Huperzine-A capsules enhance memory and learning performance in 34 pairs of matched adolescent students¹

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KEY WORDS students; huperzine-A; memory; learning; aptitude tests; adolescent behavior; double-blind method; matched-pair analysis; cholinesterase inhibitors; Wechsler scales

ABSTRACT

AIM: To study the efficacy of huperzine-A capsules (Hup) on memory and learning performance of adolescent students. METHODS: Using double-blind and matched pair method, 34 pairs of junior middle school students complaining of memory inadequacy were divided into two groups by normal psychological health inventory (PHI), similar memory quotient (MQ), same sex and class. The Hup group was administrated orally 2 capsules of Hup (each contains Hup 50 μ g) bid, and the placebo group was given 2 capsules of placebo (starch and lactose inside) bid for 4 wk. RESULTS: At the end of trial, the Hup group's MQ (115 \pm 6) was more than that of the placebo group $(104 \pm 9, P < 0.01)$, and the scores of Chinese language lesson in the Hup group were elevated markedly too. CONCLUSION: The Hup capsules enhance the memory and learning performance of adolescent students.

INTRODUCTION

Hup, a new *Lycopodium* alkaloid first isolated from Chinese herb *Huperzia serrata* (Thunb) Trev by Chinese^[1], is a potent, centrally active, and reversible cholinesterase inhibitor^[2], it was reported to ameliorate

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learning and memory retention in rodents^[3] and improve memory in aged^[4,5]. As to the efficacy in adolescents, it had not been reported at home and abroad yet, so we selected 68 junior middle school students to study that from April to June in 1997.

MATERIALS AND METHODS

Sample origin Sixty-eight students were selected from total of 1356 students of 23 classes of Zhaohui and Huilan junior middle schools in Xiaoshan City, where psychological health work had been done better.

Inclusive criteria The student complained of bad memory, the learning performance was getting worse, psychological health inventory (PHI) was normal, no mental symptom or sign, and no any nervous illness history was sampled in the study. All of them and their parents agreed to take part in this research.

Method of entering the group Sixty-eight students were matched in the same class, the same sex, and similar memory quotient (MQ), then, divided into two groups as the Hup group and the placebo (Pla) group randomly.

Method of taking medicine Students in Hup group were given 2 capsules of Hup (each contains Hup $50~\mu g$), and Pla group were given 2 capsules of Pla (starch and lactose inside), twice a day for 4 wk. Hup and Pla capsules, same in shape, color, weight, taste and the packaging, were provided by Ningbo Lihua Pharmaceutic Company Ltd. The clinicians, the teachers and the students were all blind.

Assessment All samples were evaluated with Wechsler memory scale (WMS) and treatment emergent symptom scale (TESS) at the beginning and the end of trial, we further took the quiz performance of English, Chinese, mathematics, and the average

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score of above three lessons at 'before' and 'after' trial as the another targets of efficacy.

Data analysis The statistical analysis of the results were performed by NDST (New Drug Statistical Treatment) software provided by Prof SUN Rui-Yuan. Paired t test was used for MQ, TESS and all the quiz performance, before and after the trial; group t test was used for comparison between Hup and Pla groups.

RESULTS

The intraclass correlation (ICC) ICC on WMS from 4 evaluators at the beginning of the trial was 0.82.

General data According to the matched criteria, 20 pairs of students were sampled from Zhaohui and 14 pairs from Huilan junior middle school, their background data were listed in Tab 1.

Tab 1. Comparison of the background data between two groups. n=34 pairs of students. $\bar{x}\pm s$. $^{a}P>0.05$ vs placebo.

Item	Huperzine	Placebo
Sex (M/F)	20/14	20/14
Age/a	15.0 ± 0.6^{0}	15.0 ± 0.8
Memory quotient	$92 \pm 7^{\rm a}$	94 ± 8
Psychological health invento	nty	
Lie	53 ± 9ª	54 ± 8
Fake	51 ± 12^{4}	48 ± 11
Question	46 ± 4^a	45 ± 5
Somatic disorder	$5I \pm 8^a$	50 ± 7
Depression	56 ± 7°	54 ± 7
Anxiety	55 ± 8°	52 ± 7
Psychopathy deviance	52 ± 10^{a}	50 ± 9
Hypochondriasis	55 ± 11^{a}	52 ± 10
Unrealistic	51 ± 9ª	50 ± 9
Hypomania	54 ± 10^a	51 ± 9

It showed no statistical difference on sex, age, MQ, and PHI among them.

MQ assessment There were significant differences on MQ in both groups between 'before' and 'after' the 4-wk trial (P < 0.01), but the MQ of Hup group at end of trial was significantly more than that of Pla group (P < 0.01, Tab 2).

Group t test on each factor of WMS between 2 groups showed a significant improvement in 'accumula-

Tab 2. Comparison of memory quotient between 2 groups before and after trial. n = 34. $\bar{x} \pm s$. $^{3}P > 0.05$, $^{5}P < 0.01$ vs placebo.

Memory quotient	Huperzine A	Placebo
Baseline	92 ± 7ª	94±8
4-wk trial	115 ± 6°	104 ± 9
Odds	23 ± 7^{c}	11 ± 10

tion', 'recognition', 'reproduction', 'association', 'tactual memory', and 'number of recitation' (P < 0.05), (Tab 3).

