# World Journal of *Clinical Cases*

World J Clin Cases 2023 May 16; 11(14): 3114-3368





Published by Baishideng Publishing Group Inc

W J C C World Journal of Clinical Cases

#### Contents

#### Thrice Monthly Volume 11 Number 14 May 16, 2023

#### **OPINION REVIEW**

3114 Modernising autism spectrum disorder model engineering and treatment via CRISPR-Cas9: A gene reprogramming approach

Sandhu A, Kumar A, Rawat K, Gautam V, Sharma A, Saha L

#### **REVIEW**

Burden of disability in type 2 diabetes mellitus and the moderating effects of physical activity 3128 Oyewole OO, Ale AO, Ogunlana MO, Gurayah T

#### **MINIREVIEWS**

Postoperative hypoxemia for patients undergoing Stanford type A aortic dissection 3140 Liu HY, Zhang SP, Zhang CX, Gao QY, Liu YY, Ge SL

#### **ORIGINAL ARTICLE**

#### **Case Control Study**

- 3148 Impact of extended nursing model after multi-disciplinary treatment on young patient with post-stroke Xu XY, Pang ZJ, Li MH, Wang K, Song J, Cao Y, Fang M
- 3158 Changes and significance of serum ubiquitin carboxyl-terminal hydrolase L1 and glial fibrillary acidic protein in patients with glioma

Zhu QH, Wu JK, Hou GL

#### **Retrospective Study**

Multitrack and multianchor point screw technique combined with the Wiltse approach for lesion 3167 debridement for lumbar tuberculosis

Yuan YF, Ren ZX, Zhang C, Li GJ, Liu BZ, Li XD, Miao J, Li JF

Clinical features and prognostic factors in 49 patients with follicular lymphoma at a single center: A 3176 retrospective analysis

Wu H, Sun HC, Ouyang GF

3187 Value of optical coherence tomography measurement of macular thickness and optic disc parameters for glaucoma screening in patients with high myopia

Mu H, Li RS, Yin Z, Feng ZL

#### **Observational Study**

3195 Comparative study of the clinical efficacy of all-inside and traditional techniques in anterior cruciate ligament reconstruction

An BJ, Wang YT, Zhao Z, Wang MX, Xing GY



•	World Journal of Clinical Cases
Conten	Thrice Monthly Volume 11 Number 14 May 16, 2023
3204	Positioning and design by computed tomography imaging in neuroendoscopic surgery of patients with chronic subdural hematoma
	Wang XJ, Yin YH, Zhang LY, Wang ZF, Sun C, Cui ZM
3211	Evaluation of chronic idiopathic tinnitus and its psychosocial triggers
	Hamed SA, Attiah FA, Fawzy M, Azzam M
3224	Intestinal complications in patients with Crohn's disease in the Brazilian public healthcare system between 2011 and 2020
	Sassaki LY, Martins AL, Galhardi-Gasparini R, Saad-Hossne R, Ritter AMV, Barreto TB, Marcolino T, Balula B, Yang- Santos C
	Randomized Controlled Trial
3238	Effect of non-pharmacological treatment on the full recovery of social functioning in patients with attention deficit hyperactivity disorder
	Lv YB, Cheng W, Wang MH, Wang XM, Hu YL, Lv LQ
	CASE REPORT
3248	Diagnosis of tuberculous uveitis by the macrogenome of intraocular fluid: A case report and review of the literature
	Zhang YK, Guan Y, Zhao J, Wang LF
3256	Intragastric fish bones migrate into the liver: A case report
	Dai MG, Zheng JJ, Yang J, Ye B
3261	Primary seminal vesicle adenocarcinoma with a history of seminal vesicle cyst: A case report and review of literature
	Yao Y, Liu S, He YL, Luo L, Zhang GM
3267	Immune checkpoint inhibitor therapy-induced autoimmune polyendocrine syndrome type II and Crohn's disease: A case report
	Gao MJ, Xu Y, Wang WB
3275	Late-onset mitochondrial encephalomyopathy with lactic acidosis and stroke-like episodes syndrome with mitochondrial DNA 3243A>G mutation masquerading as autoimmune encephalitis: A case report
	Wang JW, Yuan XB, Chen HF
3282	Metastatic gastric cancer from breast carcinoma presenting with paraneoplastic rheumatic syndrome: A case report
	Rech MB, da-Cruz ER, Salgado K, Balbinot RA, Balbinot SS, Soldera J
3288	Novel mutation of SPG4 gene in a Chinese family with hereditary spastic paraplegia: A case report
	Wang J, Bu WT, Zhu MJ, Tang JY, Liu XM
3295	Chronic pulmonary mucormycosis caused by rhizopus microsporus mimics lung carcinoma in an immunocompetent adult: A case report
	Guo XZ, Gong LH, Wang WX, Yang DS, Zhang BH, Zhou ZT, Yu XH



<b>O t</b>	World Journal of Clinical Cases
Conten	Thrice Monthly Volume 11 Number 14 May 16, 2023
3304	Idiopathic sclerosing mesenteritis presenting with small bowel volvulus in a patient with antiphospholipid syndrome: A case report
	Chennavasin P, Gururatsakul M
3311	Neisseria mucosa - A rare cause of peritoneal dialysis-related peritonitis: A case report
	Ren JM, Zhang XY, Liu SY
3317	Rectal prolapse in a 30-year-old bladder stone male patient: A case report
	Ding HX, Huang JG, Feng C, Tai SC
3323	Successful treatment of veno-arterial extracorporeal membrane oxygenation complicated with left ventricular thrombus by intravenous thrombolysis: A case report
	Wang YD, Lin JF, Huang XY, Han XD
3330	Successful remimazolam sedation-epidural block in an older patient with severe chronic obstructive pulmonary disease: A case report
	Yu JJ, Pei HS, Meng Y
3340	<i>De novo</i> mutation of <i>NAXE</i> ( <i>APOAIBP</i> )-related early-onset progressive encephalopathy with brain edema and/or leukoencephalopathy-1: A case report
	Ding L, Huang TT, Ying GH, Wang SY, Xu HF, Qian H, Rahman F, Lu XP, Guo H, Zheng G, Zhang G
3351	Iatrogenic atlantoaxial rotatory subluxation after thyroidectomy in a pediatric patient: A case report
	Hong WJ, Lee JK, Hong JH, Han MS, Lee SS
3356	Bladder metastasis from epidermal growth factor receptor mutant lung cancer: A case report
	Jin CB, Yang L
3362	Primary rectal mucosa-associated lymphoid tissue lymphoma treated with only endoscopic submucosal dissection: A case report
	Lee WS, Noh MG, Joo YE



