

RESEARCH

Open Access



Electroconvulsive therapy for adolescents with severe depressive episode and suicidality: retrospective comparison between responders and non-responders

Hao Ren^{1,2}, Xinglian Wang¹, Zheng Zhang¹, Xiufen Zhong^{1,3}, Qinghua Luo¹, Haitang Qiu^{1*} and Yan Huang^{4*}

Abstract

Background For adolescents with major depression who exhibit suicidal tendencies, Electroconvulsive Therapy (ECT) is increasingly adopted in clinical practice. Yet, the precise mechanisms behind its effectiveness remain elusive, and studies on factors that influence treatment outcomes are scarce.

Methods In this retrospective comparative study, we included all adolescent severe depressive episode patients with suicidal tendencies admitted to the Psychiatry Department of the First Affiliated Hospital of Chongqing Medical University between 2017 and 2021 and received ECT treatment. By collecting data on personal history, medical history, and standard treatment features, we established demographic, disease, medication, and ECT treatment factors variables. Patients were divided into effective and ineffective groups based on the Clinical Global Impressions-Improvement (CGI-I) scale scores, and differences between outcomes were compared. Logistic regression analyses were used to identify factors independently associated with ineffectiveness.

Results A total of 494 adolescent severe depressive episode patients with suicidal behavior who received ECT were included in this study. According to CGI-I scores, the treatment was effective in 361 patients (73.1%) and ineffective in 133 patients (26.9%). Logistic regression analyses showed that 8 to 12 and 12 to 16 ECT sessions reduced the risk of ineffectiveness compared to fewer than 4 sessions. The risk of ineffectiveness decreased with age and increased with comorbidity with obsessive-compulsive disorder (OCD). Compared to sertraline, escitalopram was associated with a heightened risk of futility, whereas olanzapine and aripiprazole demonstrated a reduced risk when contrasted with quetiapine.

Conclusions ECT's ineffectiveness in treating adolescent severe depressive episode with suicidal behavior decreases with age, and comorbidity with OCD significantly increases the risk of treatment failure. Fewer than 8 ECT sessions may hinder achieving satisfactory results.

Keywords Electroconvulsive therapy, Adolescent, Depression, Suicide, Retrospective study

*Correspondence:

Haitang Qiu
qiuhaitang2008@hotmail.com
Yan Huang
huangyan1011@163.com

Full list of author information is available at the end of the article



© The Author(s) 2024. **Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit <http://creativecommons.org/licenses/by/4.0/>. The Creative Commons Public Domain Dedication waiver (<http://creativecommons.org/publicdomain/zero/1.0/>) applies to the data made available in this article, unless otherwise stated in a credit line to the data.

Background

Adolescents around the globe are increasingly at risk of major depression, with the current two diagnostic standards providing specific definitions (major depressive disorder in the Diagnostic and Statistical Manual of Mental Disorders, severe depressive episode in the International Classification of Diseases). What is clear is that major depression can significantly disrupt their lives and impede social functioning. Alarming, the global prevalence may reach up to 8% [1], with the possibility of even higher rates among American adolescents [2]. In China, which boasts a larger population, the estimated prevalence stands at 2% [3]—a figure that still represents a considerable number. Currently, depressive disorders have risen to rank as the fourth leading cause of the global burden of disease [4].

As major depression has become more pervasive, the risk of suicide has correspondingly increased, posing a grave threat to life and health [5]. Notably, suicide ranks as the leading cause of death among teenagers [6]. Consequently, psychiatrists are intensively working to develop rapid and effective treatments for depressive patients exhibiting suicidality.

ECT is the major treatment approach for adult depressive disorders with acute suicidal tendencies. This method is capable of significantly reducing both symptoms of depression and the risk of suicide [7], and its use among teenagers has been consistently growing. Recent retrospective research from various countries has shown that the response rates to ECT for adolescent major depression range from 72 to 78.6% [8–11]. However, approximately one-third of patients fail to respond to this therapy. It is unclear what factors influence the clinical response to this therapy [9, 12].

In reviewing previous research on factors that influence adult ECT outcomes, older age has been identified as a demographic factor positively affecting treatment success [13]. Concurrently, factors such as accompanying psychotic symptoms [13], extended periods of depression, baseline medical treatment failure [14], suicidal ideation [15], and co-existing personality disorders [16] may be disease factor linked to treatment effectiveness. Regarding medical treatment aspects, medications that lower the threshold for seizures or shorten their duration can reduce ECT's effectiveness [17]. Consequently, benzodiazepines (BDZs) and antiepileptic drugs are often considered the main factors affecting ECT's outcomes [18], with a general consensus to discontinue these medicines before therapy to prevent any impact on efficacy [19].

Research on adolescents is notably scarce, and the few existing studies indicate that among patients with depression undergoing ECT, features accompanying psychotic symptoms are correlated with higher relief rates

[20]. Additionally, female major depressive cases with nonsuicidal self-injury (NSSI) [21], as well as those with fewer ECT sessions [11], may correlate with unsatisfactory outcomes, but many more contributing factors await clarification.

Recently, the First Affiliated Hospital of Chongqing Medical University (CQMU) has amassed a significant number of adolescent ECT cases. We intend to conduct a retrospective analysis on the ECT of depressive disorders with suicidal tendencies in teenagers, aiming to identify differences between successful and unsuccessful cases, thereby providing support for prognosis assessment and therapeutic guidance. We collected data on personal history, medical history, and standard treatment features to establish variables related to demographics, medical conditions, medications, and ECT treatment factors. Based on previous research, it is hypothesized that variables such as gender, disease duration, psychotic symptoms, suicidal behaviors, medication dosages, and ECT sessions may influence treatment effectiveness.

Methods

This study is a retrospective comparative research effort in which all patient names have been anonymized. Since the anonymized data cannot reveal the identity information of the patients, informed consent is not required. The Human Research and Ethics Committee of CQMU had approved this study (NO: 2022-K525).

Design

The study utilizes the electronic medical record system of CQMU to extract data according to predefined variables. It then investigates the relationship with ECT effectiveness using logistic regression model analyses, controlling for confounding variables.

Participants

All participants were psychiatric inpatients 18 years old or younger at CQMU, with discharge dates between December 31, 2016, and June 30, 2021 (n = 2231).