Tab 3. Comparison of factors of Wechsler memory scale between 2 groups before and after trial.

n = 34. $\bar{x} \pm s$. b P < 0.05, $^{c}P < 0.01$ vs placebo.

Item		Huperzine A	Placebo
I→100	baseline	10.6±2.5	10.9 ± 2.1
	4-wk trial	10.5 ± 2.8	10.9 ± 2.0
100→I	baseline	10.7 ± 2.3	11.3 ± 2.2
	4-wk trial	12.4 ± 1.3	12.6 ± 1.5
Accumulation	baseline	10.4 ± 1.8	10.0 ± 2.4
	4-wk trial	11.8 ± 1.2^{h}	10.7 ± 2.2
Picture memory	baseline	9.2 ± 1.8	9.9 ± 1.8
•	4-wk trial	10.8 ± 1.8	10.3 ± 1.3
Recognition	baseline	10.0 ± 2.0	9.9 ± 2.2
ŭ	4-wk trial	10.5 ± 1.7^{b}	9.3 ± 2.7
Reproduction	baseline	9.6 ± 2.2	8.9 ± 1.9
•	4-wk trial	$11.5 \pm 1.5^{\circ}$	10.3 ± 1.6
Association	baseline	9.1 ± 2.8	9.6 ± 2.3
	4-wk trial	$12.2 \pm 1.4^{\circ}$	9.9 ± 2.5
Tactual memory	baseline	10 ± 4	10.2 ± 2.8
,	4-wk trial	$13.7 \pm 2.3^{\circ}$	10.8 ± 2.7
Understanding	baseline	14 ± 4	I4 ± 4
	4-wk trial	17.6 ± 2.5	17.4 + 2.5
Number of recitation	baseline	10±3	10.3 ± 2.3
	4-wk trial	12.8 ± 2.7^{b}	11.5 ± 2.6

Learning performance The quiz scores of Chinese language lesson in the Hup group were elevated significantly (Tab 4).

Side effects Both the scores of TESS in two groups were zero, it indicated that no any side effect was found in the 4-wk trial.

DISCUSSION

This study was designed as a double blind, matched

Tab 4. Comparison of learning performance between 2 groups before and after trial. n = 34. $\bar{x} \pm s$. ${}^{b}P < 0.05$, ${}^{c}P < 0.01$ vs baseline; ${}^{c}P < 0.05$, ${}^{c}P < 0.01$ vs placebo.

Item	Huperzine A	Placebo
English language		
baseline	59 ± 20	68 ± 18
4-wk trial	$66 \pm 18^{\circ}$	70 ± 17^{b}
Odds	6 ± 8	3 ± 12
Chinese language		
baseline	59 ± 16^{f}	68 ± 9
4-wk trial	70 ± 12^{c}	70 ± 10
Odds	10 ± 9^{c}	2 ± 7
Mathematics		
baseline	58 ± 21	58 ± 22
4-wk trial	$68 \pm 21^{\circ}$	$68 \pm 18^{\circ}$
Odds	9 ± 11	10 ± 14
Average score of ab-	ove 3 lessons	
baseline	59 ± 14	64 ± 12
4-wk trial	68 ± 14^{c}	$70 \pm 12^{\circ}$
Odds	9 ± 6^{e}	6 ± 7

pair, and randomly grouping trial to observe the action There was a 601-603 of Hup in adolescent students. considerable high ICC in evaluators, and comparable background data between 2 groups, we considered that the results of this study were relative reliable.

The results of this study exhibited that Hup obviously improved the memory function of adolescent students too. At the end of trial, the MQ of Hup group was significantly elevated than that of Pla group (P < 0.01). According to the further analysis of each factor of the WMS. Hup increased the scores of 'accumulation', 'recognition', 'reproduction'. 'association', 'tactual memory', and 'number of recitation' five factors, but not 'understanding' factor. These findings were consistent with the finding of learning performance, that is, Hup enhanced the performance of English and Chinese language lessons, but not mathematics. So, we think that Hup is also a promising candidate drug for improving the memory function and learning aptitude in adolescent students.

Owing to the insufficient observation period and lack of biochemical experiments, the results of this study are preliminary, the next strict designed study will be undertaken for identifying the efficacy of the Hup in the adolescents.

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石杉碱甲胶囊提高 34 对青春期学生记忆和学习成绩1

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关键词 学生; 石杉碱甲; 记忆; 学习; 才能测试; 青春期行为;双盲法;配对分析;胆碱酯酶抑制 剂; 韦氏量表

目的: 研究石杉碱甲胶囊对初中学生记忆和学习 成绩的效能, 方法,采用双盲法,按照心理健康 正常、记忆商接近、同班、同性别的要求,将68 例初中学生配对成两组,随机各给石杉碱甲胶囊 2 粒(每粒含 50 µg)或空白胶囊 2 片(含淀粉和乳 糖), 一日两次口服, 4 周为一疗程。 以记忆商及 语文、英语和数学以及上述三课均分为观测指标 评定疗效. 结果: 石杉碱甲胶囊组疗末的记忆商 明显高于空白组(P<0.01), 语文成绩也明显提 高. 结论: 石杉碱甲胶囊提高青春期学生的记忆 和学习成绩. (责任编辑 李