### Contents

Thrice Monthly Volume 11 Number 14 May 16, 2023

#### **ABOUT COVER**

Editorial Board Member of World Journal of Clinical Cases, Jaw-Yuan Wang, MD, PhD, Professor, Surgical Oncologist, Department of Surgery, Kaohsiung Medical University Hospital, Kaohsiung Medical University, Kaohsiung 807, Taiwan. jawyuanwang@gmail.com

### **AIMS AND SCOPE**

The primary aim of World Journal of Clinical Cases (WJCC, World J Clin Cases) is to provide scholars and readers from various fields of clinical medicine with a platform to publish high-quality clinical research articles and communicate their research findings online.

WJCC mainly publishes articles reporting research results and findings obtained in the field of clinical medicine and covering a wide range of topics, including case control studies, retrospective cohort studies, retrospective studies, clinical trials studies, observational studies, prospective studies, randomized controlled trials, randomized clinical trials, systematic reviews, meta-analysis, and case reports.

#### **INDEXING/ABSTRACTING**

The WJCC is now abstracted and indexed in Science Citation Index Expanded (SCIE, also known as SciSearch®), Journal Citation Reports/Science Edition, Current Contents®/Clinical Medicine, PubMed, PubMed Central, Scopus, Reference Citation Analysis, China National Knowledge Infrastructure, China Science and Technology Journal Database, and Superstar Journals Database. The 2022 Edition of Journal Citation Reports® cites the 2021 impact factor (IF) for WJCC as 1.534; IF without journal self cites: 1.491; 5-year IF: 1.599; Journal Citation Indicator: 0.28; Ranking: 135 among 172 journals in medicine, general and internal; and Quartile category: Q4. The WJCC's CiteScore for 2021 is 1.2 and Scopus CiteScore rank 2021: General Medicine is 443/826.

### **RESPONSIBLE EDITORS FOR THIS ISSUE**

Production Editor: Hua-Ge Yu; Production Department Director: Xu Guo; Editorial Office Director: Jin-Lei Wang.

NAME OF JOURNAL World Journal of Clinical Cases	INSTRUCTIONS TO AUTHORS https://www.wignet.com/bpg/gerinfo/204
<b>ISSN</b> ISSN 2307-8960 (online)	GUIDELINES FOR ETHICS DOCUMENTS https://www.wignet.com/bpg/GerInfo/287
LAUNCH DATE April 16, 2013	GUIDELINES FOR NON-NATIVE SPEAKERS OF ENGLISH https://www.wignet.com/bpg/gerinfo/240
FREQUENCY Thrice Monthly	PUBLICATION ETHICS https://www.wignet.com/bpg/GerInfo/288
EDITORS-IN-CHIEF	PUBLICATION MISCONDUCT
Bao-Gan Peng, Jerzy Tadeusz Chudek, George Kontogeorgos, Maurizio Serati, Ja Hyeon Ku	https://www.wjgnet.com/bpg/gerinfo/208
EDITORIAL BOARD MEMBERS	ARTICLE PROCESSING CHARGE
https://www.wjgnet.com/2307-8960/editorialboard.htm	https://www.wjgnet.com/bpg/gerinfo/242
PUBLICATION DATE	STEPS FOR SUBMITTING MANUSCRIPTS
May 16, 2023	https://www.wjgnet.com/bpg/GerInfo/239
COPYRIGHT	ONLINE SUBMISSION
© 2023 Baishideng Publishing Group Inc	https://www.f6publishing.com

© 2023 Baishideng Publishing Group Inc. All rights reserved. 7041 Koll Center Parkway, Suite 160, Pleasanton, CA 94566, USA E-mail: bpgoffice@wjgnet.com https://www.wjgnet.com



W J C C World Journal of Clinical Cases

Submit a Manuscript: https://www.f6publishing.com

World J Clin Cases 2023 May 16; 11(14): 3238-3247

DOI: 10.12998/wjcc.v11.i14.3238

ISSN 2307-8960 (online)

ORIGINAL ARTICLE

#### **Randomized Controlled Trial**

# Effect of non-pharmacological treatment on the full recovery of social functioning in patients with attention deficit hyperactivity disorder

Ying-Bo Lv, Wei Cheng, Meng-Hui Wang, Xiao-Min Wang, Yan-Li Hu, Lan-Qiu Lv

Specialty type: Psychology

#### Provenance and peer review:

Unsolicited article; Externally peer reviewed.

Peer-review model: Single blind

#### Peer-review report's scientific quality classification

Grade A (Excellent): 0 Grade B (Very good): B Grade C (Good): C Grade D (Fair): 0 Grade E (Poor): 0

P-Reviewer: Joshipura KJ, United States; Sanz M, Spain

Received: March 2, 2023 Peer-review started: March 2, 2023 First decision: March 14, 2023 Revised: March 25, 2023 Accepted: April 7, 2023 Article in press: April 7, 2023 Published online: May 16, 2023



Ying-Bo Lv, Wei Cheng, Meng-Hui Wang, Xiao-Min Wang, Yan-Li Hu, Lan-Qiu Lv, Pediatric Health Care Section, Ningbo Women and Children's Hospital, Ningbo 315000, Zhejiang Province, China

Corresponding author: Lan-Qiu Lv, MD, Attending Doctor, Pediatric Health Care Section, Ningbo Women and Children's Hospital, No. 339 Liuting Street, Haishu District, Ningbo 315000, Zhejiang Province, China. lulanqiunb@sina.com

## Abstract

#### BACKGROUND

Long-term treatment of attention deficit/hyperactivity disorder (ADHD) is associated with adverse events, such as nausea and vomiting, dizziness, and sleep disturbances, and poor maintenance of late ADHD medication compromises treatment outcomes and prolongs the recovery of patients' social functioning.

#### AIM

To evaluate the effect of non-pharmacological treatment on the full recovery of social functioning in patients with ADHD.

