Developmental anomalies and associated impairments in Saudi children with cerebral palsy: a registry-based, multicenter study

Saeed Al-Garni¹, Shereen Derbala², Hatem Saad³ and Ahmed I. Maaty^{4,5*}

Abstract

Background: There are few epidemiological data to support rehabilitation programs for cerebral palsy (CP). Scarce international studies described the developmental anomalies (DAs) among children with CP. To our knowledge, the Arab countries did not publish data regarding this topic. This study aimed to describe the percentage of DAs among children with CP and detect the association between clinical subtypes and impairment severity in children with various DAs. We collected registry data of 679 children with cerebral palsy, between 2014 and 2019, from Armed Forces Hospitals, Taif, Kingdom of Saudi Arabia (KSA). We recorded demographic, perinatal, postnatal, developmental anomalies, subtypes, and impairment characteristics. We utilized the chi-square test to calculate the differences between groups.

Results: We reported significant differences between the children with and without anomalies regarding the percentages of consanguinity, preterm labor, low birth weight, and neonatal intensive care unit admission (P = 0.001, 0.002, 0.003, 0.005, respectively). Congenital dysplasia of the hip and hydrocephalus was the most frequent skeletal and nervous anomalies among children with DAs (19.1% and 12.8%, respectively). The spastic bilateral pattern was significantly higher among children with skeletal anomalies than the central nervous system/other groups (P < 0.001). The nervous anomalies group had higher frequencies of severe intellectual, motor, speech, and visual disabilities and a higher percentage of seizures than all other groups.

Conclusions: The frequency of children with anomalies in this study was comparable to previous studies. Children with CP and nervous system anomalies had more severe motor disabilities and associated impairments.

Keywords: Congenital anomalies, Motor dysfunction, Intellectual disability, Visual and hearing impairments

Background

Cerebral palsy (CP) is a collective group of permanent, non-progressive, developmental disorders affecting body movement, balance, and posture. It remains a worldwide common cause of pediatric morbidity and disability despite the scientific advances in neonatal and maternal care over the last decades. Children with CP had motor

* Correspondence: ahmed_maaty@med.suez.edu.eg; aimaaty@hotmail.com ⁴Physical Medicine, Rheumatology, and Rehabilitation, Faculty of Medicine, Suez Canal University, Ismailia, Egypt

⁵Physical Medicine and Rehabilitation, Armed Forces Center of Health Rehabilitation, Taif, Saudi Arabia

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

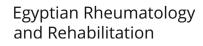
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dysfunction often accompanied by sensory, perceptive, cognitive, communicating, learning, and behavioral disorders. Congenital or acquired musculoskeletal (MSK) complications are usually present [1, 2].

To date, CP and other developmental disorders have received insufficient attention in developing countries. Epidemiological evidence of pediatric disabilities, especially CP, in KSA is scarce [3]. In a community-based study, the prevalence of CP in KSA was 2.3/1000 population [4]. Another Saudi study covered 99,788 live births in Riyadh Military Hospital has reported a CP incidence of 0.41% [5]. Along with other neurological disorders of children, healthcare providers have not given CP high

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priority amongst health problems. Inadequate perinatal and postnatal care in developing countries makes it sensible to believe that CP incidence is much higher than in developed countries [6, 7].

Several risk factors associated with CP include multiple pregnancies, intrauterine growth restriction, preterm labor, neonatal asphyxia, and developmental anomalies (DAs) [8]. DAs involved wide-ranging abnormalities of body structure or function present at birth. These anomalies were due to genetic, teratogenic, or nutritional factors, or a combination of several factors. Studies showed that DAs were more common in children with CP than in the general pediatric population [9, 10]. The DAs were mostly in the central nervous system (CNS) across all Gross Motor Function Classification System (GMFCS) levels, and the presence of DA itself correlated with severity of CP. The genetic evaluation showed that 5q21.1 locus deletion linked with some CNS anomalies such as corpus callosum hypoplasia and microcephaly [11]. Thus, parents must obtain proper genetic counseling [12, 13].

