# INCREASING PREVALENCE OF CEREBRAL PALSY AMONG CHILDREN AND ADOLESCENTS IN CHINA 1988–2020: A SYSTEMATIC REVIEW AND META-**ANALYSIS**

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**Objectives:** To investigate the pooled prevalence of cerebral palsy in China, analyse the differences between different subgroups, and explore the trend over the 32-year period from 1988 to 2020.

Methods: All potential studies related to the prevalence of cerebral palsy among children and adolescents in China were identified from 3 Englishlanguage databases and 4 Chinese-language databases. Pooled prevalence was calculated to estimate the prevalence of cerebral palsy among 0-18 years old and different geographical regions in China, using a random-effects meta-analysis model. Continuous fractional polynomial regression modelling was used to estimate the trend in prevalence of cerebral palsy over time. Subgroup analysis and meta-regression were conducted to investigate heterogeneity. Funnel plots and Egger's test were used to explore potential publication bias.

Results: The pooled prevalence of cerebral palsy over the study period among 0-18 years old and different geographical regions in China was 2.07‰ (95% confidence interval (95% CI) 1.66-2.47‰), and the prevalence of cerebral palsy was higher in males compared with females (2.25‰ vs 1.59‰), and in rural residents compared with urban residents (2.75‰ vs 1.90‰), respectively. The prevalence of cerebral palsy varied significantly between different geographical regions. In subjects with birthweights < 2.5 and > 4 kg, the prevalence of cerebral palsy was significantly higher than in subjects with birthweights between 2.5 and 4 kg. The trend in pooled prevalence of cerebral palsy increased continuously over the period studied, and could be divided into 3 stages; the mean annual increase in prevalence from 1988 to 1996 and from 2008 to 2019 was more rapid. Multivariate meta-regression found that the year of study was one of the sources of heterogeneity among overall prevalence. (p-value = 0.006).

Conclusion: The pooled prevalence of cerebral palsy over the 32-year period from 1988 to 2020 was 2.07‰. There was an increasing trend in prevalence of cerebral palsy among children and adolescents in China over this period.

Key words: cerebral palsy; children; prevalence; meta-analvsis: China.

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### LAY ABSTRACT

There are few reports regarding trends in childhood cerebral palsy in China over the past 30 years. This study searched and statistically analysed all potentially relevant studies regarding cerebral palsy. The trend in childhood cerebral palsy in China increased continuously over the period studied, and could be subdivided into 3 stages. From 1988 to 1996, and from 2008 to 2019, the mean annual increase in prevalence was more rapid than for the other years studied. Counter-measures, such as health monitoring and management of infants and young children, are recommended to reduce the prevalence of childhood cerebral palsy in China.

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reversion of heterogeneous non-progressive neurodevelopmental conditions that affect the developing foetal or infant brain (1); it is one of the most common risk factors for motor impairment (2) and limitation in manual ability, which is strongly related to limitations in daily activities (3). In line with the global trend, CP places a heavy burden of disease on children, families and society in both developed and developing countries. In Norway, the prevalence of CP decreased from 2.62 per 1,000 live births in 1999 to 1.89 per 1,000 live births in 2010 (4). A register-based cohort study in Denmark reported that the prevalence of CP was 2.2/1.000 in children of Danish-born mothers in 2018 (5). Furthermore, 2 population-based studies, from Uganda and Bangladesh, on the prevalence of CP reported prevalences of 2.9 per 1,000 children in 2017 and 3.4 per 1,000 children in 2016 (6, 7), respectively. According to a systematic review published in 2013, the global prevalence of CP was 2.11 per 1,000 live births (8).

In China, a "birth deficient registration system" has been in operation since the 1980s, but it lacks a specific registration system for CP. Lv (9) reported the prevalence of childhood CP in 1988 in China as 1.20 per 1.000

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children. With the acceleration of China's economic development and the rapid population growth in the past 30 years, CP is one of the leading diseases with a heavy burden of disease among Chinese children (10). It has been reported that the lifetime total economic loss due to all new cases of CP in 2003 in China amounted to US\$ 2-4 billion (11). Based on data from multiple provinces in China, the prevalence of CP among Chinese children has been reported to range from 1.92 to 2.46‰ in 1998 and 2013 (12, 13), respectively. A systematic review published in 2015 (14) reported the prevalence of CP among children as 1.80‰, including 21 studies published from 2000 to 2014, based on data from Pub-Med and 4 Chinese databases. However, there have been few reports on the trend in childhood CP in China in the past 30 years.

In addition, some updated original data published in recent years, has combined data from diverse populations, types of area and geographical regions. Hence, the aims of this study are to provide a comprehensive update on the trend in prevalence of CP in China, from 1988 to 2020, and to systematically analyse childhood CP by urban or rural area, five geographical regions, age group and birthweight.

### **METHODS**

This systematic review and meta-analysis adhered to the 2009 Preferred Reporting Items for Systematic Reviews and Metaanalyses (PRISMA) guidelines (15).

### Search strategy

Three English-language databases (PubMed, Embase and Web of Science) and 4 Chinese-language databases (China National Knowledge Infrastructure (CNKI), Wanfang, Weipu, and China Biology Medicine disc (CBMdisc)) were searched to identify all potential studies related to the prevalence of CP among children and adolescents in China published from database inception to 5 February 2021. The following search terms were used: cerebral palsy, mixed cerebral palsy, athetoid cerebral palsy, spastic diplegia, prevalence, morbidity, epidemiology, China, Chinese, children, and adolescent. The specific search strategies for each database are shown in Table SI1. The language of publication was restricted to English and Chinese. The reference lists of included articles and previous reviews were retrieved to identify potential studies as comprehensively as possible. Two authors, working independently, carried out the searches (SY Y and JY X).

#### Inclusion and exclusion criteria

The inclusion criteria were: (*i*) population-based studies; (*ii*) original investigations reporting data among children (under 12 years old) and adolescents (12–18 years old) in China; (*iii*)

The exclusion criteria were: (*i*) original investigations reporting data are adults (over 18 years old) or not Chinese; (*ii*) not related to CP; (*iii*) the diagnostic criteria for CP were not clear or not mentioned; (*iv*) did not report the prevalence of CP or information inadequate to evaluate the prevalence; (*v*) research plan, regarding design, research approach and research process, not clear; (*vi*) reviews, duplicate publications, randomized controlled trials (RCTs) and case-control studies.

### Data extraction

Two researchers (SY Y and JY X) independently abstracted the following information from each article: author, year of publication, medium year of data collection, geographical location of the study, sampling method, case definition, sample size, number of cases of CP, and prevalence of CP.

Subgroup information for number of cases and number of populations, including sex (male, female), age group (0–1, 1–3, 3–6 years), geographical location (East China, Central China, South China, Southwest China, Northwest China, North China, Northeast China), regions (urban or rural), birthweight (under 2.5 kg, 2.5–4 kg, over 4 kg), gestation (under 37 weeks, 37–42 weeks, over 42 weeks) and classification of CP, were collected.