#### **METHODS**

A total of 90 patients diagnosed with ADHD between May 2019 and August 2020 were included in the study and randomly assigned to either the pharmacological group (methylphenidate hydrochloride and tomoxetine hydrochloride) or the non-pharmacological group (parental training, behavior modification, sensory integration therapy, and sand tray therapy), with 45 cases in each group. Outcome measures included treatment compliance, Swanson, Nolan, and Pelham, Version IV (SNAP-IV) scores, Conners Parent Symptom Questionnaire (PSQ) scores, and Weiss Functional Impairment Rating Scale (WFIRS) scores.

#### RESULTS

The non-pharmacological interventions resulted in significantly higher compliance in patients (95.56%) compared with medication (71.11%) (P < 0.05). However, no significant differences in SNAP-IV and PSQ scores, in addition to the learning/school, social activities, and adventure activities of the WFIRS scores were observed between the two groups (P > 0.05). Patients with non-pharmacological interventions showed higher WFIRS scores for family, daily life skills, and



self-concept than those in the pharmacological group (P < 0.05).

#### **CONCLUSION**

Non-pharmacological interventions, in contrast to the potential risks of adverse events after longterm medication, improve patient treatment compliance, alleviate patients' behavioral symptoms of attention, impulsivity, and hyperactivity, and improve their cognitive ability, thereby improving family relationships and patient self-evaluation.

Key Words: Non-pharmacological treatment; Attention deficit hyperactivity disorder; Social functioning; Recovery; Weiss Functional Impairment Rating Scale scores

©The Author(s) 2023. Published by Baishideng Publishing Group Inc. All rights reserved.

**Core Tip:** This study evaluated the effect of non-pharmacological treatments on the full recovery of social functioning in patients with attention deficit hyperactivity disorder (ADHD). A total of 90 patients with ADHD were included in this study. The non-pharmacological intervention resulted in significantly higher patient compliance than the pharmacological treatment group. Patients in the non-pharmacological intervention group also had significantly higher Weiss Functional Impairment Rating Scale scores on family, daily living skills and self-concept than those receiving medication. Thus, non-pharmacological interventions had a positive impact on the overall recovery of social functioning in ADHD patients compared to long-term pharmacological treatment.

Citation: Lv YB, Cheng W, Wang MH, Wang XM, Hu YL, Lv LQ. Effect of non-pharmacological treatment on the full recovery of social functioning in patients with attention deficit hyperactivity disorder. World J Clin Cases 2023; 11(14): 3238-3247

URL: https://www.wjgnet.com/2307-8960/full/v11/i14/3238.htm DOI: https://dx.doi.org/10.12998/wjcc.v11.i14.3238

#### INTRODUCTION

Attention deficit hyperactivity disorder (ADHD) is the most common chronic neurodevelopmental disorder in childhood and adolescence[1,2], characterized by age-inappropriate distractibility, reduced attention span, overactivity, and emotional impulsivity regardless of the setting, and associated cognitive impairment, and learning difficulties. The age of onset of ADHD is uncertain, with a prevalence of 6-14 years and a peak of 8-10 years, with a significant gender difference of [3-8]: 1 between males and females. The prevalence of the disease also varies widely, ranging from 1.3% to 13.4%, with a mean prevalence of 3%. According to reports on ADHD, approximately 70% of children with ADHD develop symptoms into adolescence and are at significantly greater risk for disruptive behaviors and emotions than during childhood. The global prevalence of ADHD has been growing in the past decades [1]. Clinical studies reveal that symptoms continue in 70% of ADHD children throughout adolescence and 30% into adults, creating disturbances in family connections[9]. ADHD is a chronic condition with lifelong consequences, often associated with poor academic performance[10], emotional instability[3], and behavioral problems[4].

Studies have shown that long-term treatment of ADHD is associated with adverse events<sup>[5]</sup>, such as nausea, vomiting, dizziness, and sleep disturbances, and poor maintenance of late ADHD medication compromises treatment outcomes and prolongs the recovery of patients' social functioning[11]. The American Academy of Child and Adolescent Psychiatry advises pharmaceutical therapies for ADHD above behavioral treatment alone[6], although the American Academy of Pediatrics favors behavioral interventions, particularly for preschool-aged children[7,12]. However, controversy persists regarding the optimal intervention paradigm for ADHD.

Currently, the treatment of ADHD is mainly pharmacological. Long-term drug use is prone to insomnia, loss of appetite, headache, abdominal pain, or risk of drug dependence. Moreover, pharmacological treatment has a negligible effect on children's deficits in social functioning caused by environmental and psychological factors. In addition to pharmacological treatment, non-pharmacological treatments such as biofeedback therapy, family therapy, and combined medical-educational interventions also constitute an important part of disease management, with more targeted correction of children's behavioral disorders, more lasting and stable improvement of social functions, and higher safety compared with pharmacological treatment. In traditional Chinese medicine (TCM), the disease mainly involves the kidney and the liver, and according to clinical observation, comorbidities from the two organs represent most of the evidence. Kidney deficiency is more prominent among the symptoms



of ADHD. TCM focuses on the combination of disease and evidence and coordinates the functions of the internal organs, resulting in significant therapeutic effects.

Thus, exploring the therapeutic effects of non-pharmacological interventions is of great value and long-term significance for patients with ADHD and their families. Therefore, this study investigated the effect of non-pharmacological treatment on the full recovery of social functioning in patients with ADHD.

#### MATERIALS AND METHODS

#### Data source

Children diagnosed with ADHD between May 2019 and August 2020 were assessed for eligibility. After excluding eight cases with discontinued treatment due to adverse events, two cases that revoked their consent, and two cases whose parents refused follow-up visits, 90 ADHD patients (73 males and 17 females, aged 6-18 years) were included in the study. The included patients were randomly assigned to receive either methylphenidate hydrochloride and tomoxetine hydrochloride (pharmacological group) or parental training, behavior modification, sensory integration therapy, and sand tray therapy (nonpharmacological group), with 45 cases in each group.

The randomization was conducted using an online web-based randomization tool (freely available at http://www.randomizer.org/). For concealment of allocation, the randomization procedure and assignment were managed by an independent research assistant who was not involved in the screening or evaluation of the participants.