Researchers reported that many risk factors of DAs included diabetes mellitus (DM), thyroid dysfunction, and drug toxicity. Maternal DM significantly increased the risk of DAs, including the CNS (microcephaly and spina bifida) and the skeletal system (sacral agenesis and limb defects) [14]. Moreover, the prevalence of DAs and Down syndrome were significantly higher among patients with primary hypothyroidism than the general population. Japanese patients had a higher frequency of cardiovascular anomalies than American and Egyptian patients, and a female predominance was also observed [15]. Previous experiments reported several medications, such as anti-epileptics, anti-convulsants, and antihypertensive, implicated in neural tube defects, cardiovascular, gastrointestinal, and urinary tract DAs [16].

Aim of the study

In this study, we aimed to describe the percentage of DAs among children with CP and detect the association between clinical subtypes and impairment severity in children with various DAs.

Methods

Study design and setting

In this multicenter observational analysis, we extracted the data, between 2014 and 2019, from Military Hospitals in Taif City, KSA. These hospitals included Armed Forces Rehabilitation Center, Alhada Hospital for Armed Forces, Prince Mansour Hospital for Society, and Prince Sultan Hospital for Armed Forces. The Wipro software system contained the registered pediatric population with CP. We used these registry-based medical archives to differentiate children with CP from other recorded patients.

Patients

We reviewed 679 children with CP aged between 8 months and 17.8 years of both genders. We excluded the children with other neurological diagnoses or unconfirmed diagnosis of CP. Our definition of CP was the same as the definition used by European and Australian registries [17, 18]. Before registration, a multi-disciplinary team performed comprehensive evaluations to confirm the diagnosis and document the children's disabilities. This team involved a physiatrist, audiologist, ophthalmologist, orthotist/prosthetist, dietician, psychologist, social counselor, physical, occupational, and speech therapists. After initial assessments, we subdivided the patients into children with CP and DAs (n = 188) and children with CP and without DAs (n = 491).

Study variables

The collected data included age at registration/years, registered year, gender (male/female), consanguinity (no/ yes), mode of delivery (vaginal delivery/cesarean section), preterm defined as babies born alive < 37 weeks or 259 days gestation [19] (no/yes), low birth weight (LBW) defined as born weighing between 1500 and 2499 g [19] (no/yes), and admission to neonatal intensive care unit (NICU) (no/yes).

We classified DAs according to the World Health Organization (WHO)/the International Classification of Diseases version 11 (ICD-11) [19]. The skeletal anomalies included developmental dysplasia of the hip, congenital scoliosis, and pes planus. Reduction defects of the lower limb included femoral, tibial, or fibular hypoplasia (unilateral and bilateral). We reported the congenital deformities of the feet that involve talipes equinovarus, metatarsus varus, talipes calcaneovalgus, and hallux valgus. The nervous system anomalies included congenital hydrocephalus, microcephaly, corpus callosum agenesis, and spina bifida/ myelomeningocele). The European Surveillance of Congenital Anomalies (EUROCAT) defined microcephaly as a head circumference > 3 standard deviations (SD) below the mean [20]. We also documented the ocular (cataract), the respiratory (tracheomalacia), the circulatory (unspecified), the digestive, the genitourinary, and the chromosomal anomalies. According to the Surveillance of Cerebral Palsy in Europe (SCPE), we divided the clinical subtypes of CP into spastic unilateral or bilateral, dyskinetic, and ataxic [21].

The psychologist classified intellectual disability into mild (intelligence quotient [IQ] 50–69), moderate (IQ 36–49), severe (IQ 20–35), and profound (IQ < 20), according to the Diagnostic and Statistical Manual of Mental Disorders 5th edition (DSM-5) [22]. According

to the GMFCS, we graded the motor impairment into mild (GMFCS levels I and II), moderate (GMFCS level III), and severe (GMFCS levels IV and V) (Appendix 1) [23, 24]. According to the American Speech-Language-Hearing Association, the speech therapist categorized speech disabilities into normal speech, mildly indistinct speech, moderately indistinct speech, severely indistinct speech, and mute (no speech) [25]. The ophthalmologist categorized visual acuity (VA) into normal, mild impairment (VA < 6/12), moderate impairment (VA < 6/18), severe impairment (VA < 6/60), and blind (VA < 3/60) using ICD-11 [19]. The audiologist defined the degree of hearing loss as normal (-10-15 decibel [dB]), minor (16–25 dB), mild (26–40 dB), moderate (41–55 dB), moderately severe (56-70 dB), severe (71-90 dB), and profound (> 91 dB) using pure-tone audiometry [26]. We documented seizures as present if they occurred at least once (absent/present).