Inclusion criteria

(1) Receiving ECT. (2) Meeting the diagnostic standards in ICD-10, such as F32.2 (Severe depressive episode without psychotic symptoms) and F32.3 (Severe depressive episode with psychotic symptoms). (3) Clear suicidal intention or attempt, as indicated by the Columbia Suicide Severity Rating Scale (C-SSRS).

Exclusion criteria

(1) Diagnoses including F00-F09 (Organic mental disorders), F10-F19 (Mental and behavioral disorders resulting from psychoactive substance use), F20-F29

(Schizophrenia), F30–F31.9 (Manic episode, Bipolar affective disorder), F70–F79 (Mental retardation). (2) Absence of Clinical Global Impressions–Improvement scale (CGI-I) score.

Grouping standards

Employing the CGI-I score at discharge for evaluating overall efficacy as the basis for division, and first defining overall efficacy as: (1) Positive response: CGI-I score 1 (very much improved) or 2 (much improved), (2) Negative response: 3 (Minimally improved), 4 (No change).

Patients were then categorized into effective and ineffective groups according to the CGI-I positive or negative response, and other variable data were extracted from medical records. The procedure is illustrated in Fig. 1.

Variables and measures

Demographic variables

Age, gender, family history of psychiatric illness, family history of suicidal deaths. Family history is confined to two families and three generations, with only relatives definitively diagnosed with psychiatric illness assessed as positive.

Disease variables

Disease duration, psychotic symptoms, suicide attempts within the last month, hospitalizations at CQMU, hospitalizations at other psychiatric hospitals, Hamilton Depression Rating Scale (HAMD) score, Hamilton Anxiety Scale (HAMA) score, psychiatric comorbidities. Disease duration is the time from initial depression

symptoms to current admission. The evaluation of psychotic symptoms is limited to whether they were present during the current hospital stay. Suicide attempts within the last month refer to the actual number of intentional actions taken to end one’s life but survived in the month prior to admission [22]. HAMD and HAMA scores are ratings at the time of admission.

Medication treatment variables

Antidepressants, doses of antidepressants, fluoxetine equivalent dose, antipsychotics, doses of antipsychotics, olanzapine equivalent dose, mood stabilizers, sedative-hypnotics and other anti-anxiety drugs. Medication treatment encompasses the entire inpatient treatment period, besides mood stabilizers, they are prohibited from use during the ECT sessions period and are only prescribed in the discharge prescription after the ECTs. All medication treatment variables are selected at the time of discharge, and the units are consistently measured in mg. The doses of antidepressants and antipsychotics are converted to fluoxetine equivalent dose [23] and olanzapine equivalent dose [24, 25], respectively, to ensure comparability. The transformed variables were not entered into the statistical analysis.

Variables of ECT treatment

ECT sessions, severe side effects of ECT, previous ECT treatment history. ECT sessions were provided for the duration of the current hospitalization, without maintenance ECTs. Severe side effects of ECT are defined as those that substantially affect function, manifest clear signs upon physical examination, and necessitate intervention.

Other variables

Days hospitalized.

Electroconvulsive therapy

All patients and their lawful guardians voluntarily accepted electroconvulsive therapy after being fully informed about the risks and benefits. The lawful guardians then completed the signing of the informed consent document. Prior to treatment, all physicians in the patient’s medical team conducted a pre-treatment discussion to evaluate the indications and contraindications for ECT. The patient retains the right to demand the cessation of electroconvulsive therapy at any time.

All ECT sessions were modified and conducted using the Thymatron DGx system (SomaticsLLC, Lake Bluff, IL, USA), following a brief pulse pattern with bitemporal electrode placement. The initial electrical dose was determined by the formula age * 0.7, with subsequent electrical volumes titrated by 5% in response to seizures [26].

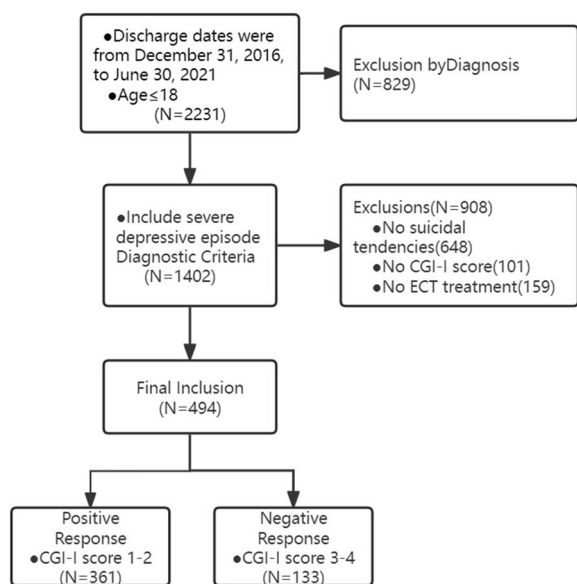


Fig. 1 Patient selection

The standard treatment frequency is two or three times a week. However, if urgent needs arise or a patient gives informed consent, the first week will have four ECT sessions, then revert to the standard frequency. Anesthesia and muscle relaxation were induced using propofol (1–1.5 mg/kg) and succinylcholine chloride (0.5–1 mg/kg), with atropine administered to regulate heart rate when required.

Statistical analyses

Data analysis was conducted using IBM SPSS Statistics 25.0. Categorical variables were described using percentages (%), and continuous variables were represented by either Mean (SD) or Median (IQR), based on conformity with normal distribution. Comparisons between groups were made using Chi-square tests, independent samples t-tests, or Mann–Whitney U tests. The outcome, CGI-I positive or negative response, was utilized as the dependent variable, with 0 for a positive response and 1 for a negative response. Logistic regression analyses were employed to assess factors associated with ineffectiveness in ECT treatment for adolescents with severe depressive episode and suicidal behavior, calculating Odds Ratios (OR) and 95% Confidence Intervals (95% CI). Due to the absence of equivalent dose conversion formulas for duloxetine and vortioxetine, their missing values were imputed using the median fluoxetine equivalent dose.

Various models were applied:

The unadjusted model assessed four variables concerning ECT treatment and days hospitalized.