The study authors were contacted for additional data or clarification, if necessary. Disagreements were resolved by consensus after discussion.

### Quality assessment

Two researchers assessed the quality of each included study using the quality assessment criteria for observational studies recommended by the Agency of Healthcare Research and Quality (AHRQ) (17). These assessment criteria included 11 criteria with 3 potential responses: yes, no and unclear. Briefly, "yes"=1 point, "no or unclear"=0 points, and a maximum score of 11 was possible for each study. The total score for each study was calculated. Studies with a score <3 points were considered low quality, 4–7 points medium quality, and 8–11 points high quality.

#### Statistical analysis

Pooled prevalence and 95% CI of children with CP were calculated. A  $\chi^2$ -based Q test and the  $I^2$  test were performed to evaluate the heterogeneity of the studies. A random-effects meta-analysis model was used when the heterogeneity was statistically significant ( $l^2 > 50$ , p-value < 0.05) (18). Furthermore, subgroup analysis was performed to evaluate the levels of prevalence of CP according to different subgroups. In addition, a continuous fractional polynomial regression model at the midpoint of each study period was used to estimate the trend in prevalence over time. Egger's test was performed to explore potential publication bias. To explore the sources of heterogeneity, sensitivity analysis, univariate and multivariable meta-regression analysis were performed including the following variables: year of publication, year of study, geographical location, male/total, mean age, quality scores, sample size, and research type. p-value < 0.05 was considered

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statistically significant. EndNote X9 (Thomson Corporation, American) was used for reference management, Excel 2008 for data arrangement, Stata 15.0 for data analysis, and Origin 2019b for graphs (Origin Lab, American).

### RESULTS

## Selection of studies

The search strategy yielded a total of 839 abstracts: 121 from PubMed, 178 from Embase, 38 from Web of Science, 174 from CNKI, 162 from Wanfang, 106 from CBM, and 60 from Weipu. A manual search of the references cited in an available systematic review of CP prevalence yielded an additional 6 abstracts. After removal of duplicates, 450 studies remained. After title and abstract screening, 112 full-texts were assessed for eligibility based on the inclusion and exclusion criteria, and 57 studies were excluded, as follows: 8 review articles, 4 studies based on non-Chinese populations, 2 studies that used recruitment from schools, 7 studies conducted in populations other than children and adolescents, 19 studies that lacked data on population,

18 studies with no diagnostic criteria for CP, and 11 studies that used data from the same study populations. A final total of 43 studies were included in the quantitative synthesis. The study selection process is shown in Fig. 1.

### *Characteristics of studies*

The 43 studies included in this systematic review were published during the years 1993 to 2020, among which, the data were collected between 1988 and 2019. The geographical locations included were East China (15 studies), Central China (8 studies), South China (5 studies), Southwest China (6 studies), Northwest China (4 studies), North China (1 studies), and Northeast China (2 studies). The age of children mostly ranged from 0 to 6 years, and almost 85.70% of the population were live births. The sample sizes of the included studies ranged from 1,712 to 12,902,002, with a total of 17,723,942 people. Quality scores for the studies ranged from 5 to 10. A detailed description of the studies is shown in Table I.

### Pooled prevalence of CP

As shown in Fig. 2, the pooled prevalence of CP was 2.07 per 1,000 persons (95% CI 1.66–2.47‰), ranging from 1.02 to 7.59 per 1,000 persons. Significant heterogeneity was found among studies (P=99.4%, *p*-value<0.001).

### Subgroup analysis

Prevalence of CP by sex and age. The pooled prevalence of CP among children per 1,000 live births was calculated by sex. The prevalence of CP was significantly higher (*p*-value <0.001) among males (2.25‰, 95% CI 1.65–2.85‰) compared with females (1.59‰, 95% CI 1.35–1.82‰), and heterogeneity was found among the different subgroups ( $I^2$  for males=99.8%, *p*-value <0.001;  $I^2$  for females=95.9%, *p*-value <0.001). Regarding the age subgroups (Fig. 3), the pooled prevalence of CP was different among the different age groups, and the highest level was found in the 0–1 year group (1.95‰, 95% CI 1.44–2.46‰) (Fig. 3).

*Prevalence of CP by geographical location and region.* The prevalence of CP varied significantly between different geographical regions. The pooled prevalence of CP was highest in Southwest China (2.80‰, 95% CI 1.91–3.70‰), followed by Northwest China (2.55‰,

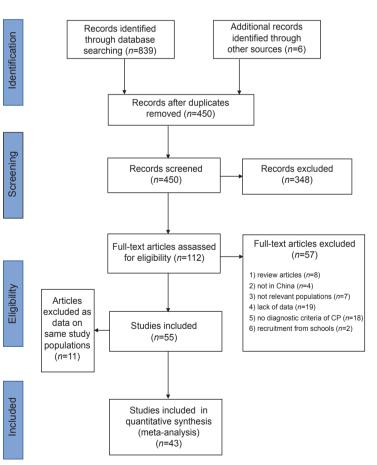


Fig. 1. Flow diagram of study selection process. CP: cerebral palsy.

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### Table I. Characteristics of studies on the prevalence of cerebral palsy (CP)