The original sample size calculation estimated that 45 patients would be needed in each group to detect a 3-point difference between groups using a 2-sided significance test with a power of 0.8 and an alpha error level of 0.05.

The normality of the sample was determined with the Shapiro-Wilk test. Exploratory analyses of descriptive statistical data were performed using the Tukey test. Inferential statistical analysis of quantitative mean data (PES/WES, ISQ, and B.L) was performed using the non-parametric Wilcoxon-Mann-Whitney U-test.

All included patients, or their guardians, provided written informed consent. The study was approved by the ethics committee of our hospital.

#### Inclusion and exclusion criteria

All patients met the diagnostic criteria established in the Diagnostic and Statistical Manual of Mental Disorders, 5th edition, regardless of sex. Patients with mental retardation, character disorder, mood disorder, tic disorder, childhood autism, and schizophrenia were excluded.

#### Clinical characteristics

Hyperactivity and impulsivity: Hyperactivity and impulsivity mostly manifest in early childhood and become obvious in elementary school. The children are hyperactive, talk much in class, and often fight with their classmates.

Inattention: Children have difficulty concentrating when learning and easily respond to external stimuli

Learning difficulties: Children with normal intelligence have learning difficulties due to the abovementioned symptoms, and some develop cognitive impairment and general analysis disorder. The onset of the disease is mostly observed before the age of seven and lasts for more than six months.

Dissonant personality or behavioral disorders: Children are more capricious, stubborn, impulsive, lack self-control, and uncomfortable with social interaction. A small number of cases still have personality and behavioral defects in adulthood.

#### Treatment methods

The pharmacological group received methylphenidate hydrochloride extended-release tablets and tomoxetine hydrochloride tablets.

The non-pharmacological group received parent training, behavior modification, sensory integration therapy, and sand tray therapy. Parent training involved four sessions, including disease awareness, pros and cons of drugs, parent-child relationship, and methods to improve attention span. Behavior modification involved two major courses covering the positive reinforcement method of behavior modification, temporary isolation method, fading method, demonstration method, cognitive behavior therapy, and applied behavior analysis.

In addition, sensory integration therapy involved 45-60 min of training per session over 3-6 mo. The training process included warm-up, vestibular sensory, proprioception, balance, hand-eye coordination, sedation, and fine motor. Warm-up included simple children's songs, finger rhymes, action songs, and



story reading. Vestibular sensor entailed rotating, moving, scooting, and crossing obstacles. Their proprioception was stimulated using "heavy work" activities, such as holding, lifting, pushing, and pulling heavy objects. Balance involved crossing the balance beam, walking along the hula hoop, one-legged games, and balance board. Furthermore, hand-eye coordination included throwing balloons and sandbags, hitting target objects, and throwing objects to partners. Sedation included relaxation activities, simple children's songs, finger rhymes, and storytelling. Finally, fine motor included making snacks, scrapbooking activities, art activities, and coloring.

Sand tray therapy involved 10–12 sessions with a unified sand tray therapist, with each session lasting about 40–60 min. The process includes instructional language, familiarization with the environment, feeling the sand, creating a sand tray, playing with the sand tray, dialogue and communication, dismantling the work, and discussion and analysis with parents. All programs were scheduled after school or during weekends for the children. All patients were followed up once a month for the first six months of drug administration and every 2–3 mo thereafter.

The two groups received Tiaoshen Yizhi Decoction consisting of 20 g of *Paeoniae Radix Alba*, 20 g of *Rehmanniae Radix Praeparata*, 6 g of *Acori Tatarinowii Rhizoma*, 20 g of *Angelicae Sinensis Radix*, 10 g of *Lycii Fructus*, 15 g of *Ostreae Concha*, 10 g of *Testudinis Carapax et Plastrum*, 15 g of *Os Draconis*, 6 g of *Polygalae Radix*, 15 g of *Margarita*, 10 g of *Uncariae Ramulus cum Uncis*, and 6 g of *liquorice root*, grounded into powder and one dose was administered daily. The powder was dissolved in 60 mL of boiling water, and half a dose was administered in the morning and in the evening.

#### Outcome measures

The compliance scale designed by our hospital was used to assess the patient's treatment compliance before the parental intervention. Compliance was rated by the parents using a 3-point scale: full compliance, partial compliance, and non-compliance.

In addition, the Swanson, Nolan, and Pelham, Version IV (SNAP-IV) was used for rating, which contains nine inattention problems and nine hyperactivity/impulsivity problems, each with a 4-point scale indicating different levels of severity.

The Conners Parent Symptom Questionnaire (PSQ) was used. The questionnaire has 48 items and includes six factors: character problems, learning problems, psychosomatic problems, impulsivity-hyperactivity, anxiety, and hyperactivity index. The higher the factor score, the more severe the problem.

Finally, the Weiss Functional Impairment Rating Scale (WFIRS) was used to assess patients' social functioning. The scale contains 50 items rated by parents on six domains of family, learning/school, life skills, self-concept, social activities, and adventure activities, each on a scale of 0–4, respectively. The lower the score, the better the social functioning

#### Statistical analysis

If the parameter beta is either a difference of means, a log odds ratio, or a log hazard ratio, then it is reasonable to assume that the beta is unbiased and normally distributed. GraphPad Prism 8 was used for image processing, and SPSS 26.0 software was used to organize the data and for statistical analysis. Measurement data were expressed as mean  $\pm$  SD and analyzed using the *t*-test. Count data were expressed as a rate (%) and compared using the  $\chi^2$  test. *P* < 0.05 indicated that differences were statistically significant.

#### RESULTS

#### Patient characteristics

There were 35 males and 10 females in the pharmacological group, aged 6–18 (9.03 ± 1.78) years, with a duration of disease of 11–35 (23.89 ± 1.77) months and Wechsler IQ scores  $\geq$  85 (90.23 ± 2.91). In total, 39 cases attended school in urban areas, while 6 cases were schooled in non-urban areas. By contrast, there were 38 males and 7 females in the pharmacological group, aged 6–18 (9.23 ± 1.65) years, with a duration of illness of 11-37 (23.48 ± 2.13) months and Wechsler IQ scores  $\geq$  85 (90.37 ± 2.75). A total of 36 cases attended school in urban areas and 9 cases in non-urban areas. The characteristics of patients in the two groups were comparable (P > 0.05) (Table 1).

