W J C C World Journal C Clinical Cases

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World J Clin Cases 2024 March 6; 12(7): 1251-1259

DOI: 10.12998/wjcc.v12.i7.1251

ISSN 2307-8960 (online)

ORIGINAL ARTICLE

Retrospective Study Clinical characteristics of testicular torsion and factors influencing testicular salvage in children: A 12-year study in tertiary center

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Peer-review report's scientific quality classification Grade A (Excellent): 0 Grade B (Very good): 0	Corresponding author: Duo-Bing Zhang, MD, Academic Research, Surgeon, Department of Urology, Suzhou Hospital Affiliated to Anhui Medical University, No. 299 Bianhe Middle Road, Suzhou 234000, Anhui Province, China. zhangduobingsz@126.com
Grade C (Good): C, C Grade D (Fair): 0 Grade E (Poor): 0	Abstract
P-Reviewer: Arslan M, Turkey; Govindarajan KK, India	BACKGROUND Testicular torsion is the most common acute scrotum worldwide and mainly occurs in children and adolescents. Studies have demonstrated that the duration of symptoms and torsion grade lead to different outcomes in children diagnosed
Received: November 26, 2023 Peer-review started: November 26,	with testicular torsion.
2023	AIM
First decision: December 29, 2023 Revised: January 9, 2024	To predict the possibility of testicular salvage (TS) in patients with testicular torsion in a tertiary center.
Accepted: February 4, 2024 Article in press: February 4, 2024 Published online: March 6, 2024	<i>METHODS</i> We reviewed the charts of 75 pediatric patients with acute testicular torsion during a 12-year period from November 2011 to July 2023 at the Suzhou Hospital
	of Anhui Medical University. Univariate and multivariate logistic regression analyses were used to determine independent predictors of testicular torsion. The data included clinical findings, physical examinations, laboratory data, color Doppler ultrasound findings, operating results, age, presenting institution status, and follow-up results.

RESULTS

Our study included 75 patients. TS was possible in 57.3% of all patients; testicular torsion occurred mostly in winter, and teenagers aged 11-15 years old accounted

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for 60%. Univariate logistic regression analyses revealed that younger age (P = 0.09), body mass index (P = 0.004), torsion angle (P = 0.013), red blood cell count (P = 0.03), neutrophil-to-lymphocyte ratio (P = 0.009), and initial presenting institution (P < 0.001) were associated with orchiectomy. In multivariate analysis, only the initial presenting institution predicted TS (P < 0.05).

CONCLUSION

The initial presenting institution has a predictive value for predicting TS in patients with testicular torsion. Children with scrotal pain should be admitted to a tertiary hospital as soon as possible.

Key Words: Testicular torsion; Testicular salvage; Pediatrics; Predictors; Case report

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Core Tip: We retrospectively collected cases of testicular torsion surgery in adolescents at a tertiary hospital and found that the initial visit to a non tertiary hospital may delay the patient's diagnosis and optimal surgical time.

Citation: Gang XH, Duan YY, Zhang B, Jiang ZG, Zhang R, Chen J, Teng XY, Zhang DB. Clinical characteristics of testicular torsion and factors influencing testicular salvage in children: A 12-year study in tertiary center. World J Clin Cases 2024; 12(7): 1251-1259 URL: https://www.wjgnet.com/2307-8960/full/v12/i7/1251.htm DOI: https://dx.doi.org/10.12998/wjcc.v12.i7.1251

INTRODUCTION

Testicular torsion (TT) is the most common acute scrotum injury worldwide and mainly occurs in children and adolescents, which causes restricted blood flow to the affected testis, resulting in testicular atrophy and testicular loss in adolescents. According to public data, the incidence of TT is estimated to be 1 in 4000 males aged < 25 years and 1 in 160 males aged > 25 years[1,2], and TT accounts for 13%–54% of cases of acute scrotal disease[3]. The incidence was high at 21.76 per 100000 population in 2010 in the country of Ireland, which has cold weather[4]. Currently, no drugs are effective for treating TT in humans, and the most effective treatment for TT is surgical exploration. However, delays in the diagnosis of TT result in loss of testes, subfertility, or infertility in patients [5]. TT is related to a variety of factors, the most important of which is the duration from testicular ischemia to surgery and the torsion angle [6-8]. In the current study, we reviewed patients who underwent orchiopexy or orchiectomy and were diagnosed with TT from 2011-2023 to summarize the clinical characteristics of TT and identify predictors of testicular salvage (TS) in patients.