Model 1 evaluated ECT treatment and demographic factors, excluding severe side effects of ECT, and days hospitalized, and included 6 variables.

Model 2 added disease variables to Model 1, excluding the same variables as in Model 1, involving 14 variables.

Model 3 combined Models 1 and 2 with medication treatment variables, excluding mood stabilizers and those omitted in Models 1 and 2, included 20 variables.

All models classified ECT sessions instances in intervals of 4 as ordinal rank variables and encoded them as nominal categorical dummy variables to specifically study their correlation with the dependent variable (details in Table 3). Subgroup analysis divided age at the median into a dichotomous variable, and in conjunction with gender, partitioned the data into four distinct stratified subsets. Each subset was analyzed according to Model 3’s parameters to validate the result stability. Statistical findings were deemed significant with $P < 0.05$ (2-tailed). Regarding omitted variables, days hospitalized and severe side effects of ECT may act as mediating variables, potentially influenced by ECT sessions.

Results

In this study, 494 adolescent patients with severe depressive episode and suicidal behavior underwent ECT treatment; 361 (73.1%) of them responded effectively. The distribution of the CGI-I score can be found in Fig. 2.

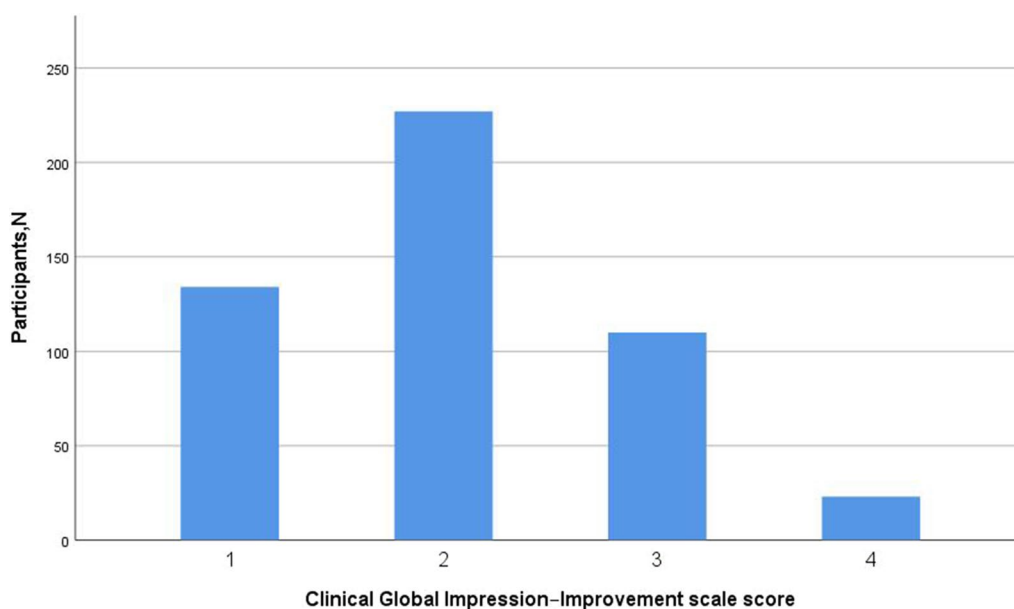


Fig. 2 Clinical Global Impression-Improvement scale score distribution

Demographic factors

410 (83.0%) of the participants were females, and the median age was 15, with an interquartile range (IQR) of 14–17 and a total range of 11–18. Age, gender, family history of psychiatric illness, and family history of suicidal deaths demonstrated no statistically significant differences between the groups, as detailed in Table 1.

Disease factors

With a median duration of 12 months (IQR: 10–24, range: 1–96), revealed a statistical difference between the effective and ineffective groups [12(8–24) vs. 24(12–24), $P=0.037$]. Comparisons for psychotic symptoms, suicide attempts within the last month, hospitalizations at CQMU, hospitalizations at other psychiatric hospitals, HAMD Score, HAMA Score, and psychiatric comorbidities all demonstrated no statistically significant differences, as detailed in Table 1.

Medication treatment factors

493 participants (99.8%) used antidepressants, and 471 (95.3%) used antipsychotics, with only the use of antipsychotics showing a statistical significance between the two groups ($P=0.008$). Comparisons between the two groups in other variables, including antidepressants, mood stabilizers, sedative-hypnotics, other anti-anxiety drugs, fluoxetine equivalent dose, and olanzapine equivalent dose, revealed no statistically significant differences. Refer to Table 1.

ECT treatment factors

The median of ECT sessions (IQR, range) was 10 (8–12, 1–18), and among the variables related to ECT treatment, only the ECT sessions demonstrated statistical significance between the two groups [10 (8–11) vs. 9 (6–12), $P=0.010$]. Of the participants, 41 (8.3%) had previously undergone ECT, and 36 (7.3%) experienced severe side effects of ECT, including slowness of thought (17, 3.4%), oblivion (13, 2.6%), headache (4, 0.8%), emesis (1, 0.2%), and haziness (1, 0.2%). Refer to Table 1.

Other factors

The median of days hospitalized was 21 (IQR 17–24, range 3–40), and a statistically significant difference was observed between the two groups [21(17–25) vs. 19(14–24)], with $P=0.018$. Refer to Table 1.

Factors associated with negative response

Unadjusted logistic regression analyses revealed that undergoing ECT sessions $8 \leq n < 12$, or $12 \leq n < 16$, reduced the risk of nonresponse compared to fewer

than 4 ($8 \leq n < 12$: OR=0.08, 95% CI 0.02–0.32, $P=0.001$; $12 \leq n < 16$: OR=0.13, 95% CI 0.03–0.68, $P=0.015$). Models 1 and 2 yielded similar results for ECT treatment variables, and Model 2 further revealed that as age increases, the risk of non-response decreases (OR=0.85, 95% CI 0.74–0.98, $P=0.026$). Additionally, comorbid OCD increased the risk of non-response (OR=9.46, 95% CI 1.50–59.65, $P=0.017$). Model 3 supported Models 1 and 2's ECT treatment variable results, including a decreased risk of non-response as age increases (OR=0.80, 95% CI 0.68–0.94, $P=0.005$), increased risk with comorbid OCD (OR=10.86, 95% CI 1.65–71.43, $P=0.013$), and medication treatment findings (Escitalopram: OR=3.76, 95% CI 1.21–11.72, $P=0.023$; Olanzapine: OR=0.44, 95% CI 0.24–0.80, $P=0.006$; Aripiprazole: OR=0.35, 95% CI 0.18–0.67, $P=0.002$). Refer to Table 2.