Statistical analysis

We performed the statistical analysis using Statistical Package for Social Sciences (SPSS), version 23.0 (IBM SPSS^{*}, Statistics 23, Armonk, NY, USA). We compared the data regarding demographic, perinatal, postnatal, clinical subtypes, and associated impairments. We applied the chi-square or Fisher's exact tests to determine the differences between groups. We set the significant level at a probability (*P*) value < 0.05.

Results

We evaluated 679 children (386 males, 293 females) with CP. The mean age (SD) at registration was 7.8 (4.6) years. We reported significant differences between children with and without DAs regarding the prevalence of consanguinity (57.4% versus 13.2%, respectively), preterm labor (27.7% versus 17.0%, respectively), LBW (13.3 versus 6.3%, respectively), and admission to NICU (17.0% versus 9.4%, respectively). Age, gender, registered year, and mode of delivery showed insignificant differences between both groups (P > 0.05), as shown in Table 1. Congenital subluxation/dysplasia of the hip and hydrocephalus were the most frequent skeletal and nervous anomalies (19.1% and 12.8%, respectively). Twelve patients (6.4%) of the DAs group had Down syndrome, and 7.0% of all anomalies occurred in other systems (Table 2).

Children with skeletal DAs had higher unilateral spastic patterns than children without DAs and higher bilateral spasticity than children with nervous and other DAs. Meanwhile, no one of them had dyskinetic or ataxic subtypes. CP with nervous DAs had more severe intellectual, motor, speech, and visual disabilities than other groups (Table 3). Interestingly, patients with chromosomal anomalies (including Down syndrome) and nervous anomalies had near proportions of severe hearing impairment (9.7% and 9.0%, respectively). Seizures were significantly higher among the nervous anomalies' group and significantly lower among the chromosomal/other anomalies group than the nonanomaly group (P = 0.003 and 0.02, respectively). The children with nervous anomalies had higher frequencies of intellectual, visual, and hearing impairments than the skeletal DAs group (P < 0.001, < 0.001, and 0.002, respectively). Motor and speech dysfunctions showed insignificant differences between skeletal and nervous systems groups, as demonstrated in Table 3.

Discussion

In the last decade, register-based studies have shown an increased interest in estimating DAs incidence among children with CP [27, 28]. However, no previous studies in KSA, to our knowledge, investigated the frequency of DAs in the CP population. So, we chose Taif city as a represented region of the Saudi people as it consisted of urban, rural, and sub-rural inhabitants. This study was the first Saudi, registry-based, multicenter research to describe DAs' rate among CP children.

The prevalence of DAs among our CP children was 27.7%. The incidence of anomalies among children with CP varied substantially among different populations with a range between 11% and 40% [28-36]. The percentage of children with CP and DAs in our study (27.7%) was similar to Australian Cerebral Palsy Registry (28%) [18], higher than Jystad et al. (25%) [30], Rankin et al. (15%) [28], Pharoah (11%) [37], Garne et al. (12%) studies [17], and lower than described by Blair et al. (32%) [32]. Racial, ethnic, and sociocultural variations between studied populations explained the discrepancy in these studies. Besides, this inconsistency may be due to different inclusion criteria of the patients. Compared to our research, Blair et al. [32] only included minor DAs. Pharoah [37] and Garne et al. [17] depended on data from CP registries. Rankin et al. [28] utilized data from CP and DA registries.