Reference	Publicat year	tion Study year	Province/s	Geographical location	Range of birth time	Age range, years	Cases with CP, n	Sample size, n	Denominator	Prevalence (‰)	Qualit scores
Lv et al. (9)	1993	1988	Sichuan	Southwest China	1974-1988	0-14	15	12,489	Children < 14 years	1.20	6
Ma et al. (40)	1996	1995	Yinchuan, Ningxia	Northwest China	1982-1995	0-12	31	16,872	Children <13 years	1.84	8
Li et al. (41)	1996	1996	Jiaxing, Zhejiang	East China	1988-1995	0-6	98	69,852	Live births	1.40	8
Song et al. (42)	2002	1997	Suzhou, Jiangsu	East China	1990-1997	0-6	92	56,231	Live births	1.64	10
Li et al. (43)	2003	1997	Jiangsu	East China	1991-1997	1-6	484	305,263	Live births	1.59	9
Li et al. (44)	2000	1997	Jilin	Northeast China	1991-1997	0-6	56	30,876	Live births	1.81	8
Liu et al. (45)	2000	1997	Jiangsu	East China	1990-1996	0-6	622	388,192	Children <7 years	1.60	7
Xie et al. (46)	2005	1997	Leshan, Sichuan	Southwest China	1991-1997	0-6	50	22,180	Live births	2.25	8
Wang et al. (47)	2002	1997	Suzhou, Jiangsu	East China	1990-1997	0-6	108	63,102	Live births	1.71	9
Zhang et al. (48)	2000	1998	Wujin, Jiangsu	East China	1991–1997	0-6	63	62,003	Live births	1.02	10
Li et al. (49)	2000	1998	Nanning, Guangxi	South China	1991-1997	1-6	39	30,485	Live births	1.28	8
Zhao et al. (50)	2002	1998	Zhejiang	East China	1992-1997	1-6	92	62,949	Live births	1.46	10
Liang et al. (51)	2002	1998	Guangxi	South China	1991–1997	1-6	193	150,806	Live births	1.28	9
Tao et al. (52)		1998	Anhui	East China	1990-1998	1-8	105	50,714	Live births	2.07	7
Lin et al. (12)	2001	1998	6 provinces in China	-	1991–1997	1-6	2009	1,047,327	Children 1–6 years	1.92	7
Zhong et al. (53)	2002	1998	Chengdu, Sichuan	Southwest China			348	148,723	Live births	2.07	9
Guo et al. (54)	2003	1998	Gansu	Northwest China	1991–1997	1-6	395	152,463	Live births	2.59	7
Chen et al. (55)	2000	1998	Hefei, Anhui	East China	1985-1996	2-13	11	7,388	Live births	1.49	10
Yan et al. (56)	2002	1999	Guangxi	South China	1991–1997	1-6	100	61,912	Live births	1.62	7
Xing et al. (57)	1999	1999	Heilongjiang	Northeast China	1991–1997	1-6	25	13,755	Live births	1.82	10
Sun et al. (58)	2001	2000	Rizhao, Shandong	East China	1993-1999	0-7	81	62,989	Live births	1.29	8
Tao (59)	2001	2000	Anhui	East China	1992-2000	1-8	193	104,120	Children 1–8 years	1.85	8
Shi et al. (60)	2004	2000	Hebei	North China	1993–1999	1-6	171	96,435	Live births	1.77	8
Tao et al. (61)	2002	2000	Anhui	East China	1992-2000	0-8	88	53,406	Live births	1.65	10
Gao et al. (62)	2003	2001	Henan	Central China	1995-2000	0-6	582	434,920	Live births	1.34	8
Yuan et al. (63)	2005	2002	Xiangtan, Hunan	Central China	1995-2002	0-7	385	179,895	Live births	2.14	9
Wang et al. (64)	2005	2002	Fujian	East China	1985-2001	1–16	298	179,393	Live births	1.49	9
Tseng et al. (65)	2018	2005	Taiwan, China	East China	2002-2008	0-6	49521	1,290,2002	Children <7 years	3.84	8
Li (66)	2010	2009	Chongqing	Southwest China			13	1,712	Live births	7.59	5
Tan et al. (67)	2011	2009	Chongqing	Southwest China			17	3,360	Live births	5.06	7
Hu et al. (68)	2016		Hunan	Central China	2005-2010		55	20,003	Live births	2.75	9
Ma et al. (69)			Henan	Central China	2005-2010		120	51,108	Live births	2.35	8
Song (70)	2014		Henan	Central China	2005-2010		69	24,500	Live births	2.82	10
Li (71)	2012		Shandong	East China	2005-2010		36	20,077	Live births	1.79	10
Li et al. (72)			Qinghai	Northwest China			54	10,000	Live births	5.40	9
Yuan et al. (73)	2019		Henan	Central China	2005-2010		120	50,596	Children <7 years	2.37	9
Peng et al.	2016		Xiamen, Fujian	East China	2010-2014		32	20,915	Live births	1.53	8
(74) Li et al. (13)	2018		12 provinces in China	-	2005-2010		797	323,858	Live births	2.46	10
Lin et al. (75)	2015	2014	Kazak Autonomous prefecture of Ili- Xinjiang	Northwest China	2006-2012	1-6	515	288,368	Live births	1.79	10
Wang et al. (76)	2016	2015	Ezhou, Hubei	Central China	2007-2014	0-6	106	71,310	Live births	1.49	10
Zhang et al. (77)	2016	2015	Foshan, Guangdong	South China	2008-2013	0-6	60	28,704	Live births	2.09	6
Li et al. (78)	2018	2016	Hainan	South China	2008-2014	0-6	80	37,862	Live births	2.11	8
Luo et al. (79)	2020	2010	Xizang	Southwest China	2009-2018	0-10	29	4,827	Live births	5.80	9

\*The quality of each included study was assessed by using the quality assessment criteria for observational studies recommended by the Agency of Healthcare Research and Quality (AHRQ) (17).

Lv,et al.(9)	1.20 (0.59, 1.81) 2.36
Li,et al.(41) ' Song, et al.(42) Li,et al.(43) Li,et al.(43) Li,et al.(45) Wang, et al.(46) Wang, et al.(48) Li,et al.(49) Zhao, et al.(50) Li,et al.(49) Zhao, et al.(51) Tao, et al.(52) Lin, et al.(52) Lin, et al.(54) Chen, et al.(55) Yan, et al.(56) Xing, et al.(57) Sun, et al.(58) Tao, et al.(61) Gao, et al.(62) Yuan, et al.(63) Wang, et al.(66) Tan, et al.(67) Hu, et al.(68) Ma, et al.(68) Ma, et al.(68) Song, et al.(71) Li,et al.(75) Yuan, et al.(75) Yuan, et al.(75) Yuan, et al.(75) Yuan, et al.(76) Zhang, et al.(77) Li,et al.(78) Luo, et al.(79) Luo, et al.(79) Verall (1-squared = 99.4%, p = 0.000)	1.84 (1.19, 2.49) 2.34 1.40 (1.12, 1.68) 2.46 1.64 (1.31, 1.97) 2.45 1.59 (1.45, 1.73) 2.48 2.25 (1.63, 2.87) 2.35 1.71 (1.39, 2.03) 2.45 1.02 (0.77, 1.27) 2.47 1.28 (0.88, 1.68) 2.43 1.46 (1.16, 1.76) 2.46 1.28 (1.10, 1.46) 2.48 2.07 (1.67, 2.47) 2.43 1.92 (1.84, 2.00) 2.49 2.07 (1.84, 2.00) 2.47 2.59 (2.33, 2.85) 2.46 1.49 (0.61, 2.37) 2.22 1.62 (1.30, 1.94) 2.45 1.82 (1.11, 2.53) 2.31 1.29 (1.01, 1.57) 2.46 1.85 (1.59, 2.11) 2.46 1.85 (1.59, 2.11) 2.46 1.85 (1.59, 2.11) 2.46 1.85 (1.59, 2.11) 2.46 1.65 (1.31, 1.99) 2.44 1.34 (1.23, 1.45) 2.48 2.75 (2.02, 3.48) 1.49 2.75 (2.02, 3.48) 1.49 2.35 (1.93, 2.77) 2.42 2.62 (2.16, 3.48) 2.33 1.79 (1.21, 2.37) 2.47 1.49 (1.31, 1.67) 2.48 3.84 (3.81, 3.87) 2.49 7.59 (3.48, 11.70) 0.69 5.06 (2.66, 7.46) 1.32 2.75 (2.02, 3.48) 2.30 2.35 (1.93, 2.77) 2.42 2.82 (2.16, 3.48) 2.33 1.79 (1.21, 2.37) 2.47 1.49 (1.21, 1.77) 2.47 2.82 (2.16, 3.48) 2.33 1.79 (1.21, 2.37) 2.42 2.63 (1.96, 6.84) 1.89 2.37 (1.95, 2.79) 2.42 1.53 (1.00, 2.06) 2.39 2.46 (2.29, 2.63) 2.48 1.79 (1.64, 1.94) 2.48 1.79 (1.64, 1.94) 2.48 1.79 (1.64, 1.94) 2.48 1.99 (1.66, 2.67) 2.47 1.58 (0.3.66, 7.94) 1.46 2.07 (1.66, 2.47) 100.00

