an impact on the child's behavior or family life recently [7].

Occlusion therapy is not always effective in treating pediatric patients, about one third of the children gain little or no visual benefit despite prolonged treatment<sup>[8,9]</sup>. Perhaps compliance is the critical factor which has an influence on the effect of occlusion therapy. But full concordance with patching is very difficult because of restriction of visual field and the adverse effects such as skin irritation, allergy and so on [10, 11]. Moreover patching is difficult to implement for the reason that patching may interfere with social activities or have an impact on educational development<sup>[9, 12]</sup>. Pharmacologic penalization is an alternative to patching for amblyopia therapy when the child is intolerant of patching. But atropine works less quickly than occlusion initially, and there are some side-effects from atropine use, including light sensitivity, allergy, dry mouth, hyperactivity and tachycardia, which need to be considered<sup>[5, 10, 11]</sup>. So it is reasonable to explore safer method for improving the effectiveness and compliance of treatment which can fully recover visual functions for amblyopes.

#### NOVEL APPROACHES TO TREAT AMBLYOPIA

Recently, perceptual learning (practice-induced improvement in performance on specific task) and the Interactive Binocular Treatment [12] (I-BiTTM, a computer-based virtual reality treatment) system are investigated as novel approaches for improving the effectiveness and compliance of treatment for amblyopia. Many studies have delivered encouraging results in recent years. Perceptual learning can improve performance on many visual tasks and learning can generalize to other tasks following some degree of inter-ocular transfer to the untrained fellow eyes [13, 14]. The Interactive Binocular Treatment system [12] encourages the two eyes to work together, which can avoid disrupting fusion during treatment [15]. Furthermore there is no evidence that they have the negative effects mentioned above as yet.

Perceptual Learning Amblyopes show abnormal performance in many visual functions except for reduced visual acuity, which include hyperacuity [16, 17], and contrast-sensitivity function et al. But the traditional methods could not recover all the abnormalities [6]. Recently, a lot of studies showed that perceptual learning could improve performance on many visual tasks in amblyopic eyes, which had more benefits than traditional methods. Perceptual learning is more active and intensive than passive patching [18], so it is more acceptable.

Several studies indicated that perceptual learning was useful for different age groups to improve visual functions. In 1997, Levi *et al* [19] found that training with Vernier task could lead to significant improvements in Vernier actuity in the amblyopic eyes, in some of these observers the improvement was

accompanied by some improvement in Snellen acuity. In addition, their results showed that adults with amblyopia retain some degree of neural plasticity. From then on perceptual learning has received more and more attention as a method for treating amblyopia. Other studies also showed that perceptual learning could induce some degree of recovery of vision in amblyopic eyes. In three recently studies, Li et al found that practicing position-discrimination could improve positional and Snellen acuities, and they also found that the improvement transferred to letter acuity [18, 20, 21]. prospective, randomized, masked, controlled study, Polat et al<sup>[22]</sup> used training tasks that emphasized lateral interactions to aim directly at the deficient connectivity in amblyopes. They demonstrated that perceptual learning could improve contrast sensitivity and visual acuity significantly in amblyopes (between 9 and 55 years old). Their results support the use of perceptual learning in amblyopia to improve vision of adults. In order to understand the basis of neural plasticity in amblyopia, Zhou et al<sup>[14]</sup> used sine-wave gratings to evaluate effects of perceptual learning in young adults and older children with amblyopia. They found that training could improve visual acuity and contrast-sensitivity functions and the improvement could generalize to the fellow eyes. Focusing on juvenile amblyopia, Li et al<sup>[21]</sup> found that perceptual learning may speed up the time of recovery. Compared with the tasks as we have said above, perceptual learning of contrast-defined letters may have limited value in treatment of amblyopia, but adults with amblyopia can also benefit from it [23]. These results support the use of perceptual learning to improve vision of amblyopes which target at the specific deficiencies. If we combine different visual-tasks for amblyopia treatment [18], perhaps we could fully recover visual functions of amblyopes. Perceptual learning is usually very specific for the exact task trained, however, it can generalize to other task. This can be explained by the reverse hierarchy theory of perceptual learning, which assumes that learning starts at high-levels of visual system, where it generalizes and progresses backwards towards input levels, where it is  $specific^{[13, 24]}$ . Now several studies had shown that the improvement with perceptual learning could generalize to untrained eye, and other untrained, higher level visual tasks. Some of them indicated that the improvement can be maintained for a long period. These evidences also support the use of perceptual learning to improve visual functions of amblyopes.