The epidemiological figures of CP and DAs varied significantly worldwide, especially between low-, middle-, and high-income countries [38]. Considerable variations in the live birth rate of anomalies between nations were due to differences in prenatal screening and the proportion of pregnancy terminations [39, 40]. A higher proportion of pregnancy termination in women diagnosed with significant fetal DAs might be one reason clarifying the reported declines in CP and DAs rates [41, 42]. The interaction between genetic and environmental factors that differed among populations could also cause these variations [43].

In our studied population, the mean age at registration was about 8.0 years. This delayed registration for

Variables	Children with DA ($n = 188$)	Children without DA ($n = 491$)	Р	
Age at registration (years), n (%)				
< 1	2 (1.1%)	11 (2.2%)	0.40	
1-	28 (14.9%)	52 (10.6%)		
3–	41 (21.8%)	116 (23.6%)		
5—	19 (10.1%)	55 (11.2%)		
7–	25 (13.3%)	76 (15.5%)		
9–	29 (15.4%)	59 (12.0%)		
11-	15 (8.0%)	43 (8.8%)		
13–	12 (6.4%)	19 (3.9%)		
15+	17 (9.0%)	60 (12.2%)		
Registered year, n (%)				
2014	9 (4.7%)	21 (4.3%)	0.12	
2015	57 (30.3%)	107 (21.8%)		
2016	32 (17.1%)	119 (24.2%)		
2017	28 (15.0%)	61 (12.4%)		
2018	23 (12.2%)	73 (14.9%)		
2019	39 (20.7%)	110 (22.4%)		
Gender, <i>n</i> (%)				
Male	117 (62.2%)	269 (54.8%)	0.08	
Female	71 (37.8%)	222 (45.2%)		
Consanguinity, <i>n</i> (%)				
No	80 (42.6%)	426 (86.8%)	0.001*	
Yes	108 (57.4%)	65 (13.2%)		
Mode of delivery, n (%)				
Vaginal delivery	128 (68.0%)	322 (65.6%)	0.14	
Caesarean section	60 (32.0%)	169 (34.4%)		
Preterm, n (%)				
No	136 (72.3%)	408 (83.0%)	0.002*	
Yes	52 (27.7%)	83 (17.0%)		
Low birth weight, n (%)				
No	163 (86.7%)	460 (93.7%)	0.003*	
Yes	25 (13.3%)	31 (6.3%)		
Admission to NICU, n (%)				
No	156 (83.0%)	445 (90.6%)	0.005*	
Yes	32 (17.0%)	46 (9.4%)		

Table 1 Demographic, perinatal, and postnatal data among children with cerebral palsy (CP)

CP cerebral palsy, DA developmental anomalies, NICU neonatal intensive care unit Significant P value

rehabilitation impaired the outcome of CP. The effectiveness of rehabilitative interventions of CP patients improved by early referral for physical therapy [44]. Maximal neural plasticity occurred during early childhood, where the therapeutic strategies had tremendous potential for long-term efficacy [45]. Moreover, early rehabilitation can prevent or modest the complex social, communication, and emotional associations that can have functional implications in later life [46]. We matched both groups (CP with and without DAs) regarding age at registration and gender (p > 0.05). So, these variables had an equal distribution among both groups, eliminating the impact of confounding variables on the study outcome [47]. Children with DAs had a higher consanguinity rate than children without DAs (13.3% versus 6.3%, respectively). The consanguinity

Table 2 Developmental anomalies (DAs) among children with cerebral palsy (CP)

	Children with DAs ($n = 188$)
DAs of the skeleton ($n = 101$)	
Congenital subluxation/dysplasia of hip	36 (19.1%)
Congenital scoliosis	20 (10.6%)
Congenital deformities of feet	16 (8.5%)
Congenital pes planus	14 (7.4%)
Reduction defects of lower limb(s)	11 (5.9%)
Multiple anomalies	4 (2.1%)
DAs of the nervous system ($n = 56$)	
Congenital hydrocephalus	24 (12.8%)
Microcephaly	17 (9.0%)
Corpus callosum agenesis	8 (4.3%)
Spina bifida/myelomeningocele	7 (3.7%)
Chromosomal anomalies ($n = 18$)	
Down syndrome	12 (6.4%)
Other syndromes	6 (3.2%)
Other systems $(n = 13)$	
Circulatory system	5 (2.7%)
Genitourinary system	3 (1.6%)
Eye	2 (1.1%)
Respiratory system	2 (1.1%)
Digestive tract	1 (0.5%)