Fig. 2. Forest plot of prevalence of cerebral palsy (CP) in Chinese children and adolescents in the random-effects model (‰). NO.: number; 95% CI: 95% confidence interval; ES: estimated statistics.

	NO. of studies I <sup>2</sup>		Р		
Subgroup			value	Prevalence (95% CI)	
Gender:Male	36	99.8	< 0.001	2.25 (1.65, 2.85)	•
Female	36	95.9	< 0.001	1.59 (1.35, 1.82)	•
Age:(0,1y]	7	83	< 0.001	1.95 (1.44, 2.46)	•
(1,3]	7	61.9	0.015	1.90 (1.65, 2.15)	•
(3,6]	7	80.4	< 0.001	1.47 (1.19, 1.75)	•
Birthweight:(0,2.5kg)	6	90.5	< 0.001	20.55 (14.01, 27.09)	
[2.5,4kg]	6	97.3	< 0.001	1.82 (1.40, 2.23)	•
(4,+)	6	86.5	< 0.001	2.02 (1.08, 2.96)	•
Region:Urban	8	98.3	< 0.001	1.90 (0.73, 3.08)	•
Rural	8	98.9	< 0.001	2.75 (1.54, 3.95)	•
Classification:Spastic	12	93.5	< 0.001	1.31 (1.07, 1.55)	•
Hypotonia	12	94.4	< 0.001	0.24 (0.15, 0.33)	•
Ataxic	12	79.2	< 0.001	0.08 (0.05, 0.11)	+
Mixed	12	87.3	< 0.001	0.14 (0.09, 0.20)	•
Location:East China	15	99.6	< 0.001	1.76 (0.99, 2.53)	•
Central China	8	93.8	< 0.001	1.97 (1.57, 2.37)	•
South China	5	78.5	< 0.001	1.63 (1.30, 1.97)	•
Southwest China	6	84.7	< 0.001	2.80 (1.91, 3.70)	•
Northwest China	4	93.9	< 0.001	2.55 (1.82, 3.29)	•
Northeast China	2	0	0.982	1.81 (1.42, 2.21)	•
North China NOTE: Weights are from random e	1 effects analy	0 Isis	1	1.77 (1.51, 2.04)	•
				-5	0 1 2 3 4 5

Fig. 3. Prevalence of cerebral palsy (CP) in children and adolescents in China (‰): subgroup meta-analysis and analysis of heterogeneity. 95% - CI: 95% confidence interval; NO.: number; y: years.

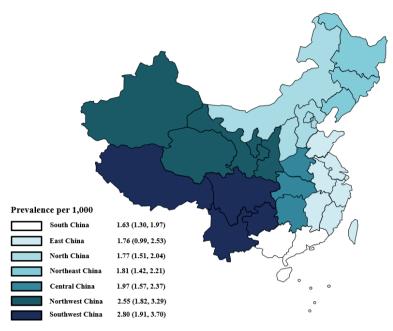


Fig. 4. Prevalence of cerebral palsy (CP) in children and adolescents in different geographical locations of China.

95% CI 1.82–3.29‰), and was lowest in South China (1.63‰, 95% CI 1.3–1.97‰) (*p*-value <0.001, Fig. 3 and Fig. 4). Urban residents had a much lower prevalence of CP (1.90‰, 95% CI 1.75–3.95‰) compared with rural residents (2.75‰, 95% CI 0.73–3.08‰, Fig. 3).

*Prevalence of CP by classification of CP.* As for the prevalence of CP by different classifications of CP, spastic was highest (1.31‰, 95% CI 1.07–1.55‰) and ataxic the lowest (0.08‰, 95% CI 0.05–0.11‰).

*Prevalence of CP by birthweight.* The prevalence of CP among children with birthweights under 2.5 kg (20.55‰, 95% CI 14.01–27.09‰) and over 4

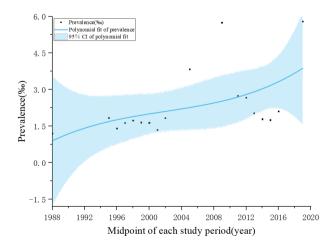


Fig. 5. Long-term trend in prevalence of cerebral palsy (CP) in Chinese children and adolescents over time. 95% CI: 95% confidence interval.

kg (2.02‰, 95% CI 1.08–2.96‰) was significantly higher than those of birthweight 2.5–4 kg (1.82‰, 95% CI 1.40–2.23‰), especially for the group under 2.5 kg (*p*-value < 0.001, Fig. 3).

### Prevalence of CP over time

Continuous fractional polynomial regression modelling on the midpoint of each study period was used to estimate the trend in CP in Chinese children over time (Fig. 5). The trend increased continuously over time, and could be divided into 3 stages. From 1988 to 1996, and 2008 to 2019, the mean annual increases in prevalence were approximately 9.14% and 38.13%, respectively, and between 1996 and 2008 the prevalence increased by approximately 9% per year.

### Publication bias

Egger's test showed that there was significant publication bias in this metaanalysis (t=-4.6, p-value<0.05) (Fig. 6).

### Sensitivity analysis and meta-regression analysis

Sensitivity analysis was conducted with 43 studies that were distributed on both sides of the pooled prevalence (1.02-7.59%). After excluding the 3 articles with 5 or 6 quality scores and the 2 articles that were not cross-sectional studies, the prevalence of CP was similar to the overall pooled estimates (2.05%) and 2.07%, which strengthened the credibility of the original analysis results.

A meta-regression analysis was performed to evaluate the potential risk factors for the prevalence of CP (*p*-va-

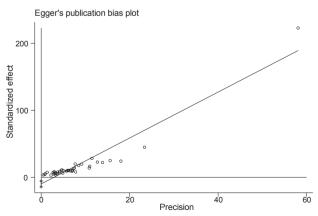


Fig. 6. Egger's publication bias plot of the prevalence of cerebral palsy (CP) (p-value < 0.05).