MATERIALS AND METHODS

General information and surgery details

After receiving Institutional Review Board approval from the Ethics Committee of Anhui Medical University Suzhou Hospital (A2023006), we queried a maintained database of patients who underwent scrotal exploration at a tertiary referral medicine center from January 2011 to July 2023. In the current study, 75 patients were enrolled. Prior to study commencement, we obtained informed consent from patients who were preoperatively diagnosed with TT using Doppler ultrasonography (D-USG) (Supplementary Figure 1). Even young children with small testes, where testicular blood flow is difficult to detect, undergo D-USG to prevent misdiagnosis. The exclusion criteria for the retrospective study included: (1) Patients younger than one year old; (2) Patients who had severe hepatic or renal dysfunction or heart disease; and (3) Patients who underwent scrotal surgery before TT.

The patient was placed in the supine position, and after anesthesia was administered, the scrotum on the affected side was opened, and a warm, damp gauze was placed on the testis. If no blood supply was restored to the testicle after 15-20 min, an orchiectomy was performed. If the testicular blood supply was restored and bright red blood was observed on the needle prick, bilateral orchiopexy was performed.

According to type of surgery, patients were divided into two groups: TS and Orchiectomy groups.

Statistical methods

Categorical data were presented as numbers and percentages. Means and standard deviations were calculated for the continuous variables. Statistical analysis was performed using SPSS (version 22.0; IBM, Armonk, NY, United States). A student's t-test was used to perform parametric comparisons of normally distributed data. The Mann-Whitney U test was used to compare means between non-normally distributed groups. Fisher's exact test and chi-square test were used to compare categorical data between the two groups. Two-sided test values of P < 0.05 were considered statistically significant. To identify the predictive factors of orchiectomy and develop a novel predictive model, univariable and



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multivariable binary logistic regression analyses were used.

RESULTS

Clinical parameters of patients

Patients with TT confirmed by Doppler ultrasound who were all minors aged 1–18 years were included in the study cohort, and we ultimately collected clinical information from 75 patients through the application of exclusion and inclusion criteria. Among them, 32 underwent orchiectomy, and TS occurred in 43 patients. The mean age of the Orchiectomy group was 12.53 ± 3.29 years younger than 14.42 ± 2.41 years in the TS group (Table 1). The peak TT rate was observed between 11–15 years old (Figure 1A). The probability of TS increased with age (Figure 1B). The incidence of TT was high in the winter (Figure 1C). Over 84% of the patients had testicular pain, and 10% had scrotal swelling. A previous history of TT occurred in only 5% of the patients. The time from the event to hospitalization ranged from 4–115 min. The twisted testicle side and location of the TT were comparable between the TS and Orchiectomy groups.

When comparing the blood test results and testicular color of the affected testes, we found that all patients in the Orchiectomy group presented with a black color. The neutrophil-to-lymphocyte ratio (NLR) was higher in the Orchiectomy group than in the Salvage group.

We also compared differences between the Orchiectomy and Salvage groups. We find that the initial presenting institution accounts. More than 50% of the patients in the Orchiectomy group visited a primary hospital for the first time, while < 20% of the patients in the TS group visited a primary hospital (Table 2).

Independent predictors for testicular salvage rate

Interestingly, we found that over 80% of the patients in the Salvage group were sent to tertiary hospitals, while only 65% of the patients in the Orchiectomy group were sent to tertiary hospitals. The guardian type (father or mother) and residence (urban or rural) of the children had no predictive effect on testicular salvage.

Univariate analysis showed that age, body mass index, torsion angle, red blood cell count, NLR, and the initial presenting institution were predictive of orchiectomy (Table 3). Univariate and subsequent multivariate logistic regression analyses showed that the initial presenting institution correlated with the risk of a salvageable testis (Table 3).