Factors associated with negative response in subgroup analysis

In the subgroup analysis conducted using Model 3, all four examined strata provided evidence that 8 to 12 sessions of ECT can reduce the risk of treatment inefficacy. This was observed in the following subgroups:

Female: OR=0.09, 95% CI 0.02–0.49, $P=0.006$
 Male: OR=0.00, 95% CI 0.00–0.08, $P=0.006$
 Age ≤ 15 : OR=0.12, 95% CI 0.01–0.83, $P=0.032$
 Age > 15 : OR=0.01, 95% CI 0.00–0.09, $P=0.001$

Supplementary to the primary findings, in the female stratum, disease duration may increase the risk of inefficacy (OR=1.02, 95% CI 1.00–1.03, $P=0.049$). In the age ≤ 15 stratum, suicide attempts within the last month was found to be positively associated with an increased risk of inefficacy (OR=2.03, 95% CI 1.09–3.79, $P=0.025$). Conversely, in the age > 15 stratum, an increase in HAMD Score may indicate a reduced risk of inefficacy (OR=0.92, 95% CI 0.85–0.99, $P=0.033$). The OR values for the remaining results were consistent with the direction of the original findings. For a detailed overview, refer to Table 3.

Discussion

In this retrospective clinical study, conducted over the past 5 years at the Department of Psychiatry in CQMU, an exhaustive analysis was performed on adolescents with severe depressive episode and suicidal tendencies who underwent ECT. The research utilized a substantial dataset to distinguish between effective and ineffective treatment outcomes. Consistent results were noted across various statistical models, and the reliability of these findings was substantiated through targeted

Table 1 Compare the differences between responders and non-responders to ECT treatment in terms of demographic factors, disease factors, medication treatment factors, ECT treatment factors, and other factors

Variables	Total ECT (N = 494)	Positive response in ECT (n = 361)	Negative response in ECT (n = 133)	Statistics		
				Chi square test (X)	M-W U test (Z)	P
Demographic						
Gender				0.415		0.520
Female	410 (83.0%)	302 (83.7%)	108 (81.2%)			
Male	84 (17.0%)	59 (16.3%)	25 (18.8%)			
Age (year)	15 (14,17)	15 (14,17)	15 (13,17)		-0.464	0.643
Family history of psychiatric illness						
Positive	39 (7.9%)	26 (7.2%)	13 (9.8%)	0.884		0.347
Negative	455 (92.1%)	335 (92.8%)	120 (90.2%)			
Family history of suicidal deaths						
positive	18 (3.6%)	11 (3.0%)	7 (5.3%)	1.359		0.244
Negative	476 (96.4%)	350 (97.0%)	126 (94.7%)			
Disease						
Disease duration (months)	12 (10,24)	12 (8,24)	24 (12,24)		-2.088	0.037
Psychotic symptoms	116 (23.5%)	83 (23.0%)	33 (24.8%)	0.179		0.672
Suicide attempts within the last month (n)	0 (0,0)	0 (0,0)	0 (0,0)		-0.553	0.574
Hospitalizations (n)						
CQMU	1 (1,1)	1 (1,1)	1 (1,1)		-1.376	0.142
Other psychiatric hospitals	0 (0,0)	0 (0,0)	0 (0,0)		-1.174	0.258
HAMD Score	33.15 ± 7.39	33.29 ± 7.28	32.75 ± 7.70			0.470
HAMA Score	21.64 ± 7.20	21.64 ± 7.05	21.64 ± 7.63			0.996
Psychiatric comorbidities						
OCD	14 (2.8%)	8 (2.2%)	6 (4.5%)	9.798		0.081
Posttraumatic stress disorder (PTSD)	5 (1.0%)	2 (0.6%)	3 (2.3%)			
Eating disorder (ED)	3 (0.6%)	3 (0.8%)	0			
Attention deficit hyperactivity disorder (ADHD)	2 (0.4%)	2 (0.6%)	0			
Anxiety disorder (AD)	1 (0.2%)	0	1 (0.8%)			
Anxiety disorder (AD)	3 (0.6%)	1 (0.3%)	2 (1.5%)			
Medication treatment						
Antidepressants						
Sertraline	493 (99.8%)	360 (99.7%)	133 (100.0%)	12.308		0.254
Fluoxetine	298 (60.3%)	217 (60.3%)	81 (60.9%)			
Escitalopram	113 (22.9%)	88 (24.4%)	25 (18.8%)			
Venlafaxine	23 (4.7%)	15 (4.2%)	8 (6.0%)			
Paroxetine	34 (6.9%)	27 (7.5%)	7 (5.3%)			
Fluvoxamine	6 (1.2%)	3 (0.8%)	3 (2.3%)			
Bupropion	1 (0.2%)	0	1 (0.8%)			
Duloxetine	2 (0.4%)	1 (0.3%)	1 (0.8%)			
Citalopram	6 (1.2%)	3 (0.8%)	3 (2.3%)			
Vortioxetine	5 (1.0%)	4 (1.1%)	1 (0.8%)			
Vortioxetine	3 (0.6%)	1 (0.3%)	2 (1.5%)			
Mirtazapine	3 (0.6%)	1 (0.3%)	1 (0.8%)			
Antipsychotics						
Quetiapine	471 (95.3%)	345 (95.6%)	126 (94.7%)	17.948		0.008
Olanzapine	178 (36.0%)	116 (32.1%)	62 (46.6%)			
Aripiprazole	153 (31.0%)	120 (33.2%)	33 (24.8%)			
Amisulpride	128 (25.9%)	101 (28.0%)	27 (20.3%)			
Risperidone	1 (0.2%)	0	1 (0.8%)			
Paliperidone	5 (1.0%)	5 (1.4%)	0			
Paliperidone	5 (1.0%)	2 (0.6%)	3 (2.3%)			

Table 1 (continued)