#### Treatment compliance

Before treatment, there was no significant difference in compliance between the two groups (P > 0.05). After treatment, 43 (95.56%) of the 45 patients in the non-pharmacological group were compliant. The number of partially compliant patients in the non-pharmacological group increased to 22 (48.89%), while the number of non-compliant patients decreased to 2 (4.44%), indicating that non-pharmacological treatment can improve patients' treatment compliance compared with pharmacological intervention, and the difference was statistically significant (P < 0.05) (Table 2).

Zaishidena® WJCC | https://www.wjgnet.com

#### Lv YB et al. Effect of non-drug therapy on ADHD

Table 1 Patient characteristics					
Characteristics	Pharmaceutical group ( $n = 45$ )	Non-pharmaceutical group (n = 45)	t value	P value	
Sex, n (%)			0.653	0.419	
Male	35 (77.78)	38 (84.45)			
Female	10 (22.22)	7 (15.55)			
Age (yr, mean)	6-18 (9.03 ± 1.78)	6-18 (9.23 ± 1.65)	0.553	0.582	
Duration of disease (mo, mean)	11-35 (23.89 ± 1.77)	11-37 (23.48 ± 2.13)	0.993	0.323	
Wechsler intelligence score (score, mean)	≥ 85 (90.23 ± 2.91)	≥ 85 (90.37 ± 2.75)	0.235	0.815	
School, <i>n</i> (%)			0.720	0.396	
Urban	39 (86.67)	36 (80.00)			
Non-urban	6 (13.33)	9 (20.00)			

Table 2 Treatment compliance, n (%)					
Treatment compliance	Pharmaceutical group ( <i>n</i> = 45)	Non-pharmaceutical group ( <i>n</i> = 45)	X <sup>2</sup>	P value	
Before treatment			0.179	0.673	
Complete compliance	1 (2.22)	2 (4.44)			
Partial compliance	21 (46.67)	18 (40.00)			
Non-compliance	23 (51.11)	25 (55.55)			
Compliance	22 (48.89)	20 (44.44)			
After treatment					
Complete compliance	12 (26.67)	21 (46.67)	15.195	< 0.001	
Partial compliance	20 (44.44)	22 (48.89)			
Non-compliance	13 (28.89)	2 (4.44)			
Compliance	32 (71.11)	43 (95.56)			

#### **SNAP-IV**

There was no statistically significant difference in SNAP-IV scores between the two groups of patients before treatment (P > 0.05). SNAP-IV scores were significantly reduced in both groups after treatment; the reduction was greater in the non-pharmacological intervention group (16.85  $\pm$  2.48) than in the pharmacological group (17.69 ± 2.28). However, the differences in SNAP-IV scores between the two groups before and after treatment were not statistically significant (P > 0.05) (Figure 1).

#### Conners parenting inventory

After treatment, patients in the pharmacological group had character problems scores of  $0.65 \pm 0.11$ , learning problems scores of  $0.88 \pm 0.25$ , psychosomatic problems scores of  $3.35 \pm 1.05$ , impulsivityhyperactivity scores of 1.02  $\pm$  0.51, anxiety scores of 1.98  $\pm$  1.21, and hyperactivity index scores of 0.85  $\pm$ 0.36. Patients in the non-pharmacological group had character problems scores of  $(0.61 \pm 0.08)$ , learning problems 0.81 ± 0.11, psychosomatic problems scores of 3.21 ± 0.77, impulsivity-hyperactivity scores of  $0.87 \pm 0.12$ , anxiety scores of  $1.87 \pm 0.89$ , and hyperactivity scores of  $0.79 \pm 0.35$ . There were no significant differences in Conners' PSQ scores between the two groups (P > 0.05) (Table 3).

#### Social functioning

After treatment, the family score in the pharmacological group ( $0.78 \pm 0.52$ ) and non-pharmacological group ( $0.46 \pm 0.41$ ) were statistically significantly different (t = 3.242, P < 0.05). There was a statistically significant difference (t = 5.335, P < 0.05) in the life skills score of the pharmacological group ( $0.99 \pm 0.38$ ) and the non-pharmacological group  $(0.69 \pm 0.31)$ . In addition, the difference in the self-concept score of the pharmacological group  $(0.95 \pm 0.42)$  and the non-pharmacological group  $(0.65 \pm 0.35)$  was statistically significant (t = 3.436, P < 0.05). However, there was no significant difference in Weiss scores between the two groups in learning/school, social activities, and risk-taking activities (P > 0.05). Family, life skills, and self-concept scores indicated that the non-pharmacological treatment group scored significantly better than the pharmacotherapy group (Table 4).



Table 3 Conners scores (mean ± SD)					
	Pharmaceutical group ( <i>n</i> = 45)	Non-pharmaceutical group (n = 45)	t value	P value	
Character problems	$0.65 \pm 0.11$	$0.61 \pm 0.08$	1.973	0.052	
Learning problems	$0.88 \pm 0.25$	$0.81 \pm 0.11$	1.719	0.089	
Psychosomatic problems	$3.35 \pm 1.05$	3.21 ± 0.77	0.721	0.437	
Impulsivity-hyperactivity	$1.02\pm0.51$	$0.87 \pm 0.12$	1.921	0.058	
Anxiety	$1.98 \pm 1.21$	$1.87 \pm 0.89$	0.491	0.625	
Hyperactivity index	$0.85 \pm 0.36$	$0.79 \pm 0.35$	0.802	0.425	

#### Table 4 Weiss scores (mean ± SD)

	Pharmaceutical group (n = 45)	Non-pharmaceutical group (n = 45)	t value	P value
Family	$0.78 \pm 0.52$	$0.46 \pm 0.41$	3.242	0.002
Learning/school	$0.52 \pm 0.44$	$0.42 \pm 0.35$	1.193	0.236
Life skills	$0.99 \pm 0.38$	$0.69 \pm 0.31$	5.335	< 0.001
Self-concept	$0.95 \pm 0.42$	$0.65 \pm 0.35$	3.436	0.001
Social activities	$0.45\pm0.51$	$0.32 \pm 0.21$	1.581	0.117
Adventure activities	$0.22 \pm 0.21$	$0.20 \pm 0.18$	0.485	0.629



Figure 1 Swanson, Nolan, and Pelham, Version IV scores.