DISCUSSION

TT is a common urological emergency in adolescents that often results in testicular loss, decreased fertility, and even loss of fertility if the diagnosis and management of TT are delayed. It has been reported that the overall incidence is approximately 4 in 100000 people worldwide, with a higher incidence of TT in Ireland, between 16.8 and 26.3[4]. At present, the initial causes of TT remain unknown, and mouse models have been used to study the causes of TT and protective factors of TT[9]. Some researchers have reported that TT causes dysfunction in mitochondrial oxidative phosphorylation[10] and that Insl3 could be a protective factor against TT[11]. Which clinical characteristics predict a higher risk for testicular orchiectomy? In the Orchiectomy group in our study, the testicular color was black, and the NLR was higher than that in the TS group. Multiple studies have also reported that torsion grade is associated with testicular vitality[12-14]. In this study, the degree of torsion was associated with TS. The median torsion grade was 568.12 ± 195.501 in the Orchiectomy group compared to 425.93 ± 235.747 in the TS group. Recently, Delgado-Miguel *et al*[15] confirmed that laboratory tests of the NLR should be considered a predictor of pediatric TT in cases with clinical and ultrasound suspicion, which may help anticipate the urgent surgical treatment required in these patients[15]. Therefore, testicular color, torsion grade, and laboratory tests will help urologists judge the severity of TT.

TT is a urological emergency; therefore, social factors may delay the diagnosis and treatment of patients with TT. Accurate and prompt diagnosis and surgical treatment are important for TS. TT can occur at any age, but teenagers are the main victims, especially children who are 12–16 years old[16]. Similarly, we found that patients aged 11–15 account for 60% of the cohort (Figure 1A). We found a correlation between age and TS, with younger children being more likely to undergo orchiectomy. This may be because younger children are unable to judge the severity of pain and accurately describe pain symptoms in words[17]. Similarly, Bayne *et al*[18] reported that misdiagnosed patients tended to be younger and more likely to have developmental, cognitive, or social disorders than those in an acute setting[18].

In addition to young age and atypical symptoms, what factors could lead to a delay in TT and failure of TS? In our study, we found that the initial presenting institution was important. TT is a urology emergency; therefore, sociological characteristics, such as tertiary hospital location, were also considered. Univariate and multivariate logistic regression analyses in the current study showed that only patients who visited the tertiary facility shortly after symptom onset had a higher salvage rate, which was similar to research in Brazil[19]. Tertiary hospitals often have better Doppler ultrasound equipment, more experienced ultrasound diagnosticians, and urologists at higher levels, which results in lower rates of misdiagnosis. In 2016, Douglas pointed out the inclination to refer adolescent patients with TT away from regional hospitals to a tertiary pediatric center and experienced urologists[20]. However, in northern and central China, many children live in rural areas far from tertiary hospitals in cities, which often causes delays in TT treatment.

In addition to preventing delays in diagnosis, we must avoid long durations before surgery. If patients with TT are diagnosed and treated promptly, the probability of testicular loss will decrease significantly. The "golden six-hour" window for TT treatment signifies that if patients reach the hospital within six hours of symptom onset, the salvage