Variables	Total ECT (N=494)	Positive response in ECT (n=361)	Negative response in ECT (n=133)	Statistics		
				Chi square test (X)	M-W U test (Z)	P
Ziprasidone	1 (0.2%)	1 (0.3%)	0	1.636		0.662
Mood stabilizers	15(3.0%)	10(2.8%)	5(3.8%)			
Lithium	5 (1.0%)	3 (0.8%)	2 (1.5%)			
Valproate	2 (0.4%)	2 (0.6%)	0	3.559		0.169
Lamotrigine	8 (1.6%)	5 (1.4%)	3 (2.3%)			
Sedative-hypnotics	66 (13.4%)	41 (11.4%)	25 (18.8%)			
BDZs	45 (9.1%)	30 (8.3%)	15 (11.3%)	0.251		0.888
Non-BDZs	19 (3.8%)	11 (3.0%)	8 (6.0%)			
Other anti-anxiety drugs	61 (12.3%)	43 (11.9%)	18 (13.5%)			
Tandospirone	40 (8.1%)	28 (7.8%)	12 (9.0%)	0.174	0.174	0.862
Buspirone	21 (4.3%)	15 (4.2%)	6 (4.5%)			
Fluoxetine equivalent dose (mg)	60 (40,80)	60 (40,80)	60 (40,80)			
Olanzapine equivalent dose (mg)	3.33 (2.50,6.67)	3.33 (2.50,6.67)	3.75 (1.64,7.50)	-0.202		0.840
ECT treatment						
ECT sessions (n)	10 (8,12)	10 (8,11)	9 (6,12)	7.666	2.568	0.010
Serious side effects of ECT	36 (7.3%)	31 (8.6%)	5 (3.8%)			
Slowness of thought	17 (3.4%)	15 (4.2%)	2 (1.5%)			
Oblivion	13 (2.6%)	11 (3.0%)	2 (1.5%)	0.52	0.52	0.471
Headache	4 (0.8%)	4 (1.1%)	0			
Emesis	1 (0.2%)	1 (0.3%)	0			
Haziness	1 (0.2%)	0	1 (0.8%)	2.374		0.018
Previous ECT treatment history	41 (8.3%)	28 (7.8%)	13 (9.8%)			
Other						
Days hospitalized	21 (17,24)	21 (17,25)	19 (14,24)			

subgroup analysis. This comprehensive examination encompassed non-responder characteristics, including demographic information, specific disease characteristics, medication interventions, and ECT treatment outcomes.

Within the scope of demographic variables, despite the narrow 7-year age range in the sample (11 to 18 years), a significant impact of age on treatment effectiveness emerged. This finding is in line with the consistent trend observed in prior meta-analyses [13], suggesting that the risk of ECT inefficacy decreases with age. Adolescence represents a crucial brain development phase characterized by dynamic alterations in brain structure and functionality [27]. A possible explanation for the varying ECT responses across different age groups could be the linear relationship between age and the degree of brain maturation. Yet, current scientific literature has not identified distinct brain differences in adolescents with depression across varying age groups [28]. As a result, some scholars argue that age might act as an intermediary factor connected with specific predictive symptoms, rather than directly impacting treatment effectiveness [29]. Further, other age-related aspects, such as advancing

school grades creating unique environmental stressors, evolving and stable personality characteristics, increased social engagement, tendencies toward substance use, and greater access to online information, may all serve as mediators between age and treatment success.

Concerning disease variables, the study found that the presence of comorbid OCD significantly increases the risk of ECT treatment ineffectiveness compared to those without any comorbidities. Furthermore, we then compared the variables between the two ineffective patients and the three effective patients, and no statistically significant difference was observed. These findings align with outcomes from other studies on ECT treatment for bipolar depression [30] and mania [31], which similarly reported diminished efficacy in patients suffering from depression with comorbid OCD. The exact cause for this phenomenon remains unclear.

In investigating medication treatment factors, the study found notable differences among various medication. Specifically, when compared to sertraline, escitalopram was associated with an increased risk of ECT ineffectiveness. Conversely, olanzapine and aripiprazole were found to reduce the risk of ineffectiveness compared to

Table 2 Logistic regression analyses: factors associated with ineffectiveness in ECT treatment for adolescents with severe depressive episode and suicidal behavior

Variables	Unadjusted model	Model 1	Model 2	Model 3
ECT sessions (n)				
4 < n	1.00 (Ref)	1.00 (Ref)	1.00 (Ref)	1.00 (Ref)
4 ≤ n < 8				
8 ≤ n < 12	0.08 (0.02–0.32)**	0.09 (0.03–0.29)**	0.08 (0.02–0.29)**	0.05 (0.01–0.20)**
12 ≤ n < 16	0.13 (0.03–0.68)*	0.16 (0.05–0.56)**	0.17 (0.05–0.61)**	0.08 (0.02–0.34)**
16 ≤ n				
Age			0.85 (0.74–0.98)*	0.80 (0.68–0.94)**
Psychiatric comorbidities				
None			1.00 (Ref)	1.00 (Ref)
OCD			9.46 (1.50–59.65)*	10.86 (1.65–71.42)*
Antidepressants				
Sertraline				1.00 (Ref)
Escitalopram				3.76 (1.21–11.72)*
Antipsychotics				
Quetiapine				1.00 (Ref)
Olanzapine				0.44 (0.24–0.79)**
Aripiprazole				0.35 (0.18–0.67)**

Results presentation: OR (95% CI). *P < 0.05; **P < 0.01

Unadjusted model: Included 4 variables such as ECT sessions, Severe side effects of ECT, Previous ECT treatment history, and Days hospitalized

Model 1: Comprised 6 variables including ECT sessions, Previous ECT treatment history, Age, Gender, Family history of psychiatric illness, and Family history of suicidal deaths

Model 2: Built upon Model 1 with a total of 14 variables, adding Disease duration, Psychotic symptoms, Suicide attempts within the last month, Hospitalizations at CQMU, Hospitalizations at other psychiatric hospitals, HAMD Score, and HAMA Score, Psychiatric comorbidities

Model 3: Expanded Model 2 to include 20 variables by additionally incorporating Antidepressants, Fluoxetine equivalent dose, Antipsychotics, Olanzapine equivalent dose, Sedative-hypnotics, and Other anti-anxiety drugs