#### DISCUSSION

ADHD is a psychiatric disorder with significant symptoms that interfere with the child's daily life and learning. ADHD belongs to the categories of "forgetfulness", "deafness", and "false annoyance" in TCM. Approximately 20%-25% of children with ADHD exhibit symptoms that continue into adulthood. Early intervention once symptoms are detected is important to alleviate the symptoms and reduce the impairment of social functioning of the patients[13]. Pharmacotherapy is an effective treatment for ADHD that significantly relieves the core symptoms but is associated with adverse effects [7,14]. The efficacy of non-pharmacological interventions is more stable and efficient than pharmacological treatment.

This study found that the compliance of the non-pharmaceutical group (95.56%) was significantly higher than that of the pharmaceutical group (71.11%) after treatment. This difference in compliance suggests that, after treatment, patients in the non-pharmacological group, could follow their parents' advice and actively cooperate with the treatment, leading to significant improvements in their emotions and physical discomfort. Moreover, there were no significant differences in the SNAP-IV scores, PSQ scores, and the learning/school, social activities, and adventure activities of the WFIRS scores between the two groups. Patients who received non-pharmacological therapies had higher WFIRS ratings for family, life skills, and self-concept than those who received drugs. Compared to pharmaceutical therapies, non-pharmacological interventions provide more enrichment for children's competence development, family connection improvement, and self-evaluation. This finding contrasts with previous research findings, which may be attributed to personalized variances from the small sample size of this study.



In this study, non-pharmacological interventions included parent training, behavior modification, sensory integration therapy, and sand tray therapy. Parenting training is an established treatment for children with ADHD, whereas behavioral therapies focus on the functional recovery of ADHD children. Prior research has demonstrated marked symptom alleviation with parenting treatments for ADHD, and this correlation remains significant when trials with concurrent medication are omitted [15]. Different parenting behavior training programs have been proven effective, such as the 3P Positive Parenting Program[16], the New Forest Parenting Program[17], and the Barkley Program[18]. Some patients with ADHD experience perceptual problems and hand-eye coordination deficits. Research has shown that sensory-motor training could correct hyperactivity, impulsivity, and attention deficit, and adjust the vestibular response deficits of sensory integration disorders (vestibular balance organs form vague images and effects in the brain), tactile defenses (frontal cortical sensory acuity and control difficulties), use disorders in children (blurred body image formed in the brain by proprioceptive kinesthesia and vestibular balance organs, causing clumsy coordination of the five senses, especially hand-eye coordination and reading and writing difficulties), visuospatial shape perception disorder (the same principle as the use disorder, coupled with a lack of dexterity in the visual cortical coordination of the brain), and gravitational insecurity, which were reflected in the results of the current study.

In addition, sand tray therapy improves the ADHD hyperactivity index, character problems, hyperactivity/impulsivity, and anxiety in children. The reason may be that during the sand tray game, children with ADHD focus on the sand in their hands under the supervision of the host tester, which may gradually waken the children's inner ability of self-healing and self-development, effectively reducing anxiety and improving their emotional stability. The succession of sandbox imagery portrayed in the sandbox produces a continual dialogue between the sandbox player's conscious and unconscious thoughts, resulting in an effective improvement of character issues closely associated with personality. The combination of the above non-pharmacological interventions promotes emotional stability, gradually improves former interpersonal sensitivity, personality paranoia, and impulsive and aggressive behavior, and enhances self-esteem and self-confidence, which in turn facilitate the full recovery of patients' social functioning.

According to TCM theory, normal life activities and the mental state of the human body is a state of calmness of Yin and Yang[19,20]. Children's internal organs are delicate and vigorous and frequently suffer from a relative lack of essence, blood, fluid, and other material bases, predisposing them to a state of excitement, hyperactivity, impatience, irritability, and mental disturbance[21,22]. In TCM, treatment for ADHD lies on nourishing Yin and Yang, tonifying the liver, and benefiting the kidneys, for which Tiaoshen Yizhi Decoction was developed in our hospital for ADHD patients. The combination of drugs in the decoction focused on nourishing Yin and Yang without harming the spleen, and Angelicae Sinensis Radix was used to invigorate the blood. In addition, some herbs with sedative functions were used to prevent drowsiness in the children. Moreover, calm and warm medicines were used to avoid damage to the spleen and stomach, and protection of the righteous energy was emphasized in the use of drugs[23].

Our study has some limitations. The diversity of non-pharmacologic treatments increases the difficulty in controlling the operational criteria. Assessing the effectiveness of such non-pharmacologic treatments is closely related to the complexity of the intervention and the influence of different operational providers. Despite our rigorous control of the means and measures of non-pharmacological interventions, uncontrollable variation in the implementation of the treatment may still occur. Second, non-pharmacologic treatment alone is slow and time-consuming, so it is crucial that non-pharmacologic treatment be combined with appropriate pharmacologic treatments. The primary outcome measure in this study was based on self-reported symptoms and function. Therefore, we lacked more objective clinician-based measures. In addition, the sample size of ADAH patients in the study was not large enough, and future studies could be based on a larger sample size. Finally, long-term follow-up information could also be included in future studies to assess the long-term effects of non-pharmacological treatments on patients with ADAH.

No precise etiology or pathogenesis of ADHD has been identified to date, and a growing body of data suggests that the syndrome involves a combination of biopsychosocial factors. Therefore, combining pharmacological and non-pharmacological treatments is more appreciated for the biopsychosocial model. Compared with foreign studies, research in this area in China is still in its initial stage, and there are many problems to be addressed. Therefore, further promotion of research on the non-pharmacological treatment of ADHD in China is required.

#### CONCLUSION

In contrast to the potential risks of adverse events after long-term medication, non-pharmacological interventions improve the treatment compliance of patients, alleviate patients' behavioral symptoms of attention, impulsivity, and hyperactivity, and improve patients' cognitive ability, thereby enhancing family relationships and patient self-evaluation.



#### **ARTICLE HIGHLIGHTS**

#### Research background

Long-term treatment of attention-deficit hyperactivity disorder (ADHD) is associated with adverse events. Therefore, non-pharmacological treatment has attracted a lot of attention as a new treatment, but its impact on the full recovery of social functioning of ADHD patients is still unknown.