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Table 1 Com	parisons of clinical	parameters between th	ne orchiectomy arou	up and salvage grou	p. n (%)
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Characteristic	Orchiectomy (n = 32)	Salvage (<i>n</i> = 43)	Total (<i>n</i> = 75)	P value
Age (yr), median (range)	12.53 ± 3.29 (2-18)	14.42 ± 2.41 (9-18)	13.61 ± 2.97 (2-18)	0.347
BMI (kg/m ²), mean \pm SD	18.75 ± 1.57	20.10 ± 1.97	19.53 ± 1.92	P < 0.01
Initial symptom				0.662
Testicular pain	25 (78.13)	38 (88.37)	63 (84.00)	
Oscheocele	5 (15.63)	3 (6.97)	8 (10.66)	
Nausea and vomiting	1 (3.12)	1 (2.33)	2 (2.67)	
Others	1 (3.12)	1 (2.33)	2 (2.67)	
Previous history of testicular torsion				0.417
No	29 (90.63)	41 (95.35)	70 (93.33)	
Yes	3 (9.37)	2 (4.65)	5 (6.67)	
Time from the event to the hospitalization (min), median (range)	42 (4-115)	37 (4-105)	41 (4-115)	0.791
The side of the twisted testicle				0.888
Left	25 (78.13)	33 (76.74)	58 (77.33)	
Right	7 (21.87)	10 (23.26)	17 (22.67)	
Intratesticular blood flow				0.657
Invisible	28 (87.50)	39 (90.70)	67 (89.33)	
Visible	4 (12.50)	4 (9.30)	8 (10.66)	
Testicular parenchymal echotexture				0.761
Nonhomogeneous	30 (93.75)	41 (95.35)	71 (94.67)	
Homogeneous	2 (6.25)	2 (4.65)	4 (5.37)	
Location of testicular torsion				0.739
Intrathecal	31 (96.88)	41 (95.35)	72 (96.00)	
Extrathecal	1 (3.12)	2 (4.65)	3 (4.00)	
Torsion angle	540 (90-1080)	360 (45-720)	540 (45-1080)	0.49
Testicular color of the affected testis				P < 0.01
Black	32 (100)	30 (69.77)	62 (82.67)	
Maroon	0	7 (16.27)	7 (9.33)	
Bright red	0	5 (11.63)	5 (6.67)	
Other	0	1 (2.33)	1 (1.33)	
Normal testicular fixation				0.289
No	1 (3.12)	4 (9.30)	5 (6.67)	
Yes	31 (96.88)	39 (90.70)	70 (93.33)	
Hospital stay time, median (range)	4 (2-8)	5 (1-9)	5 (1-9)	0.309
Red blood cell count ($10^9/L$), mean ± SD	4.58 ± 0.33	4.76 ± 0.35	4.69 ± 0.35	0.028
White blood cell count (10 ⁹ /L), median (range)	11.235 (5.16-19.31)	11.06 (5.00-22.08)	11.06 (5.00-22.08)	0.894
NLR, median (range)	3.055 (0.96-8.90)	5.28 (1.11-9.15)	4.23 (0.96-9.15)	0.014
LMR, median (range)	2.81 (0.49-6.14)	2.89 (1.25-6.36)	2.82 (0.49-6.36)	0.894

BMI: Body mass index; NLR: Neutrophil-to-lymphocyte ratio; LMR: Lymphocyte-to-monocyte ratio.

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Characteristic	Orchiectomy (n = 32)	Salvage (<i>n</i> = 43)	Total (<i>n</i> = 75)	P value
Season of events				0.933
Spring	7 (21.87)	11 (25.58)	18 (24.00)	
Summer	5 (15.63)	6 (13.95)	11 (14.67)	
Autumn	6 (18.75)	6 (13.95)	12 (16.00)	
Winter	14 (43.75)	20 (46.52)	34 (45.33)	
Patient's residence				0.689
Rural area	9 (28.12)	14 (32.56)	23 (30.67)	
Urban area	23 (71.88)	29 (67.44)	52 (69.33)	
Patient's guardian				0.749
Father	13 (40.63)	16 (37.21)	29 (38.67)	
Mother	10 (31.25)	17 (39.53)	27 (36.00)	
Other	9 (28.12)	10 (23.26)	19 (25.33)	
Initial presenting institution				P < 0.01
Primary hospital	18 (56.25)	8 (18.60)	26 (34.67)	
Tertiary hospital	14 (43.75)	35 (81.49)	49 (65.33)	
Median treatment cost [United States dollars, median (range)]	764.085 (526.75-1266.32)	726.19 (381.04-1363.83)	736.79 (381.04-1363.83)	0.739

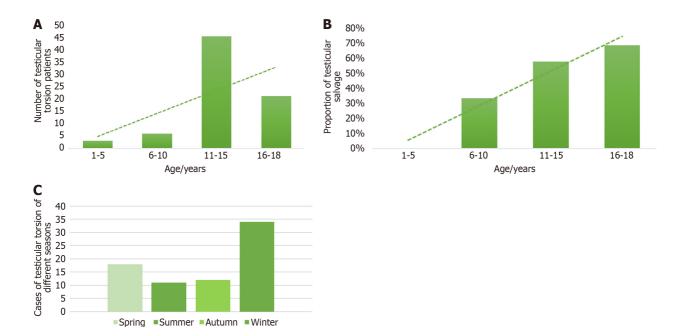


Figure 1 Proportion of testicular salvage in testicular torsion patients. A: Age related distribution rate of testicular retention in testicular torsion patients; B: Age related distribution rate of proportion of testicular salvage; C: Age related distribution rate of proportion of testicular salvage.

incidence can reach 90%[21]. Dias Filho *et al*[19] reported that presentation delay was a significant predictor of orchiectomy[19]. If surgical exploration is delayed, the TS rate decreases to 50% when symptoms last beyond 12 h. When symptoms last for over 24 h, the TS rate is reduced to less than 10%[22].