Note: Mediating variables, Severe side effects of ECT and Days hospitalized were excluded in Models 1, 2, and 3. Mood stabilizers were excluded in Model 3 because they did not participate in the entire the ECT sessions

quetiapine. These findings contrast with previous meta-analyses concerning medication treatments for adolescents. Two primary divergences were noted: (1) Previous studies have recommended escitalopram for treating major depressive disorder without an associated suicide risk [32]; (2) The efficacy of atypical antipsychotics (AAPs) in augmenting treatment-resistant depression appeared similar across different types, inconsistent with our findings [33]. It's essential to recognize that our medication data were obtained at the time of discharge, necessitating a careful interpretation of the results. Upon examining the data, we identified a subset of patients on escitalopram who had previously exhibited a poor response to sertraline. This prior non-responsiveness could have influenced our results. However, the reported findings specifically center on escitalopram, potentially constraining a more holistic understanding of the medication-related outcomes. Furthermore, our research noted the inclusion of mood stabilizers in the discharge prescriptions, which could be intended for managing suicidal ideation, treatment-resistant depression, and significant irritability. It is important to highlight that

while antiepileptic mood stabilizers are not recognized for their efficacy in alleviating suicidal tendencies [34], lithium salts may offer such benefits [35, 36]. This distinction draws attention to the need for more extensive research to verify the therapeutic potential of lithium in these contexts.

In the study's examination of variables related to ECT treatment, a consistent pattern emerged: the risk of treatment ineffectiveness declined with 8 to 16 ECT sessions, a finding reinforced across different models and corroborated by subgroup analysis. This consistent reduction in risk was particularly stable within the 8 to 12 session range. Though other retrospective studies conducted in different regions of China did not provide the overall average ECT sessions for direct comparison between effective and ineffective groups, they did observe similar disparities (e.g., the effective group averaged 7.4 sessions versus an ineffective group averaged 6.6 sessions, $P=0.046$) [11]. An Israeli retrospective study further emphasized this pattern, revealing that a significant number of patients only responded to treatment after more than 12 ECT

Table 3 Stratified by age and gender in model 3: analysis of factors associated with the ineffectiveness of ECT treatment adolescent severe depressive episode with suicidal behavior

Variables	Female	Male	Age ≤ 15	Age > 15
ECT sessions (n)				
4 < n	1.00 (Ref)	1.00 (Ref)	1.00 (Ref)	1.00 (Ref)
4 ≤ n < 8		0.00 (0.00–0.09)*		
8 ≤ n < 12	0.09 (0.02–0.49)**	0.00 (0.00–0.81)**	0.11 (0.01–0.83)*	0.01 (0.00–0.09)**
12 ≤ n < 16		0.00 (0.00–0.15)*		0.00 (0.00–0.07)**
16 ≤ n				0.00 (0.00–0.23)**
Age	0.76 (0.64–0.91)**			
Disease duration	1.02 (1.00–1.03)*			
Suicide attempts within the last month			2.03 (1.09–3.79)*	
HAMD Score				0.92 (0.85–0.99)*
Psychiatric comorbidities				
None	1.00 (Ref)			1.00 (Ref)
OCD	9.19 (1.11–76.20)*			60.61 (3.28–1121.44)**
Antidepressants				
Sertraline				1.00 (Ref)
Fluoxetine				5.01 (1.03–24.41)*
Escitalopram				21.29 (3.35–135.38)**
Venlafaxine				7.38 (1.69–32.34)**
Paroxetine				40.55 (1.70–965.04)*
Antipsychotics				
Quetiapine	1.00 (Ref)		1.00 (Ref)	1.00 (Ref)
Olanzapine	0.41 (0.21–0.81)*		0.38 (0.17–0.87)*	0.29 (0.09–0.94)*
Aripiprazole	0.29 (0.14–0.63)**		0.27 (0.10–0.71)**	

Results presentation: OR (95% CI). *P < 0.05; **P < 0.01

sessions [12]. The finding underscores the importance of an adequately extensive ECT regimen to circumvent potential false-negative outcomes [12]. From the conducted research, it is revealed that the effective range of electroconvulsive therapy sessions for adolescents with severe depressive episode exceeds the AACAP's recommended range of 10–12 sessions [37]. This insight stresses the need for ongoing assessment of treatment response in adolescent patients in clinical practice, as it enables the development of personalized treatment plans. It's worth noting, however, that ECT treatments involving fewer than 8 sessions might not produce satisfactory therapeutic outcomes.

Conclusions

In this retrospective comparative study, we discovered that the risk of ineffective ECT in treating adolescent severe depressive episode with suicidal tendencies diminishes with increasing age. Furthermore, comorbidity with OCD notably elevates the risk of treatment failure, and administering fewer than 8 ECT sessions may hinder the achievement of satisfactory outcomes.

Limitations

1. Unavailability of ECT Parameters: The study lacks detailed ECT parameters, such as electrode placement, dosage, type, and frequency, obtainable from the system. This limits the ability to replicate or standardize the treatment, constraining the generalizability of the findings.
2. Exclusion of Psychological Factors: The absence of stress and family relationship variables might overlook critical factors influencing ECT treatment's effectiveness in adolescents. This omission could lead to a partial understanding of the outcomes.
3. No Information on Previous Medication Usage: The study does not include data on prior medication usage, and it was unable to evaluate treatment-resistant depression, possibly hindering the accurate interpretation of current medication-related results and creating potential confounding effects.
4. Restrictions Due to Study Design: The retrospective study did not utilize the Structured Clinical Interview for Diagnosis, leading to potential inaccuracies in

comorbidity rates and limiting the interpretation of the results.

5. **Absence of Laboratory Testing:** Laboratory tests including biomarkers were not available for this study. Such an omission restricts our ability to fully interpret the available data, potentially missing out on vital insights into the physiological responses to treatment.
6. **Single Measurement Instrument:** The study used only the CGI-I for efficacy evaluation and grouping. The omission of tools like HAMD or the Suicide Severity Rating Scale may reduce outcome interpretation accuracy and clarity.