	Meta- regression			Adjusted R-squared
Covariate	coefficient	95% CI	p-value	(%)
Univariate analyses				
Year of studying	0.0334	0.0093 to 0.0574	0.008	23.46
Geographical location <sup>a</sup>	0.089	-0.0266 to 0.2047	0.128	3.48
Male/total%	2.4178	-4.7218 to 9.5573	0.498	1.84
Mean age	-0.0942	-0.2585 to 0.0700	0.253	2.22
Quality scores	-0.0149	-0.1830 to 0.1533	0.859	4.82
Sample size	-2.16E-07	-1.29E-06 to 8.62E-07	0.688	6.87
Research type <sup>a</sup>	0.2253	-0.7403 to 1.1909	0.640	1.66
Multivariable analyses				24.12
Year of study	0.0387	0.0121 to 0.0654	0.006	
Geographical location <sup>a</sup>	0.1101	-0.0058 to 0.2260	0.062	
Male/total%	4.8524	-1.8162 to 11.5210	0.148	
Mean of age	-0.0144	-0.1719 to 0.1432	0.854	
Quality score	-0.0729	-0.2344 to 0.0885	0.365	
Sample size	3.91E-08	-9.42E-07 to 1.02E-06	0.936	
Research type <sup>a</sup>	-0.1032	-0.1006 to 0.8000	0.818	

<sup>a</sup>Studies restricted to East China = 1, Central China = 2, South China = 3, Southwest China = 4, Northwest China = 5, North China = 6, Northeast China = 7 and data from cross-sectional studies = 1, baseline investigations from prospective studies = 2.

lue=0.049, P=87.48%) (Table II). In the univariate analyses, the heterogeneity was not modified by geographical location, male/total, mean age, quality scores, sample size or research type, but the year of study was significantly associated with the prevalence of CP (p-value=0.008 for the year of study). As shown by multivariable analysis, the year of study was significantly associated with prevalence of CP (p-value=0.006) (Table II).

### **DISCUSSION**

A total of 43 studies were included in this systematic review, for which the data were collected between 1988 and 2019. The pooled prevalence of CP was 2.07‰, which was a little higher than reported in a review in China, in 2015 (1.8‰) (14), and lower than the mean of 2.21‰ worldwide in 2013 (8). In addition, the continuous fractional polynomial regression model showed that the prevalence of CP among children in China has increased continuously over the 32-year period from 1988 to 2020.

There are a number of reasons associated with the increasing prevalence of CP in China. With economic development, medical conditions have gradually improved. The availability of neonatal intensive care units and high-technology diagnostic procedures in China has led to increased survival of premature infants, which may be associated with higher risk of CP (19, 20). Urbanization entered a period of accelerated development in the 1990s (21). According to the sixth national census in 2010, half of China's population was living in urban areas (22). Urbanization not only transforms the urban environment, increasing air pollution, occupational and traffic hazards, but encourages

changes in people's lifestyle (22), factors which are associated with the prevalence of CP (23). With economic development and improved living conditions in China, women are consuming increasing amounts of sugar and fats, which may be risk factors for gestational hypertension and diabetes mellitus (23). Lastly, previous estimates suggest that the contribution of genetic variants to the burden of CP is approximately 12% (24). Established environmental risk factors for CP may interact with predisposing genetic variants, potentiating and multiplying the risk of CP (25). Furthermore, familial clustering of CP has been described as the risk of CP for the sibling of a child with CP, and this is increasing (26).

In 2013, China applied the policy allowing marital couples in which at least 1 of the partners was an only-child to have 2 children, and in 2015. China finally ended all 1-birth restrictions and moved to a nationwide 2-child policy (27), which resulted in an increase in multiparous mothers and mothers aged 35 years and over (28). However, multiple pregnancies, elderly maternal age (35 years and over), and parity of 3 or more were the risk factors for preterm infants (23, 29). In addition, the model indicated a sharply increasing trend in the prevalence of CP before 1996. However, only 3 studies from 1988 to 1996 were included in the analysis, which might have introduced some heterogeneity. In addition, the denominator used in 1988 was the number of the children <14 years of age, while the denominator in 1995 was the number of children < 13 years, which may have led to a sharp increase in prevalence before 1996 when these 2 age groups were considered as denominators.

The longitudinal trend in childhood CP in China for this period is different from that in more developed countries, whose trends decreased. Population-based databases and registers in Europe and Australia show that the trend in childhood CP was decreasing (30). In Norway, the prevalence of CP decreased from 2.62 per 1,000 live births in 1999 to 1.89 per 1,000 live births in 2010 (4). In Sweden, the prevalence of childhood CP has been continuously reported (31, 32) to have been decreasing significantly since the 1980s up to the birth-year period 1995-1998, although this overall trend ceases in the birth-year period 1999-2002 (31–33). In Asia, the overall prevalence of childhood CP increased from 1988 to 1997 and decreased from 1998 to 2007 in Okinawa, Japan (34), which was different from the increasing trend in China. In South Korea, the prevalence of childhood CP has decreased significantly from 2007 to 2103 (35).

The current subgroup analysis showed that the prevalence of CP was significantly higher among males (2.25‰) than females (1.9‰) (*p*-value <0.001). This

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difference between the sexes is consistent with results from previous studies in other populations (35, 36). In addition, this research indicated that the prevalence rate was highest among children in the age group 0–1 years old (1.95‰), while the rate in other groups fluctuated. There is agreement that a threshold age is generally not given for the appearance of symptoms early in life; the great majority of children with CP present with symptoms as infants or toddlers, and the diagnosis of CP is made before the age of 2 years (16). It is also important to note that the rate of spastic CP was the highest type, which was similar to the results of studies in countries other than China (37).

The prevalence of CP varied significantly in different geographical regions. The pooled prevalence of CP was highest in Southwest China, followed by Northwest China and Central China. The prevalence showed a decreasing trend from west to east of China. In addition, urban residents had a much lower prevalence of CP (2.54‰) compared with rural residents (1.9‰). Such differences might be related to unbalanced regional development, such as variability in medical conditions and economic development. In urban areas, pregnant women may be more able to attend prenatal examinations and consultations (38).

The analysis of CP according to birthweight showed, as expected, that the prevalence decreased significantly among children with a birthweight > 2,500 g, which was similar to the trend seen in Japan and other European countries (34, 36, 39). Before the early 1980s, Hagberg (37) found that these very low birthweight children with CP were, in general, severely dyskinesic and frequently had additional serious impairments, such as being seriously multi-impaired (SMR), and having infantile hydrocephalus and epilepsy. Therefore, more attention should be paid to health services for low-birthweight infants and their families.

### Strengths and limitations

This systematic review was performed using a complete search strategy and rigorous inclusion and exclusion criteria, which adhered to 2009 PRISMA guidelines (15). In comparison with previous reviews (14), this review included studies extracted from more databases, a larger sample size and national representativeness, and further subgroup analysis on the heterogeneity was performed. The longitudinal trend in children with CP in China over the 32-year period from 1988 to 2020 was analysed.