#### Interactive Binocular Treatment (I-BiTTM ) System

However, nearly in all these studies mentioned above observers were required to covered the fellow eye during training. Therefore they can not avoid the negative impact on children that accompanies occlusion therapy completely<sup>[7]</sup>. The Interactive Binocular Treatment (I-BiTTM) system was developed by Eastgate *et al* <sup>[12]</sup> to treat amblyopia with both eyes stimulated simultaneously. This system incorporates virtual reality (VR) technology and specific software which provides interactive games (2D or 3D) and videos to the patient via a stereo (binocular) display. Waddingham *et al* <sup>[15]</sup> demonstrated that this system could improve vision in the

amblyopic eyes of children within a short period of time while avoiding occlusion of the nonamblyopic eye. This is an exciting new technique, which is positive and enthusiastic for patients.

#### **CONCLUSION**

The most common forms of treatment for amblyopia are occlusion therapy and pharmacologic penalization. But these methods can not recover all the visual deficits of amblyopes and there are some disadvantages of the treatment which need to be considered, including adverse effects, compliance and social stigma. Perceptual learning and the Interactive Binocular Treatment (I-BiTTM) system can both improve vision in the amblyopic eyes. Ophthalmologists can use these new effective approaches clinically which can avoid the disadvantages of conventional therapy. However, before these approaches is used widely, there are still many questions to be resolved.

#### REFERENCES

- 1 Hubel DH, Wiesel TN. The period of susceptibility to the physiological effects of unilateral eye closure in kittens. *J Physiol* 1970; 206 (2): 419-436
- 2 Attebo K, Mitchell P, Cumming R, Smith W, Jolly N, Sparkes R. Prevalence and causes of amblyopia in an adult population. *Ophthalmology* 1998;105(1):154-159
- 3 Brown SA, Weih LM, Fu CL, Dimitrov P, Taylor HR, McCarty CA. Prevalence of amblyopia and associated refractive errors in an adult population in Victoria, Australia. *Ophthalmic Epidemiol* 2000; 7 (4): 249-258
- 4 Newman DK, East MM. Prevalence of amblyopia among defaulters of preschool vision screening. *Ophthalmic Epidemiol* 2000;7(1):67-71
- 5 Wu C, Hunter DG. Amblyopia: diagnostic and therapeutic options. Am J Ophthalmol 2006;141(1):175-184
- 6 Huang C, Tao L, Zhou Y, Lu ZL. Treated amblyopes remain deficient in spatial vision: a contrast sensitivity and external noise study. *Vision Res* 2007;47(1):22-34
- 7 Webber AL. Amblyopia treatment: an evidence-based approach to maximising treatment outcome. Clin Exp Optom 2007;90(4):250-257
- 8 Clarke MP, Wright CM, Hrisos S, Anderson JD, Henderson J, Richardson SR. Randomised controlled trial of treatment of unilateral visual impairment detected at preschool vision screening. *BMJ* 2003;327 (7426):1251-1254
- 9 Webb BS, McGraw PV, Levi DM. Learning with a lazy eye: a potential treatment for amblyopia. *Br J Ophthalmol* 2006;90(4): 518 10 Holmes JM, Clarke MP. Amblyopia. *Lancet* 2006;367(9519):1343-1351
- 11 Stewart CE. Treatment of amblyopia; evidence-based medicine at long last. *Techniques in Ophthalmology* 2005;3(2);94-99
- 12 Eastgate RM, Griffiths GD, Waddingham PE, Moody AD, Butler TK, Cobb SV, Comaish IF, Haworth SM, Gregson RM, Ash IM, Brown SM. Modified virtual reality technology for treatment of amblyopia. *Eye* 2006;20(3):370-374
- 13 Fahle M. Perceptual learning: specificity versus generalization. *Curr Opin Neurobiol* 2005;15(2):154-160
- 14 Zhou Y, Huang C, Xu P, Tao L, Qiu Z, Li X, Lu ZL. Perceptual learning improves contrast sensitivity and visual acuity in adults with anisometropic amblyopia. *Vision Res* 2006;46(5):739-750
- 15 Waddingham PE, Butler TK, Cobb SV, Moody AD, Comaish IF, Haworth SM, Gregson RM, Ash IM, Brown SM, Eastgate RM, Griffiths GD. Preliminary results from the use of the novel Interactive binocular treatment (I-BiT (TM)) system, in the treatment of strabismic and anisometropic amblyopia. *Eye* 2006;20(3):375-378