TT often occurs in cold weather at low temperatures. In cold zone countries, such as Ireland and Canada, the incidence rate of TT is relatively higher than that in other countries[4,23,24]. Even in tropical countries, TT follows a seasonal pattern and is more frequent in the colder months of the year[25]. Why do low temperatures cause TT? Researchers have suggested that contraction of the cremasteric muscle at low temperatures increases the risk of TT[26]. In Japan, researchers found that over 70% of TT cases occurred in winter[27], and in Brazil, the TT incidence was higher in the southern area than in the northern area (the cold zone in the south)[24]. According to a report in Taiwan, 73.7% of the patients had TT when the diurnal temperature change was $\geq 6^{\circ}$ C. Compared to the torsion rate for diurnal temperature

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Table 3 Univariate and multivariate logistic regression analyses of related factors related to testicular salvage

	Univariate analysis	Univariate analysis		/sis
Characteristic	OR (95%CI)	P value	OR (95%CI)	P value
Age (yr), median (range)	1.16 (0.977-1.378)	0.09		
BMI (kg/m²), mean ± SD	1.54 (1.14-2.06)	< 0.01	1.24 (0.46-3.39)	0.669
Initial symptom				
Testicular pain	Ref			
Scrotal swelling	0.39 (0.09-1.80)	0.230		
Nausea and vomiting	0.66 (0.04-11.01)	0.771		
Other	0.66 (0.04-11.01)	0.771		
Previous history of testicular torsion				
No	Ref			
Yes	0.72 (0.14-3.85)	0.706		
Time from the event to the hospitalization (min), median (range)	0.99 (0.97-1.01)	0.423		
The side of the twisted testicle				
Left	Ref			
Right	1.08 (0.36-3.24)	0.888		
Intratesticular blood flow				
Invisible	Ref			
Visible	0.72 (0.17-3.12)	0.658		
Testicular parenchymal echotexture				
Nonhomogeneous	Ref			
Homogeneous	0.73 (0.10-5.49)	0.761		
Operative time (min), median (range)	0.98 (0.95-1.00)	0.073		
Location of testicular torsion				
Intrathecal	Ref			
Extrathecal	1.51 (0.13-17.44)	0.740		
Torsion angle	1.00 (0.99-1.00)	< 0.05		
Testicular color of the affected testis				
Black	Ref			
Maroon	123358712.82 (0.00-Inf)	0.994		
Bright red	123358712.25 (0.00-Inf)	0.995		
Other	123358712.35 (0.00-Inf)	0.998		
Normal testicular fixation	· · · · · · · · · · · · · · · · · · ·			
No	Ref			
Yes	0.31 (0.03-2.96)	0.312		
Hospital stay time, median (range)	1.18 (0.96-1.44)	0.112		
Red blood cell count (10 ⁹ /L), mean ± SD	5.27 (1.18-23.57)	< 0.05	6.00 (0.10-348.34)	0.387
White blood cell count $(10^9/L)$, median (range)	0.98 (0.86-1.13)	0.811	(
NLR, median (range)	1.34 (1.08-1.67)	< 0.01	3.78 (0.96-14.91)	0.057
LMR, median (range)	1.21 (0.85-1.73)	0.285		0.007
Season of events	(0.00 1.0)			
Spring	Ref			
opinie	INCI			

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Summer	0.76 (0.17-3.49)	0.728		
Autumn	0.64 (0.15-2.78)	0.548		
Winter	0.91 (0.28-2.92)	0.873		
Patient's residence				
Rural area	Ref			
Urban area	0.81 (0.30-2.20)	0.681		
Patient's guardian				
Father	Ref			
Mather	1.38 (0.47-4.03)	0.554		
Other	0.90 (0.28-2.88)	0.863		
Initial diagnosis institution				
Primary hospital	Ref		Ref	
Tertiary hospital	5.62 (1.99-15.88)	< 0.01	12.58 (1.12-141.05)	< 0.05
Median treatment cost [United States dollars, median (range)]	1.00 (1.00-1.00)	0.650		