This review has several potential limitations. Firstly, some factors could affect the heterogeneity of included studies, including different denominators, study design, diagnostic criteria for CP, genetic background, nutrition levels, etc. Secondly, owing to the lack of a population-based registration system for CP, the nationwide prevalence of CP in China is unknown; the study covered only part of mainland China in evaluating the trend in CP. Thirdly, possible biases in selection and information, sample representativeness, specificity of diagnostic tools, and missing data in this cross-sectional survey might have resulted in underestimation of the pooled results, and, as only Chineseand English-language literature were included, this may also have led to publication bias.

### Conclusion

This study shows a trend of continuous increase in the prevalence of CP in children and adolescents over the 32-year period from 1988 to 2020 in China, especially since 2008. Multiple counter-measures should be taken to reduce the prevalence of CP in China. Pregnant women should be provided with prenatal examinations, consultations and health education to raise awareness of CP, especially in northern China and rural inland areas. Furthermore, it is essential to strengthen the health monitoring and management of infants and young children. Therefore, more research is necessary among children and adolescents with CP.

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### **REFERENCES**

- van Lieshout P, Candundo H, Martino R, Shin S, Barakat-Haddad C. Onset factors in cerebral palsy: a systematic review. Neurotoxicology 2017; 61: 47–53.
- Wichers M, Hilberink S, Roebroeck ME, van Nieuwenhuizen O, Stam HJ. Motor impairments and activity limitations in children with spatic cerebral palsy: a Dutch populationbased study. J Rehabil Med 2009; 41: 367–374.
- van Eck M, Dallmeijer AJ, van Lith IS, Voorman JM, Becher JG. Manual ability and its relationship with daily activities in adolescents with cerebral palsy. J Rehabil Med 2010; 42: 493–498.
- Hollung SJ, Vik T, Lydersen S, Bakken IJ. Decreasing prevalence and severity of cerebral palsy in Norway among children born 1999 to 2010 concomitant with improvements in perinatal health. Eur J Paediatr Neurol 2018; 22: 1–8.
- Petersen TG, Forthun I, Lange T, Villadsen SF, Nybo Andersen AM, Uldall P, et al. Cerebral palsy among children of immigrants in Denmark and the role of socioeconomic status. Eur J Paediatr Neurol 2019; 233: 507–516.
- Kakooza-Mwesige A, Andrews C, Peterson S, Wabwire Mangen F, Eliasson AC, Forssberg H. Prevalence of cerebral palsy in Uganda: a population-based study. Lancet Glob Health 2017; 5: 1–8.
- 7. Khandaker G, Muhit M, Karim T, Smithers-Sheedy H, Novak I, Jones C, Badawi N. Epidemiology of cerebral palsy in

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Bangladesh: a population-based surveillance study. Dev Med Child Neurol 2018; 61: 1–9.

- Oskoui M, Coutinho F, Dykeman J, Jette N, Pringsheim T. An update on the prevalence of cerebral palsy: a systematic review and meta-analysis. Dev Med Child Neurol 2013; 55: 509–519.
- Lv SF HL, Tian AP, Duan H, Ma YM. [Investigation of neurological diseases in 12489 children in five cities and counties of Sichuan Province.] Hua Xi Yi Xue Za Zhi 1993; 8: 204–205 (in Chinese).
- GBD 2019 Mortality and Causes of Death 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: 1135–1159.
- 11. Wang B, Chen Y, Zhang J, Li J, Guo Y, Hailey D. A preliminary study into the economic burden of cerebral palsy in China. Health Policy 2008; 87: 223–234.
- 12. Li S, Lin Q, Liu JM, Zhang SX, Hong SX, Jiang MF, et al. [Prevalence of childhood cerebral palsy in six provinces in China.] Zhonghua Yi Xue Za Zhi 2001; 81: 1220–1223 (in Chinese).
- 13. Li XJ, Qiu HB, Jiang ZM, Pang W, Guo J, Zhu LL, et al. [Epidemiological characteristics of infantile cerebral palsy in 12 provinces and cities of China.] Zhonghua Shi Yong Er Ke Lin Chuang Za Zhi 2018; 33: 378–383 (in Chinese).
- Qi MM, Lai XH, Li ZK, et al. Meta analysis of the prevalence of cerebral palsy in children in China Xun Zheng Hu Li 2015; 2: 63–67.
- Moher D, Shamseer L, Clarke M, Ghersi D, Liberati A, Petticrew M, et al. Preferred reporting items for systematic review and meta-analysis protocols (PRISMA-P) 2015 statement. Syst Rev 2015; 4: 1.
- Li X. [Proceedings of the 6th National Child Rehabilitation, 13th National Conference on the Health of Pediatric Cerebral Palsy and International academic Exchange Conference.] 2014, Apr 11–13; Henan, China. China: CNKI; 2014 (in Chinese).
- Rostom A, Dubé C, Cranney A, Saloojee N, Sy R, Garritty C, et al. Celiac disease. Evid Rep Technol Assess (Summ) 2004; 6: 1–6.
- Higgins JP, Thompson SG. Quantifying heterogeneity in a meta-analysis. Stat Med 2002; 21: 1539–1558.
- Trønnes H, Wilcox AJ, Lie RT, Markestad T, Moster D. Risk of cerebral palsy in relation to pregnancy disorders and preterm birth: a national cohort study. Dev Med Child Neurol 2014; 56: 779–785.
- Reddihough D, Collins K. The epidemiology and causes of cerebral palsy. Aust J Physiother 2003; 49: 7–12.
- Gong P, Liang S, Carlton EJ, Jiang Q, Wu J, Wang L, et al. Urbanisation and health in China. Lancet 2012; 379: 843–852.
- 22. Spruill TM. Chronic psychosocial stress and hypertension. Curr Hypertens Rep 2010; 12: 10–16.
- Xue J, Chen LZ, Xue L, Zhou Q. [Meta-analysis of risk factors for childhood cerebral palsy during pregnancy.] Zhongguo Dang Dai Er Ke Za Zhi 2013; 15: 535–540 (in Chinese).
- 24. Rajab A, Yoo S, Abdulgalil A, Kathiri S, Ahmed R, Mochida G, et al. An autosomal recessive form of spastic cerebral palsy (CP) with microcephaly and mental retardation. Am J Med Genet C Semin Med Genet 2006; 140: 1504–1510.
- Alastair H. MacLennan SCT, Jozef Gecz. Cerebral palsy: causes, pathways, and the role of genetic variants. Am J Obstet Gynecol 2016; 214: 670–671.
- Korzeniewski S, Slaughter J, Lenski M, Haak P, Paneth N. The complex aetiology of cerebral palsy. Nat Rev Neurol 2018; 14: 528–543.
- Zeng Y, Hesketh T. The effects of China's universal twochild policy. Lancet 2016; 388: 1930–1938.
- Li HT, Xue M, Hellerstein S, Cai Y, Gao Y, Zhang Y, et al. Association of China's universal two child policy with changes in births and birth related health factors: national,

descriptive comparative study. BMJ 2019; 366: 4680.

