Cerebral palsy is a group of persistent movements or postural disturbances secondary to a nonprogressive lesion in a developing brain, and is considered the main cause of motor disability in children, affecting 1 in 500-1000 newborns. It is the most common cause of severe physical disability in childhood. Spasticity is a common and disabling symptom for many patients with cerebral palsy. Therapy for spasticity is symptomatic with the aim of increasing functional capacity and relieving discomfort. Spasticity treatment by orally administered drugs and intramuscular chemodenervation agents has become more frequent. Most oral medications to treat spasticity have been inadequately studied in children, especially those with cerebral palsy. Since its first use in pediatric patients, reported in 1993, botulinum toxin (BTX-A), a relatively recent addition to the available medical interventions for children with cerebral palsy, has rapidly gained acceptance as a treatment of spasticity. The clinical effects of BTX-A have been reported to include decreased spasticity and increased range of motion. However, no consensus exists among clinicians about how an optimal dose of BTX-A should be determined and there are no standard guidelines on doses of BTX-A in children. Doses of 2-6 U/kg with a maximum total dose of 29 U/kg have been reported. Although there are no standard guidelines on doses in children. The current practice is to inject BTX-A at higher doses than reported in the past. Larger dose of BTX-A is used more frequently, which is considered safe, more effective, and better tolerated by children. Titration of the dose of BTX-A is necessary because muscle spasticity affects different patients in different ways. The dosage of BTX-A must be individualized for each patient.
For the walking child with spastic cerebral palsy, the objective is to counter or minimise the negative long term effect of spasticity and weakness during the period of growth. Simple measures of stretching, walking, and orthotic use can be supplemented by oral muscle relaxants and particularly botulinum toxin type A. The use of intramuscular botulinum toxin to block neuromuscular transmission in spastic muscles has become more popular since its initial use with pediatric patients, reported in 1993. Its prime use has been in calf muscles to overcome dynamic equinus of the ankle in children with hemiplegia or diplegia. The effects of botulinum toxin are transitory and injections must be repeated every 4 to 6 months.

With the advent of botulinum toxin type A (BTX-A) as an adjuvant treatment of patients with equinus foot, several studies have been reported demonstrating that BTX-A can be one of the main tools to improve prognosis in children with cerebral palsy. Botulinum toxin, a relatively recent addition to the available medical interventions for children with cerebral palsy, has rapidly gained acceptance as a treatment that temporarily reduces focal muscle spasticity. The clinical effects of BTX-A have been reported to include decreased spasticity and increased range of motion. These effects may be critically important when considering the influence of spasticity, limited muscle length, and restricted range of motion on the growing bones of young children. Controlled studies, demonstrating the efficacy of BTX-A in treatment of childhood cerebral palsy have led to approval for this indication in many European countries. The greatest pitfalls are created by the lack of established protocols for the walking child with spastic cerebral palsy.

Doses of 2-6 U/kg body weight per muscle with a maximum total dose of 29 U/kg have been reported. Although there are no standard guidelines on doses in children, a review of previous publications indicates that the dose that has been used for children with cerebral palsy has increased over time. The current practice is to inject several muscles at each injection session, with smaller patients receiving higher doses than reported in the past.

The dosage of BTX-A must be individualized for each patient. A recent consensus statement recommended a dose of 12 units of Botox (Allergan, USA) per kg bodyweight, with a maximum of 300 units Botox per child on one occasion, and 50 units per injection site. The recommended treatment interval is 6-12 months, with a minimum of 3 months. It has become apparent that larger doses used less frequently are safe, more effective, better tolerated by children, more acceptable to their families, and the functional outcomes are longer lasting. Dosages of 16-24 units per kg bodyweight are now in use in experienced hands for double or triple level lower limb sites. However, the dose-efficacy relationship might only be linear for a distinct range. Clinicians should be aware of unwanted effects during the application of the drug.

The mean dose of BTX-A used varies between treatment centres. This cannot be entirely explained on the basis of different patient populations. The use of a small dose of BTX-A may reflect a cautious attitude of the practitioner or an attempt to reduce the cost of treatment. An alternative explanation is that BTX-A is used in conjunction with other antispasticity treatments. Similarly, the variability between examiners in the evaluation of the clinical effectiveness of BTX-A may be a factor.

The clinician may inject a large dose of BTX-A to achieve a desired reduction in muscle tone in a given patient but may use a smaller dose in another subject, although both patients have the same degree of muscle spasticity. This need to titrate the dose of BTX-A is necessary because muscle spasticity affects different patients in different ways and the treatment decision in each case is usually based upon therapeutic goals. The choice of therapeutic interventions is also influenced by locally available expertise and resources.

Three randomised, double-blind dose-ranging studies have examined the effects of various doses of BTX-A on gait measures in patients with cerebral palsy. In these studies, the higher doses of the preparations were more effective in improving gait measures than the lower doses.

What is the dose of BTX-A in cerebral palsy? There is some debate concerning optimum treatment of BTX-A in the non-ambulatory child. The greatest pitfalls are created by a lack of established protocols for the walking child with spastic cerebral palsy. There is currently no consensus among clinicians about how an optimal dose of BTX-A should be determined, and there are no standard guidelines on doses of BTX-A in children.

There is little doubt that BTX-A has been an outstanding advance in recent years, creating important opportunities to enable our therapy and treatments to be significantly more effective in the management of cerebral palsy. More
controlled trials are now needed to confirm the anti-
spasticity effects of BTX-A with larger patients groups for
more accurate results.

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