BMI: Body mass index; NLR: Neutrophil-to-lymphocyte ratio; LMR: Lymphocyte-to-monocyte ratio; OR: Odds ratio.

changes < 6°C, the relative risk of TT at \geq 6 °C was 1.8 (P < 0.05)[28]. This finding is similar to the results of the present study. We found that TT occurred more frequently in winter than in the other seasons, and winter was the season with the most severe temperature changes in northern and central China.

From the perspective of disease characteristics, the degree of TT and laboratory indicators of blood tests affect the success rate of TS in TT. However, it should be noted that TT is a urologic emergency, and the delay in diagnosis due to sociological characteristics leads to the loss of testes. In rural areas or small towns, the lack of tertiary hospitals causes delays in diagnosis. Therefore, for better treatment, we should popularize emergency knowledge among parents and children^[29], and provide corresponding diagnosis and treatment training for doctors in rural areas to avoid misdiagnosis of TT.

This study has several limitations. First, it is a retrospective study, which is subject to selection bias. Secondly, the study is single-centered with a limited number of cases included. In the future, larger-scale multicenter studies will be needed to identify factors that influence the outcomes of TT.

CONCLUSION

The initial presenting institution has a predictive value for predicting outcomes in children with acute TT. Children with scrotal pain should be referred to a tertiary hospital as soon as possible for an accurate diagnosis and timely surgery.

ARTICLE HIGHLIGHTS

Research background

Testicular torsion (TT) is currently the leading cause of testicular removal in adolescents, with an incidence rate of 1 in 4000 among males under the age of 25. Delayed medical consultation for TT and misdiagnosis could lead to orchiectomy. Therefore, we aim to analyze the factors influencing orchiectomy or testicular salvage by reviewing the data of patients who presented at our center.

Research motivation

Various factors, such as delayed presentation to the healthcare facility and excessive degree of torsion, can lead to orchiectomy in adolescents with TT. Hence, we conducted a retrospective analysis of data from our center to identify the factors affecting the prognosis of TT.

Research objectives

Factors to predict the possibility of testicular salvage (TS) in patients with TT in our tertiary center.

Research methods

By collecting clinical data from patients with TT treated at our center, we performed univariate and multivariate



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regression analyses to compare between the TS and the orchiectomy group.

Research results

Our study has identified that younger age, body mass index, torsion angle, red blood cell count and neutrophil-tolymphocyte ratio have predictive value for TS. And we found a new factor that the type of institution where patients first presented as a predictive factor for the outcome of TT; patients who initially presented to a tertiary-care hospital were more likely to have testicular preservation.

Research conclusions

The initial presenting institution is a new predictive factor for outcome of TT. Patients with scrotal pain should be presented to a tertiary hospital as soon as possible.

Research perspectives

Future research will focus on enhancing public health education regarding scrotal emergencies to prevent delays in the presentation of patients with conditions such as TT.

FOOTNOTES

Co-first authors: Xiang-Hui Gang and Yuan-Yuan Duan.

Author contributions: Gang XH study conception and design, analysis and data interpretation, drafting of the manuscript, and critical revision; Duan YY, Jiang ZG and Teng XY study conception and design, data acquisition, and critical revision; Zhang B, Zhang R, and Chen J data acquisition, analysis, interpretation, and critical revision; Zhang DB study conception and design, analysis and data interpretation, drafting of the manuscript, and critical revision.

Supported by Anhui Province Translational Medicine Research Fund Project, No. 2021zhyx-C59 and No. 2021zhyx-C75.

Institutional review board statement: The study was reviewed and approved by the Ethics Committee of Anhui Medical University Suzhou Hospital Institutional Review Board, Approval No. A2023006.

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

Conflict-of-interest statement: All the authors report no relevant conflicts of interest for this article.

Data sharing statement: No additional data are available.

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S-Editor: Li L L-Editor: A P-Editor: Zhao S

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