Abbreviations

ECT	Electroconvulsive therapy
NSSI	Nonsuicidal self-injury
AACAP	The American Academy of Child and Adolescent Psychiatry
C-SSRS	Columbia Suicide Severity Rating Scale
CGI-I	Clinical Global Impressions–Improvement scale
HAMD	Hamilton Depression Rating Scale
HAMA	Hamilton Anxiety Scale
CQMU	The First Affiliated Hospital of Chongqing Medical University
OCD	Obsessive–compulsive disorder
PTSD	Posttraumatic stress disorder
ED	Eating disorder
ADHD	Attention deficit hyperactivity disorder
AD	Anxiety disorder
AAPs	Atypical antipsychotics

Acknowledgements

Thanks to the two anonymous reviewers for their constructive comments on the previous manuscript.

Author contributions

HR had full access to all of the data in the study and took responsibility for the integrity of the data and accuracy of the data analysis. Concept and design: All authors. Acquisition, analysis, or interpretation of data: HR. Statistical analysis: HR and YH. Drafting of the manuscript: HR. Critical revision of the manuscript for important intellectual content: All authors. Supervision: HQ and YH. All authors have read and approved the final manuscript.

Funding

This study was supported by the National Natural Science Foundation of China (grant/award number: 81901373).

Availability of data and materials

The data-sets analyzed during this study are available from the corresponding author on reasonable request.

Declarations

Ethics approval and consent to participate

The study was approved by the first affiliated hospital of Chongqing medical university (No: 2022-K525).

Consent for publication

All authors declare that they consent for publication.

Competing interests

The authors declare that they have no competing interests.

Author details

¹The First Affiliated Hospital of Chongqing Medical University, Chongqing, China. ²Chongqing Changshou District, Mental Health Center, Chongqing, China. ³Chongqing Mental Health Center, Chongqing, China. ⁴Chongqing Tradit Chinese Medicine Hospital, Chongqing, China.

Received: 24 August 2023 Accepted: 29 December 2023

Published online: 20 January 2024

References

1. Shorey S, Ng ED, Wong CHJ. Global prevalence of depression and elevated depressive symptoms among adolescents: a systematic review and meta-analysis. *Br J Clin Psychol.* 2022;61(2):287–305. <https://doi.org/10.1111/bjc.12333>.
2. Daly M. Prevalence of Depression Among Adolescents in the U.S. From 2009 to 2019: analysis of trends by sex, race/ethnicity, and income. *J Adolesc Health.* 2022;70(3):496–9. <https://doi.org/10.1016/j.jadohealth.2021.08.026>.
3. Li FH, Cui YH, Li Y, Guo LT, et al. Prevalence of mental disorders in school children and adolescents in China: diagnostic data from detailed clinical assessments of 17,524 individuals. *J Child Psychol Psychiatry.* 2022;63(1):34–46. <https://doi.org/10.1111/jcpp.13445>.
4. GBD 2019 Diseases and Injuries Collaborators. Global burden of 369 diseases and injuries in 204 countries and territories, 1990–2019: a systematic analysis for the Global Burden of Disease Study 2019. *Lancet.* 2020;396(10258):1204–22. [https://doi.org/10.1016/S0140-6736\(20\)30925-9](https://doi.org/10.1016/S0140-6736(20)30925-9).
5. Serra G, De Crescenzo F, Maisto F, Galante JR, et al. Suicidal behavior in juvenile bipolar disorder and major depressive disorder patients: systematic review and meta-analysis. *J Affect Disord.* 2022;311:572–81. <https://doi.org/10.1016/j.jad.2022.05.063>.
6. World Health Organization. PREFACE In: Preventing suicide: a global imperative. WHO Western Pacific Region. 2014. P.03. ISBN 978-92-4-156477-9 (NLM classification: HV 6545)
7. Kaster TS, Blumberger DM, Gomes T, Sutradhar R, et al. Risk of suicide death following electroconvulsive therapy treatment for depression: a propensity score-weighted, retrospective cohort study in Canada. *Lancet Psychiatry.* 2022;9(6):435–46. [https://doi.org/10.1016/S2215-0366\(22\)00077-3](https://doi.org/10.1016/S2215-0366(22)00077-3).
8. Karayağmurlu A, Coşkun M, Elboğa G, Ghaziuddin N, et al. Efficacy and safety of electroconvulsive therapy in adolescents: a retrospective chart review study from Turkey. *J ECT.* 2020;36(1):54–9. <https://doi.org/10.1097/YCT.0000000000000602>.
9. Pierson MD, Mickey JD, Gilley BL, Weeks RH. Outcomes of youth treated with electroconvulsive therapy: a retrospective cohort study. *J Clin Psychiatry.* 2021;82(2):19m13164. <https://doi.org/10.4088/JCP.19m13164>.
10. Grover S, Raju V, Chakrabarti S, Sharma A, et al. Use of electroconvulsive therapy in adolescents: a retrospective study. *Indian J Psychol Med.* 2021;43(2):119–24. <https://doi.org/10.1177/0253717620956730>.
11. Si Q, Zhang XY, Lei JX, Chen CX, et al. Electroconvulsive therapy efficacy in adolescents with mental illness: a retrospective comparison. *Front Psychiatry.* 2022;13: 990660. <https://doi.org/10.3389/fpsy.2022.990660>.
12. Maoz H, Nitzan U, Goldwyn Y, Krieger I, et al. When can we predict the outcome of an electroconvulsive therapy course in adolescents? A Retrospective Study. *J ECT.* 2018;34(2):104–7. <https://doi.org/10.1097/YCT.0000000000000469>.
13. Van Diermen L, Van Den Amele S, Kamperman AM, Sabbe BCG, et al. Prediction of electroconvulsive therapy response and remission in major depression: meta-analysis. *Br J Psychiatry.* 2018;212(2):71–80. <https://doi.org/10.1192/bjp.2017.28>.
14. Haq AU, Sitzmann FA, Goldman LM, Maixner FD, et al. Response of depression to electroconvulsive therapy: a meta-analysis of clinical predictors. *J Clin Psychiatry.* 2015;76(10):1374–84. <https://doi.org/10.4088/JCP.14r09528>.
15. Sienaert P, Brus O, Lambrichts S, Lundberg J, et al. Suicidal ideation and ECT, ECT and suicidal ideation: a register study. *Acta Psychiatr Scand.* 2022;146(1):74–84. <https://doi.org/10.1111/acps.13425>.