#### Research motivation

Clarifying the effects of non-pharmacological treatments on the social functioning of ADAH patients is of great value and long-term significance to ADHD patients and their families.

#### Research objectives

This study aimed to investigate the impact of non-pharmacological treatments on the full recovery of social functioning in patients with ADHD.

#### Research methods

A total of 90 patients diagnosed with ADHD were enrolled in the study and randomly assigned to either the pharmacological group or the non-pharmacological group, with 45 cases in each group. Treatment adherence, Swanson, Nolan and Pelham Fourth Edition (SNAP-IV) scores, Connors Parental Symptom Questionnaire (PSQ) scores, and Weil Functional Impairment Rating Scale (WFIRS) scores were measured.

#### Research results

Non-pharmacological interventions resulted in significantly higher compliance in patients compared to pharmacological intervention. No significant differences in the SNAP-IV scores, PSQ scores, and the learning/school, social activities, and adventure activities of the WFIRS scores were observed between the two groups. Patients in the non-pharmacological group showed higher WFIRS scores of family, daily life skills, and self-concept compared to those in the pharmacological group.

#### **Research conclusions**

In contrast to the potential risks of adverse events after long-term medication, non-pharmacological interventions improve patient treatment compliance, alleviate patients' behavioral symptoms of attention, impulsivity, and hyperactivity, and improve their cognitive ability, thereby improving family relationships and patient self-evaluation.

#### Research perspectives

This study demonstrates the positive impact of non-pharmacological treatment compared to long-term medication on the full recovery of social functioning in patients with ADAH.

#### FOOTNOTES

Author contributions: Lv YB and Cheng W proposed concepts for this study; Wang MH and Lv LQ collected data; Lv YB, Wang XM, and Hu YL contributed to formal analysis; Lv YB and Lv LQ contributed to the survey; Lv YB, Lv LQ, and Wang XM contributed to this method; Lv YB, Lv LQ, Hu YL, and Wang MH supervised the study; Lv LQ validated this study; Lv YB and Cheng W contributed to the visualization of research; Lv YB and Lv LQ initially drafted this manuscript; Lv YB, Cheng W, Wang MH, and Wang XM reviewed and edited the manuscript.

Supported by Ningbo Science and Technology Plan Project Public Welfare Plan (Municipal Level), No: 2019C50099; Ningbo Medical Key Supporting Discipline Child Health Science, No: 2022-F26.

Institutional review board statement: The study was approved by the Ethics Committee of Ningbo Women and Children's Hospital.

Clinical trial registration statement: This study is registered at https://www.researchregistry.com/browse-theregistry#home/registrationdetails/63f45f91a0f7f7002b4f0915/.

Informed consent statement: All study participants, or their legal guardian, provided informed written consent.

**Conflict-of-interest statement:** We declare that there are no conflicts of interest.

Data sharing statement: No additional data are available.

CONSORT 2010 statement: The authors have read the CONSORT 2010 statement, and the manuscript was prepared and revised according to the CONSORT 2010 statement.



Open-Access: This article is an open-access article that was selected by an in-house editor and fully peer-reviewed by external reviewers. It is distributed in accordance with the Creative Commons Attribution NonCommercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited and the use is noncommercial. See: https://creativecommons.org/Licenses/by-nc/4.0/