- Goldenberg RL, Culhane JF, Iams JD, Romero R. Epidemiology and causes of preterm birth. Lancet 2008; 371: 75–84.
- Elodie Sellier SM, Smithers-Sheedy H, Platt MJ. European and Australian cerebral palsy surveillance networks working together for collaborative research. Neuropediatrics 2020; 51: 105–112.
- Himmelmann K, Hagberg G, Uvebrant P. The changing panorama of cerebral palsy in Sweden. X. Prevalence and origin in the birth-year period 1999–2002. Acta Paediatr 2010; 99: 1337–1343.
- Himmelmann K, Hagberg G, Beckung E, Hagberg B, Uvebrant P. The changing panorama of cerebral palsy in Sweden. IX. Prevalence and origin in the birth-year period 1995–1998. Acta Paediatr 2005; 94: 287–294.
- Hagberg B, Hagberg G, Beckung E, Uvebrant P. Changing panorama of cerebral palsy in Sweden. VIII. Prevalence and origin in the birth year period 1991–1994. Acta Paediatr 2001; 90: 271–277.
- 34. Touyama M, Touyama J, Kinjo Y. Trends in the prevalence and characteristics of unilateral spastic cerebral palsy in patients born between 1988 and 2007 in Okinawa, Japan. No To Hattatsu 2017; 49: 11–14.
- Kim SW, Jeon HR, Shin JC, Youk T, Kim J. Incidence of cerebral palsy in Korea and the effect of socioeconomic status: a population-based nationwide study. Yonsei Med J 2018; 59: 781–786.
- 36. Sellier E, Platt MJ, Andersen GL, Krägeloh-Mann I, De La Cruz J, Cans C. Decreasing prevalence in cerebral palsy: a multi-site European population-based study, 1980 to 2003. Dev Med Child Neurol 2016; 58: 85–92.
- Smithers-Sheedy H, McIntyre S, Gibson C, Meehan E, Scott H, Goldsmith S, et al. A special supplement: findings from the Australian Cerebral Palsy Register, birth years 1993 to 2006. Dev Med Child Neurol 2016; 58: 5–10.
- Hao P, Han X, You L, Zhao Y, Yang L, Liu Y. Maternal health services utilization and maternal mortality in China: a longitudinal study from 2009 to 2016. BMC Pregnancy Childbirth 2020; 20: 220.
- 39. Hagberg B, Hagberg G, Olow I, von Wendt L. The changing panorama of cerebral palsy in Sweden. V. The birth year period 1979–1982. Acta Paediatr Scand 1989; 78: 283–290.
- 40. Ma HZ, Ding XH, Guo SY, Ma DM, Han SZ. [Study on the incidence and etiology of infantile cerebral palsy in Yinchuan, Ningxia.] Ningxia Yi Xue Za Zhi 1996; 18: 35 (in Chinese).
- Li S, Hong SX, Zhao P, Chen X, Yang L, Huang JJ, et al. [Investigation report on infantile cerebral palsy in Haiyan Heping Lake.] Zhongguo You Sheng You Yu Za Zhi 1996; 7: 97–99 (in Chinese).
- 42. Song Y, Gu GN. [Epidemiological investigation of cerebral palsy in 6-year-old children in Suzhou.] Suzhou Da Xue Xue Bao Za Zhi 2002; 22: 404–405 (in Chinese).
- 43. Li S, Hong SX, Wang TM, Liu HL, Zhao FL, Lin Q, et al. [Premature, low birth weight, small for gestational age and childhood cerebral palsy.] Zhonghua Er Ke Za Zhi 2003; 41: 344–347 (in Chinese).
- Li ZX, Zhao SJ, Zhou J, Chen TT. [Epidemiological investigation of cerebral palsy in 30 000 scattered children aged 6 years old in Jilin area.] Xian Dai Kang Fu Za Zhi 2000; 4: 846–847 (in Chinese).
- 45. Liu J, Li Z, Lin Q, Zhao P, Zhao F, Hong S, et al. Cerebral palsy and multiple births in China. Int J Epidemiol 2000; 29: 292–299.
- Xie GP, Li X, Chen L, Xu HA. [Epidemiological investigation and analysis of children with cerebral palsy in community.] Zhongguo Fu You Bao Jian 2005; 20: 31–32 (in Chinese).
- 47. Wang HJ, Yang XL. [Preliminary analysis of the prevalence and related risk factors of cerebral palsy in children aged 0-6 years old.] Zhongguo She Qu Yi Xue Za Zhi 2002; 8: 15–16 (in Chinese).