- 16 Fronius M, Sireteanu R, Zubcov A. Deficits of spatial localization in children with strabismic amblyopia. *Graefes Arch Clin Exp Ophthalmol* 2004;242(10):827-839
- 17 Kelly SL, Buckingham TJ. Movement hyperacuity in childhood amblyopia. *Br J Ophthalmol* 1998;82(9):991-995
- 18 Li RW, Young KG, Hoenig P, Levi DM. Perceptual learning improves visual performance in juvenile amblyopia. *Invest Ophthalmol Vis Sci* 2005;46(9):3161-3168
- 19 Levi DM, Polat U, Hu YS. Improvement in Vernier acuity in adults with amblyopia-Practice makes better. *Invest Ophthalmol Vis Sci* 1997; 38(8):1493-1510
- 20 Li RW, Levi DM, Klein SA. Perceptual learning improves efficiency by re-tuning the decision 'template' for position discrimination. *Nat Neurosci* 2004;7(2):178-183
- 21 Li RW, Provost A, Levi DM. Extended perceptual learning results in substantial recovery of positional acuity and visual acuity in juvenile amblyopia. *Invest Ophthalmol Vis Sci* 2007;48(11): 5046-5051
- 22 Polat U, Ma-Naim T, Belkin M, Sagi D. Improving vision in adult amblyopia by perceptual learning. *Proc Natl Acad Sci USA* 2004; 101 (17);6692-6697
- 23 Chung STL, Li RW, Levi DM. Identification of contrast-defined

letters benefits from perceptual learning in adults with amblyopia. Vision  $Res\ 2006\ ; 46\ (22)\ ; 3853\ -3861$ 

24 Ahissar M, Hochstein S. The reverse hierarchy theory of visual perceptual learning. *Trends Cogn Sci* 2004;8(10):457-464

## 弱视治疗方法的研究进展

李 华1,向庆龄1,廖 孟2,刘陇黔2

(作者单位: 1402160 中国重庆市,重庆医科大学附属永川医院眼科; 2610041 中国四川省成都市,四川大学华西医院眼科)作者简介:李华,硕士,主治医师,研究方向:斜弱视、视光学。通讯作者:刘陇黔,博士,教授,研究方向:斜弱视、视光学.bq15651@ hotmail. com

#### 摘要

遮盖疗法和阿托品压抑疗法是最常用的弱视治疗方法。 但是这些传统的方法并不能恢复弱视患者的所有视功能, 而且还有一定的缺点,包括副作用、依从性和社会影响。 近年来,人们逐渐开始应用一些可以提高弱视治疗的有效 性和依从性的新方法来治疗弱视患者。

关键词:弱视;知觉学习;虚拟现实;有效性;依从性