rates in KSA and other Arab countries are significantly higher than in the Americans, Europeans, South-Africans, Eastern-Asians, and Oceanic countries [48]. The prevalence of consanguinity in the Saudi population is approximately 56% [49]. Consanguinity played an essential role in the etiopathogeneses of CP and DAs [10, 50–52]. Other significant differences between children with and without DAs included higher preterm labor, LBW, and NICU admission rates. Our findings confirmed the observations of earlier studies regarding CP/ DAs risk factors [45, 53–56]. We found that the delivery mode was not significantly different between both groups, which was the same observation in the Turkish population [57].

In our study, most DAs presented in the skeletal system (53.7%), and about 30% appeared in the nervous system. This observation contrasted with prior studies where the CNS anomalies were the most common DAs [30, 36]. This result most likely reflected a referral pattern rather than a difference in epidemiology. Physicians referred CP children, to our rehabilitation center for physical therapy and functional rehabilitation, mainly due to MSK dysfunction. CP rehabilitation goals were primarily to minimize MSK complications rather than treating the central neurological deficit [44].

We can easily say that the most common DAs in our patients affecting the MSK and CNS. These findings are comparable to previous hospital-based studies [36, 58, 59]. Skeletal and nervous anomalies are widespread anomalies since they are physically apparent, and parents seek early medical advice [58, 59]. Out of 188 patients with DAs, only 5 (2.7%) had cardiovascular DAs, much lower than other studies (10-14%) [28, 30]. Survival bias explained these findings—this type of selection bias disproportionate the survivors of a particular outcome and ignored the non-survivors. Focusing on the survivors resulted in a false-negative estimate of probability [60]. Unfortunately, our results may underestimate the actual incidence of DAs among the CP population.

Our data revealed that congenital subluxation and dysplasia of the hip account for 19.1% of all anomalies in CP patients with DAs. Hydrocephalus was the most recorded DAs of the nervous system (12.8%). The muscle weakness affecting the limb positioning in utero increased the risk of congenital hip affection in CP children. Likewise, the detectable risk of hydrocephaly is the consequence of prenatal brain affection. The cooccurrence of these DAs with CP approved that cerebral dysfunction happens during fetal growth. The brain impairment led to neuronal migration abnormalities,