16. Hein M, Mungo A, Loas G. Nonremission after electroconvulsive therapy in individuals with major depression: role of borderline personality disorder. *J ECT*. 2022;38(4):238–43. <https://doi.org/10.1097/YCT.0000000000000857>.
17. Sackeim HA, Prudic J, Devanand DP, Nobler MS, et al. A prospective, randomized, double-blind comparison of bilateral and right unilateral electroconvulsive therapy at different stimulus intensities. *Arch Gen Psychiatry*. 2000;57(5):425–34. <https://doi.org/10.1001/archpsyc.57.5.425>.
18. Brus O, Cao Y, Gustafsson E, Hultén M, et al. Self-assessed remission rates after electroconvulsive therapy of depressive disorders. *Eur Psychiatry*. 2017;45:154–60. <https://doi.org/10.1016/j.eurpsy.2017.06.015>.
19. Weiss A, Hussain S, Ng B, Sarma S, et al. Royal Australian and New Zealand College of Psychiatrists professional practice guidelines for the administration of electroconvulsive therapy. *Aust N Z J Psychiatry*. 2019;53(7):609–23. <https://doi.org/10.1177/0004867419839139>.
20. Rask O, Nordenskjöld A, Johansson BA, Rad PM. Electroconvulsive therapy in children and adolescents: results from a population-based study utilizing the Swedish National Quality Register. *Eur Child Adolesc Psychiatry*. 2022. <https://doi.org/10.1007/s00787-022-02123-2>.
21. Rootes-Murdy K, Carlucci M, Tibbs M, Wachtel LE, et al. Non-suicidal self-injury and electroconvulsive therapy: outcomes in adolescent and young adult populations. *J Affect Disord*. 2019;250:94–8. <https://doi.org/10.1016/j.jad.2019.02.057>.
22. World Health Organization. Introduction In: Preventing suicide: a global imperative. WHO Western Pacific Region. 2014. p. 12
23. Hayasaka Y, Purgato M, Magni LR, et al. Dose equivalents of antidepressants: Evidence-based recommendations from randomized controlled trials. *J Affect Disord*. 2015;180:179–84. <https://doi.org/10.1016/j.jad.2015.03.021>.
24. Leucht S, Samara M, Heres S, Davis JM. Dose Equivalents for Antipsychotics: The DDD Method. *Schizophr Bull*. 2016;42(Suppl 1):S90–94. <https://doi.org/10.1093/schbul/sbv167>.
25. WHOC-ATC/DDD Index. WHO collaborative center for drug statistics methodology. https://www.whocc.no/atc_ddd_index/. Accessed 12 Feb 2023.
26. Kranaster L, Hoyer C, Janke C, Sartorius A. Bispectral index monitoring and seizure quality optimization in electroconvulsive therapy. *Pharmacopsychiatry*. 2013;46(4):147–50. <https://doi.org/10.1055/s-0032-1331748>.
27. Giedd NJ, Raznahan A, Alexander-Bloch A, Schmitt E, et al. Child psychiatry branch of the National Institute of Mental Health longitudinal structural magnetic resonance imaging study of human brain development. *Neuropsychopharmacology*. 2015;40(1):43–9. <https://doi.org/10.1038/npp.2014.236>.
28. Cullen KR, Westlund KM, Klimes-Dougan B, Mueller BA, et al. Abnormal amygdala resting-state functional connectivity in adolescent depression. *JAMA Psychiat*. 2014;71(10):1138–47. <https://doi.org/10.1001/jamapsychiatry.2014.1087>.
29. Heijnen WT CJ, Kamperman MA, Tjokrodipo LD, Hoogendijk WJG, et al. Influence of age on ECT efficacy in depression and the mediating role of psychomotor retardation and psychotic features. *J Psychiatr Res*. 2019;109:41–7. <https://doi.org/10.1016/j.jpsychires.2018.11.014>.
30. Popiolek K, Bejerot S, Brus O, Hammar A, et al. Electroconvulsive therapy in bipolar depression—effectiveness and prognostic factors. *Acta Psychiatr Scand*. 2019;140(3):196–204. <https://doi.org/10.1111/acps.13075>.
31. Popiolek K, Bejerot S, Landén M, Nordenskjöld A. Association of Clinical and demographic characteristics with response to electroconvulsive therapy in Mania. *JAMA Netw Open*. 2022;5(6): e2218330. <https://doi.org/10.1001/jamanetworkopen.2022.18330>.
32. Hetrick SE, McKenzie JE, Bailey AP, Sharma V, et al. New generation antidepressants for depression in children and adolescents: a network meta-analysis. *Cochrane Database Syst Rev*. 2021;5(5): CD013674. <https://doi.org/10.1002/14651858.CD013674.pub2>.
33. Zhou X, Keitner GI, Qin B, Ravindran AV, et al. Atypical antipsychotic augmentation for treatment-resistant depression: a systematic review and network meta-analysis. *Int J Neuropsychopharmacol*. 2015;18(11):pyv060. <https://doi.org/10.1093/ijnp/pyv060>.
34. Wilkinson ST, et al. Pharmacological and somatic treatment effects on suicide in adults: a systematic review and meta-analysis. *Depression Anxiety*. 2021;39:100–12. <https://doi.org/10.1002/da.23222>.
35. Del Matto L, Muscas M, Murru A, et al. Lithium and suicide prevention in mood disorders and in the general population: a systematic review. *Neurosci Biobehav Rev*. 2020;116:142–53. <https://doi.org/10.1016/j.neubiorev.2020.06.017>.
36. D’Anci KE, et al. Treatments for the prevention and management of suicide. *Ann Intern Med*. 2019;171:334. <https://doi.org/10.7326/M19-0869>.
37. Ghaziuddin N, Kutcher SP, Knapp P, Bernet W, et al. Practice parameter for use of electroconvulsive therapy with adolescents. *J Am Acad Child Adolesc Psychiatry*. 2004;43(12):1521–39. <https://doi.org/10.1097/01.chi.0000142280.87429.68>.

Publisher’s Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.