#### p. 10 of 10 S. Yang et al.

- Zhang QH, Chen YJ, Shen XY. [Epidemiological investigation and related factors analysis of cerebral palsy in 0-6-year-old children in Wujin city.] Zhongguo Fu You Bao Jian Za Zhi 2000; 10: 639–641 (in Chinese).
- 49. Li GY, Huang YX, Liu W, Ning CJ, Zhou L, Pan CH. [Investigation report on epidemiology of cerebral palsy in children aged 1 to 6 years in Wucheng District of Nanning City.] Yu Fang Yi Xue Za Zhi 2000; 22: 213–215 (in Chinese).
- 50. Zhao ZY, Zhou XJ, Shui QX, Chen K, Le YL, Zheng KH. [Investigation on the prevalence of cerebral palsy in children in Zhejiang Province.] Zhejiang Yu Fang Yi Xue Za Zhi 2002; 14: 8–9 (in Chinese).
- 51. Liang YL, Guo XM, Yang GG, Yan XM, Li XL, Li GY, et al. [Study on the prevalence of cerebral palsy in 1-year-old and 6-year-old children in Guangxi.] Zhonghua Yu Fang Yi Xue Za Zhi 2002; 36: 21–23 (in Chinese).
- 52. Tao FM, Ni JH, Zhang HB, Fu J, Yin HP, Xu Z, et al. [Epidemiological study on the current situation and regional distribution of cerebral palsy in children aged 1 to 8 years old in rural areas of Anhui Province.] Zhongguo Gong Gong Wei Sheng Za Zhi 1999; 15: 3–5 (in Chinese).
- 53. Zhong YQ, Wu F, Wu KM, Wen RK, Hou GF, Peng DZ, et al. [A case-control study on risk factors of 308 children with cerebral palsy in Leshan, Sichuan.] Zhonghua Yu Fang Yi Xue Za Zhi 2002; 36: 323–325 (in Chinese).
- 54. Guo JX, Liu SW, Zhu Y, Feng XY, Wang TM, S. L. [Study on the prevalence and intervention countermeasures of cerebral palsy in children aged 1–6 years old in Gansu province.] Zhongguo Fu You Bao Jian Za Zhi 2003; 18: 418–419 (in Chinese).
- 55. Chen Y, Ni JF, Zhang JP, Dai W, Tao FB. [Study on the status of cerebral palsy in children aged 2 and 13 years old in Shushan Town, Hefei.] Anhui Yi Ke Da Xue Xue Bao Za Zhi 2000; 35: 38–39 (in Chinese).
- 56. Yan XM, Su LH, Lan DP, Liang YL, Guo XM, Yang GL, et al. [Analysis of prevalence and related factors of cerebral palsy in children in Hengxian County.] Zhongguo You Sheng You Yu Za Zhi 2002; 13: 11–13 (in Chinese).
- 57. Xing BQ, Wang DM, Long HF, Liu YH. [Epidemiological investigation and analysis of cerebral palsy in children in Gannan County.] Qiqi Haer Yi Xue Yuan Xue Bao Za Zhi 1999; 20: 3–5 (in Chinese).
- Sun X, An MX, Xu YT. [Investigation and analysis on the prevalence of cerebral palsy in 22 villages and towns in Rizhao area.] Zhongguo You Sheng Yu Yi Chuan Za Zhi 2001; 9: 115–116 (in Chinese).
- Tao FB. [Study on epidemiology and early screening methods of Cerebral Palsy in Children master's thesis [dissertation].] Shanghai: Fudan University; 2001 (in Chinese).
- 60. Shi RF, Tang HX, Wang KL, Zhang YQ, Cheng RG, Yang GR, et al. [A sample survey was conducted on the risk factors of cerebral palsy in 95435 children.] Lin Chuang Yan Jiu Za Zhi 2004; 1: 434–435 (in Chinese).
- 61. Tao FB, Xu J, Deng GZ, Liu ZF, Xu YQ, Yin HP, et al. [ A study on the prevalence of cerebral palsy in children aged 1 to 8 years old in two cities of Anhui Province.] Zhong-guo Gong Gong Wei Sheng Za Zhi 2002; 18 (in Chinese).
- 62. Gao L, Meng Y, Zhao SY, Zhou CJ, Yuan CZ, Wang YH, et al. [Epidemiological investigation and analysis of children with cerebral palsy in Henan province.] Zhongguo Shi Yong Er Ke Za Zhi 2003; 18: 464–467 (in Chinese).
- 63. Yuan HB, Cheng LY, Wang L, Huang X. [Differences in pre-

valence and high risk factors of cerebral palsy in children between urban and rural areas.] Zhongguo Lin Chuang Kang Fu Za Zhi 2005; 9: 160–161 (in Chinese).

- 64. Wang YY, Cai LR, Zhou YA, Huang YL, Chen LN, Gao YY, et al. [Epidemiological investigation of cerebral palsy in children in Quanzhou City, Fujian Province.] Zhonghua Liu Xing Bing Xue Za Zhi 2005; 26: 832 (in Chinese).
- 65. Tseng SH, Lee JY, Chou YL, Sheu ML, Lee YW. Association between socioeconomic status and cerebral palsy. PLoS One 2018; 13: e0191724.
- 66. Li HM. [Investigation and analysis of cerebral palsy in children aged 0–6 years old in Zhongliangshan area.] Zhong Wai Yi Liao Za Zhi 2010; 24: 65 (in Chinese).
- 67. Tan Y, Zhang R, Qiu HY. [Investigation and analysis of cerebral palsy in children in Fuling district.] Chongqing Yi Xue Za Zhi 2011; 40: 1510–1511 (in Chinese).
- 68. Hu JH, Zhang HJ, Qin R, Yan H, Wang PQ, Pang W, et al. [Epidemiological investigation and analysis of cerebral palsy in children aged 1–6 years old in Hunan province.] Zhongguo You Sheng Yu Yi Chuan Za Zhi 2014; 24: 120–123 (in Chinese).
- 69. Ma JQ. [Analysis of risk factors of cerebral palsy in children in Henan Province master's thesis.] Zhengzhou(HN): Zhengzhou Da Xue 2014 (in Chinese).
- Song SY. [Epidemiological investigation of cerebral palsy in children aged 1–6 years in Xinxiang area, Henan Province.] Zhonghua Shi Yong Er Ke Lin Chuang Za Zhi 2014; 9: 451–454 (in Chinese).
- 71. Li L. [Analysis of cerebral palsy in children aged 1–6 years old in rural areas of Zaozhuang master's thesis [dissertation].] Changchun(JL): Jilin University; 2012 (in Chinese).
- 72. Li Z, Wang XJ, Yang FW, Lv ZH, Yao SP, Li YL, et al. [Epidemic characteristics and standardized prevention and treatment of cerebral palsy in children in Qinghai province.] Zhonghua Shi Yong Er Ke lin Chuang Za Zhi 2017; 32: 374–376 (in Chinese).
- Yuan J, Wang J, Ma J, Zhu D, Zhang Z, Li J. Paediatric cerebral palsy prevalence and high-risk factors in Henan province, Central China. J Rehabil Med 2019; 51: 47–53.
- Peng GL, Zhang DX, Hu CX, Cai SY, Wang YD. [Epidemiological investigation of cerebral palsy in 0-5-year-old children in Xiamen.] Zhongguo Fu You Bao Jian Za Zhi 2016; 31: 5456–5458 (in Chinese).
- 75. Lin WY, Yang Y, Tu WL. [Epidemiological investigation of cerebral palsy in children aged 1 to 6 years old in Yili Prefecture, Xinjiang.] Zhongguo Er Tong Bao Jian Za Zhi 2005; 23: 1188–1191 (in Chinese).
- 76. Wang F, Huang XP, Xiong LY. [Investigation and analysis on the prevalence and clinical types of cerebral palsy in children in EZhou.] Yi Xue Xin Xi Za Zhi 2016; 29: 127–128 (in Chinese).
- 77. Zhang ML, Liu ZH, Cai SY, Zhou Y, Li LY, Zheng CY, et al. [A sampling survey on the prevalence of cerebral palsy in children in Foshan, Guangdong.] Shi Yong Yi Xue Za Zhi 2016; 32: 2775–2776 (in Chinese).
- Ii L, Jia QZ, Sun Y, Lv Y, Li XL, Xu LC, et al. [Epidemiological investigation of 6-year-old children with cerebral palsy in Hainan Province.] Zhongguo Kang Fu Yi Xue Za Zhi 2018; 33: 376–379 (in Chinese).
- Luo ZX, Cai SZ, Shi L, Huang D, Lu YH, Cilen DJ, et al. [Epidemiological investigation of cerebral palsy in Maizhokunggar.] Xizang Medicine 2020; 41: 89–91 (in Chinese).