Variables	CP with skeleton DAs (n = 101)	CP with nervous DAs (<i>n</i> = 56)	CP with other DAs (<i>n</i> = 31)	CP without DAs (n = 491)	P1	P2	P3	P4	P5
Clinical subtypes									
Spastic unilateral	21 (20.8%)	14 (25.0%)	16 (51.6%)	69 (14.0%)	< 0.001*	0.02*	0.01*	< 0.001*	0.002*
Spastic bilateral	80 (79.2%)	36 (64.3%)	15 (48.4%)	381 (77.6%)					
Dyskinetic	0 (0.0%)	6 (10.7%)	0 (0.0%)	22 (4.5%)					
Ataxic	0 (0.0%)	0 (0.0%)	0 (0.0%)	19 (3.9%)					
Impairments									
Intellectual disability									
None	76 (75.2%)	22 (39.4%)	18 (58.1%)	278 (56.7%)	< 0.001*	0.001*	0.0004*	0.98	< 0.001*
Mild-moderate	14 (13.9%)	16 (28.5%)	9 (29.0%)	150 (30.5%)					
Severe-profound	11 (10.9%)	18 (32.1%)	4 (12.9%)	63 (12.8%)					
Motor impairment									
Mild	62 (61.4%)	31 (55.3%)	19 (61.3%)	422 (85.9%)	< 0.001*	< 0.001*	< 0.001*	0.001*	0.65
Moderate	10 (9.9%)	8 (14.3%)	4 (12.9%)	20 (4.1%)					
Severe	29 (28.7%)	17 (30.4%)	8 (25.8%)	49 (10.0%)					
Speech impairment									
None	61 (60.5%)	29 (51.8%)	18 (58.1%)	350 (71.3%)	0.001*	0.007*	0.009*	0.01*	0.33
Mild-moderate	23 (22.7%)	12 (21.4%)	9 (29.0%)	55 (11.2%)					
Severe-no speech	17 (16.8%)	15 (26.8%)	4 (12.9%)	86 (17.5%)					
Visual impairment									
None	85 (84.2%)	21 (37.5%)	23 (74.2%)	214 (43.6%)	< 0.001*	< 0.001*	< 0.001*	< 0.001*	< 0.001*
Mild-moderate	12 (11.8%)	29 (51.8%)	5 (16.1%)	273 (55.6%)					
Severe-blind	4 (4.0%)	6 (10.7%)	3 (9.7%)	4 (0.8%)					
Hearing impairment									
None	86 (85.1%)	34 (60.6%)	21 (67.7%)	393 (80.1%)	< 0.001*	0.2	< 0.001*	< 0.001*	0.002*
Mild-moderate	13 (12.9%)	17 (30.4%)	7 (22.6%)	94 (19.1%)					
Severe-profound	2 (2.0%)	5 (9.0%)	3 (9.7%)	4 (0.8%)					
Seizures									
No	77 (76.2%)	29 (51.8%)	28 (90.3%)	349 (71.1%)	0.0007*	0.29	0.003*	0.02*	0.002*
Yes	24 (23.8%)	27 (48.2%)	3 (9.7%)	142 (28.9%)					

Table 3 Cerebral palsy (CP) subtypes and impairments among children with and without developmental anomalies (DAs)

P1 comparison between skeleton DA group, nervous DA group, and no DA group; P2 comparison between skeleton DA group and no DA group; P3 comparison between nervous DA group and no DA group; P4 comparison between other DA group and no DA group; P5 comparison between skeleton DA group and nervous DA group

^{*}Significant *p* value

encephalopathy, and other cerebral pathologies presented clinically as CP [37].

Following former CP studies, we found that spastic CP was the most common clinical subtype (93.1%) [5, 6, 28, 51]. Ataxic CP was the least frequent type, with a prevalence of 2.8%. This result was consistent with Rankin et al. [28] and Taha and Mahdi [51] observations but differed from Garne's study, where the dyskinetic type was more frequent than other types [17]. Our systematic neurodevelopmental evaluation of each patient documented the motor impairment subtypes' spastic, ataxic, or dyskinetic. The clinical features of motor affection were defined for each

extremity differentiating unilateral from bilateral involvement (spastic unilateral or bilateral) [21]. The present study emphasized the results of Jystad and his colleagues [30] that children with skeletal DAs had a higher frequency of spastic CP than other children. Spasticity is a crucial factor in MSK complications of CP because spastic agonist muscles overcome the weaker antagonist muscles causing contractures and deformities. Hip flexors/adductors, hamstrings, gastrocnemius, soleus, and paraspinal muscles are most affected. Spasticity causes several bony and articular changes, including hip subluxation/ dislocation, foot deformities, and scoliosis [61].

We found that children with the DAs of CNS had more severe psychomotor, vision, hearing, speech impairments, and a higher frequency of seizures than other children. Jystad et al. [30] observed that children with CNS-anomalies had more severe motor impairments, impaired speech function, a higher proportion of epilepsy, and profound vision and hearing impairments than children without anomalies. These findings were also in line with Blair et al. [32], who reported a higher risk for severe CP and associated impairments in children with CNS anomalies. Comparably, Manlongat et al. [36] found that poor functional outcomes were more frequent in children with neurological anomalies than in children without neurological anomalies. Non-verbal communication, epilepsy, blindness, and bilateral disorders were observed exclusively in children with neurological DAs.