#### Country/Territory of origin: China

**ORCID number:** Ying-Bo Lv 0009-0007-7692-8664; Lan-Qiu Lv 0009-0008-1814-0833.

S-Editor: Wang JL L-Editor: A P-Editor: Yuan YY

#### REFERENCES

- Polanczyk GV, Willcutt EG, Salum GA, Kieling C, Rohde LA. ADHD prevalence estimates across three decades: an updated systematic review and meta-regression analysis. Int J Epidemiol 2014; 43: 434-442 [PMID: 24464188 DOI: 10.1093/ije/dyt261]
- Drechsler R, Brem S, Brandeis D, Grünblatt E, Berger G, Walitza S. ADHD: Current Concepts and Treatments in 2 Children and Adolescents. Neuropediatrics 2020; 51: 315-335 [PMID: 32559806 DOI: 10.1055/s-0040-1701658]
- Sobanski E, Banaschewski T, Asherson P, Buitelaar J, Chen W, Franke B, Holtmann M, Krumm B, Sergeant J, Sonuga-3 Barke E, Stringaris A, Taylor E, Anney R, Ebstein RP, Gill M, Miranda A, Mulas F, Oades RD, Roeyers H, Rothenberger A, Steinhausen HC, Faraone SV. Emotional lability in children and adolescents with attention deficit/hyperactivity disorder (ADHD): clinical correlates and familial prevalence. J Child Psychol Psychiatry 2010; 51: 915-923 [PMID: 20132417 DOI: 10.1111/j.1469-7610.2010.02217.x]
- Biederman J, Petty CR, Dolan C, Hughes S, Mick E, Monuteaux MC, Faraone SV. The long-term longitudinal course of 4 oppositional defiant disorder and conduct disorder in ADHD boys: findings from a controlled 10-year prospective longitudinal follow-up study. Psychol Med 2008; 38: 1027-1036 [PMID: 18205967 DOI: 10.1017/S0033291707002668]
- 5 Sonuga-Barke EJ, Brandeis D, Cortese S, Daley D, Ferrin M, Holtmann M, Stevenson J, Danckaerts M, van der Oord S, Döpfner M, Dittmann RW, Simonoff E, Zuddas A, Banaschewski T, Buitelaar J, Coghill D, Hollis C, Konofal E, Lecendreux M, Wong IC, Sergeant J; European ADHD Guidelines Group. Nonpharmacological interventions for ADHD: systematic review and meta-analyses of randomized controlled trials of dietary and psychological treatments. Am J Psychiatry 2013; 170: 275-289 [PMID: 23360949 DOI: 10.1176/appi.ajp.2012.12070991]
- Pliszka S; AACAP Work Group on Quality Issues. Practice parameter for the assessment and treatment of children and 6 adolescents with attention-deficit/hyperactivity disorder. J Am Acad Child Adolesc Psychiatry 2007; 46: 894-921 [PMID: 17581453 DOI: 10.1097/chi.0b013e318054e724]
- Subcommittee on Attention-Deficit/Hyperactivity Disorder; Steering Committee on Quality Improvement and Management, Wolraich M, Brown L, Brown RT, DuPaul G, Earls M, Feldman HM, Ganiats TG, Kaplanek B, Meyer B, Perrin J, Pierce K, Reiff M, Stein MT, Visser S. ADHD: clinical practice guideline for the diagnosis, evaluation, and treatment of attention-deficit/hyperactivity disorder in children and adolescents. Pediatrics 2011; 128: 1007-1022 [PMID: 22003063 DOI: 10.1542/peds.2011-2654]
- National Collaborating Centre for Mental Health (UK). Attention Deficit Hyperactivity Disorder: Diagnosis and Management of ADHD in Children, Young People and Adults. Leicester (UK): British Psychological Society (UK), 2009 [PMID: 22420012]
- Harpin VA. The effect of ADHD on the life of an individual, their family, and community from preschool to adult life. 0 Arch Dis Child 2005; 90 Suppl 1: i2-i7 [PMID: 15665153 DOI: 10.1136/adc.2004.059006]
- Biederman J, Petty CR, Fried R, Kaiser R, Dolan CR, Schoenfeld S, Doyle AE, Seidman LJ, Faraone SV. Educational 10 and occupational underattainment in adults with attention-deficit/hyperactivity disorder: a controlled study. J Clin Psychiatry 2008; 69: 1217-1222 [PMID: 18681752 DOI: 10.4088/jcp.v69n0803]
- Miklós M, Futó J, Komáromy D, Balázs J. Executive Function and Attention Performance in Children with ADHD: 11 Effects of Medication and Comparison with Typically Developing Children. Int J Environ Res Public Health 2019; 16 [PMID: 31658722 DOI: 10.3390/ijerph16203822]
- Sprich SE, Safren SA, Finkelstein D, Remmert JE, Hammerness P. A randomized controlled trial of cognitive behavioral 12 therapy for ADHD in medication-treated adolescents. J Child Psychol Psychiatry 2016; 57: 1218-1226 [PMID: 26990084 DOI: 10.1111/jcpp.12549]
- 13 Felt BT, Biermann B, Christner JG, Kochhar P, Harrison RV. Diagnosis and management of ADHD in children. Am Fam Physician 2014; 90: 456-464 [PMID: 25369623]
- Manos MJ, Giuliano K, Geyer E. ADHD: Overdiagnosed and overtreated, or misdiagnosed and mistreated? Cleve Clin J 14 Med 2017; 84: 873-880 [PMID: 29173249 DOI: 10.3949/ccjm.84a.15051]
- 15 Coates J, Taylor JA, Sayal K. Parenting Interventions for ADHD: A Systematic Literature Review and Meta-Analysis. J Atten Disord 2015; 19: 831-843 [PMID: 24915737 DOI: 10.1177/1087054714535952]
- Sanders MR. Triple P-Positive Parenting Program: towards an empirically validated multilevel parenting and family 16 support strategy for the prevention of behavior and emotional problems in children. Clin Child Fam Psychol Rev 1999; 2: 71-90 [PMID: 11225933 DOI: 10.1023/a:1021843613840]



- Sonuga-Barke EJ, Daley D, Thompson M, Laver-Bradbury C, Weeks A. Parent-based therapies for preschool attention-17 deficit/hyperactivity disorder: a randomized, controlled trial with a community sample. J Am Acad Child Adolesc *Psychiatry* 2001; **40**: 402-408 [PMID: 11314565 DOI: 10.1097/00004583-200104000-00008]
- 18 Storebø OJ, Elmose Andersen M, Skoog M, Joost Hansen S, Simonsen E, Pedersen N, Tendal B, Callesen HE, Faltinsen E, Gluud C. Social skills training for attention deficit hyperactivity disorder (ADHD) in children aged 5 to 18 years. Cochrane Database Syst Rev 2019; 6: CD008223 [PMID: 31222721 DOI: 10.1002/14651858.CD008223.pub3]
- 19 Schein J, Childress A, Gagnon-Sanschagrin P, Maitland J, Bedard J, Cloutier M, Guérin A. Treatment Patterns Among Patients with Attention-Deficit/Hyperactivity Disorder and Comorbid Anxiety and/or Depression in the United States: A Retrospective Claims Analysis. Adv Ther 2023 [PMID: 36913128 DOI: 10.1007/s12325-023-02458-5]
- Fu D, Guo HL, Hu YH, Chen F. A precision medication study of atomoxetine in children with attention deficit 20 hyperactivity disorder: CYP2D6 genetic testing and therapeutic drug monitoring. Zhongguo Dang Dai Er Ke Za Zhi 2023; 25: 98-103 [PMID: 36655671 DOI: 10.7499/j.issn.1008-8830.2208092]
- 21 Çelik HEA, Küçükgöncü S, Erdoğan A, Özerdem A. Response Inhibition and Interference Control in Adult Attention Deficit Hyperactivity Disorder. Noro Psikiyatr Ars 2023; 60: 3-8 [PMID: 36911564 DOI: 10.29399/npa.28192]
- Chen SC, Cheng HL, Han LF, Wu GT, Zhang RY, Suen LK, Chen X, Yeung WF. Parent-administered pediatric tuina for 22 the treatment of attention deficit hyperactivity disorder symptoms: Process evaluation of a pilot randomized controlled trial. Complement Ther Med 2022; 70: 102854 [PMID: 35842070 DOI: 10.1016/j.ctim.2022.102854]
- Pauli-Pott U, Skoluda N, Nater UM, Becker K, Derz F, Kaspar E, Kaspar Z, Kehm K, Kött M, Mann C, Schurek P, 23 Pott W, Schloß S. Long-term cortisol secretion in attention deficit hyperactivity disorder: roles of sex, comorbidity, and symptom presentation. Eur Child Adolesc Psychiatry 2023 [PMID: 36917355 DOI: 10.1007/s00787-023-02180-1]





## Published by Baishideng Publishing Group Inc 7041 Koll Center Parkway, Suite 160, Pleasanton, CA 94566, USA Telephone: +1-925-3991568 E-mail: bpgoffice@wjgnet.com Help Desk: https://www.f6publishing.com/helpdesk https://www.wjgnet.com