The WHO recommended implementing community genetics programs to prevent DAs at the population level. This program's policies included neonatal screening for preventable diseases, detecting carriers of common recessive conditions, and removing environmental factors causing these disorders [62]. Successful programs in the Middle East and North/South Africa included Human Health in Africa, the Qatar Genome Program, and the Mexico National Institute of Genomic Medicine [63].

This study had several strengths. A confirmed diagnosis of CP was accomplished by the multi-disciplinary team before registration using a thorough neurodevelopmental assessment. Most of the findings were highly statistically significant (P < 0.001). These results indicated that the observations from data generated by significant tests were not likely to occur randomly or by chance but due to a specific cause [64]. In contrast to Manlongat et al. [36], we collected the data from registry-based medical archives, not from manual medical records. We kept the missing data to the minimum. We identified DAs by clinical examinations, X-rays, neuroimaging, ultrasonography, echocardiography, and genetic testing.

We found some limitations worth to be mentioned. The study did not include all CP children. Hospitals of the Ministry of Health had their cases, which might not refer to us. Additionally, we may not determine some children with mild CP [55]. Furthermore, the retrospective nature of the current study may be one of the potential limitations. However, we treated the disadvantage of retrospective design by matching children with and without DAs for age and gender, which enhanced the equal representation of subjects and reduced selection bias [65].

Conclusions

The proportion of children with DAs in Taif, KSA (27.7%) was comparable to that identified in many other

countries. Children with CP and DAs, mainly nervous system DAs, had more severe motor disabilities and associated impairments. We proposed that DAs must be one of the primary information included in every newly developed CP registry in other regions. These data will accelerate early intervention to minimize the severity of associated disabilities and improve CP patients' quality of life. Further research may provide successful opportunities for primary prevention.

Abbreviations

CNS: Central nervous system; CP: Cerebral palsy; DAs: Developmental anomalies; dB: Decibel; DSM-5: Diagnostic and Statistical Manual of Mental Disorders 5th edition; EUROCAT: European Surveillance of Congenital Anomalies; GMFCS: Gross Motor Function Classification System; ICD-11: International Classification of Diseases version 11; IQ: Intelligence quotient; KSA: Kingdom of Saudi Arabia; LBW: Low birth weight; MSK: Musculoskeletal; NICU: Neonatal intensive care unit; P: Probability; SD: Standard deviation; SCPE: Surveillance of Cerebral Palsy in Europe (SCPE); SPSS: Statistical Package for Social Sciences (SPSS); VA: Visual acuity; WHO: World Health Organization

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Authors' contributions

SA, SD, and AIM initiated the study question, design, and protocol. SD and AIM performed the data collection. All authors agreed both to be personally accountable for the author's contributions and to ensure that questions related to the accuracy or integrity of any part of the work, even ones in which the author was not personally involved, are appropriately investigated, resolved, and the resolution documented in the literature. AIM completed the data management and statistical analysis, and HS revised the draft and approved the final manuscript.

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Availability of data and materials

The authors confirm that the data supporting the findings of this study are available on request.

Ethics approval and consent to participate

This research has been reviewed and accepted by the Directors and the Research Ethical Committee of Armed Forces Hospitals (*Reg. No. H-02-T-078, Ref. REC.T. 2020-461*). We prepared the study according to the principles expressed in the Helsinki Declaration. All parents signed/fingerprinted an informed written consent included the aim and the benefits of the study.

Consent for publication

Not applicable.

Competing interests

The authors declare that they have no competing interests.

Author details

¹Department of Rehabilitation, Armed Forces Rehabilitation Center, Taif, Saudi Arabia. ²Rheumatology and Rehabilitation Department, Faculty of Medicine, Misr University for Science and Technology, Giza, Egypt. ³Rheumatology and Rehabilitation, Al-Azhar Faculty of Medicine, Cairo, Egypt. ⁴Physical Medicine, Rheumatology, and Rehabilitation, Faculty of Medicine, Suez Canal University, Ismailia, Egypt. ⁵Physical Medicine and Rehabilitation, Armed Forces Center of Health Rehabilitation, Taif, Saudi Arabia.

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