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Improvements in Lower Limb Spasticity-Related Pain in Children/ Adolescents With Cerebral Palsy After IncobotulinumtoxinA Injections

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But its efficacy is lacking and even symptomatic intake to cure comorbidities or autistic symptoms itself doesn't bring long-term benefit. There should be emphasis on psycho-correctional and physical interventions including chiropractic care. Using high-velocity low-amplitude chiropractic intervention may be more beneficial in comparison to using solely psycho-correctional techniques. Although translating insights from research data into daily clinical practice is not an easy task. It takes a huge effort in terms of medical staff's education as well as clarifying the benefits of chiropractic care to patients and healthcare professionals.

Treatment: Muscle and movement

P-247 | Absence of neutralizing antibody formation during incobotulinumtoxinA treatment of spasticity in botulinum toxin-naïve children with cerebral palsy: Pooled analysis of three phase 3 studies

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Introduction: The development of neutralizing antibodies (NABs) has been linked to secondary non-response to botulinum neurotoxin type A (BoNT-A) injections. This may be of special concern when treating conditions like pediatric spasticity. We investigated NAB formation in three large Phase 3 studies with incobotulinumtoxinA, a BoNT-A lacking complexing proteins, in children/adolescents with cerebral palsy (CP).

Patients and methods: Patients with lower-limb (LL), upper-limb (UL), or combined LL/UL spasticity (2–17 years; uni- or bilateral CP; Ashworth Scale score ≥ 2 in clinical patterns for treatment) received total body weight incobotulinumtoxinA ≤ 16 –20 U/kg (max. 400–500 U) depending on the study (TIM: NCT01893411; TIMO: NCT01905683; XARA: NCT02002884) and Gross Motor Function Classification System level I–V, for up to six injection cycles (ICs). Occurrence of NABs against BoNT-A was investigated in those ≥ 21 kg at screening and end of study. Blood samples were analyzed using a fluorescence immunoassay (FIA) for antibodies; positive samples were tested for NABs using a hemidiaphragm assay.

Results: 907 patients received treatment. 386/403 (95.8%) and 318/422 (75.4%) with bodyweight ≥ 21 kg were tested

using FIA at screening and end of study, respectively. 150/403 (37.2%) and 167/422 (39.6%) were toxin-naïve. Eleven patients tested positive for NABs at screening and/or end of study, all of whom had previously been treated with other BoNT-As (onabotulinumtoxinA/abobotulinumtoxinA). No patient developed a secondary non-response to incobotulinumtoxinA. No toxin-naïve patients developed NABs after incobotulinumtoxinA treatment.

Conclusions: NAB formation was not observed in toxin-naïve children/adolescents with CP treated with up to six ICs of incobotulinumtoxinA.

P-248 | Improvements in lower limb spasticity-related pain in children/adolescents with cerebral palsy after incobotulinumtoxinA injections

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Introduction: The effects of incobotulinumtoxinA on lower limb (LL) spasticity-related pain (SRP) over multiple treatment cycles (ICs) in children/adolescents (C/As) with cerebral palsy (CP) were analyzed using pooled data from three prospective phase 3 trials.

Patients and methods: C/As aged 2–17 years with CP-associated LL spasticity received incobotulinumtoxinA for 4 ICs. SRP was assessed with the Questionnaire on Pain caused by Spasticity (QPS) using C/A- (direct or via interviewer) and parent/caregiver (P/C)-completed LL modules. The pain population included all C/As with a key QPS item score >0 at baseline; post-baseline scores of 0 indicated complete pain relief.

Results: Data from 331 C/As and 841 P/Cs with data for at least one key QPS item were included. LL general SRP was reported by 178 C/As at baseline; 35.3%/49.4% of patients treated with incobotulinumtoxinA were pain-free by week 4 of IC1/IC4 ($p < 0.001$ vs baseline for all ICs), at which times C/A-reported mean LL QPS general item intensity scores had improved by 2.1/2.8 points ($p < 0.001$ vs baseline for all ICs). P/Cs observed LL general SRP in 568 C/As at baseline; 25.2%/34.1% of patients treated with incobotulinumtoxinA were pain-free by week 4 of IC1/IC4 ($p < 0.001$ vs baseline for all ICs). C/A-reported and P/C-observed improvements were generally greater with demanding tasks than at rest and more pronounced with increasing incobotulinumtoxinA ICs.

Conclusion: In addition to muscle tone regulation, incobotulinumtoxinA provides sustained pain relief across multiple ICs for children with CP and LL SRP, even when they were engaged in demanding tasks.

P-249 | Improvements in upper limb spasticity-related pain in children/adolescents with cerebral palsy after incobotulinumtoxinA injections

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Introduction: The effects of incobotulinumtoxinA on upper limb (UL) spasticity-related pain (SRP) over multiple treatment cycles (ICs) in children/adolescents (C/As) with cerebral palsy (CP) were analyzed using pooled data from two prospective phase 3 trials.

Patients and methods: C/As aged 2–17 years with CP-associated UL spasticity received incobotulinumtoxinA for 4 ICs. SRP was assessed with the Questionnaire on Pain caused by Spasticity (QPS) using C/A- (direct or via interviewer) and parent/caregiver (P/C)-completed UL modules. The pain population included all C/As with a key QPS item score >0 at baseline; post-baseline scores of 0 indicated complete pain relief.

Results: Data from 155 C/As and 444 P/Cs with data for at least one key QPS item were included. UL general SRP was reported by 69 C/As at baseline; 39.7%/41.8% of patients treated with incobotulinumtoxinA were pain-free by week 4 of IC1/IC4 ($p < 0.001$ vs baseline for all ICs), at which times C/A-reported mean UL QPS general item intensity scores had improved by 1.7/2.2 points ($p < 0.001$ vs baseline for all ICs). P/Cs observed UL general SRP in 294 C/As at baseline; 28.3%/38.2% of patients treated with incobotulinumtoxinA were pain-free by week 4 of IC1/IC4 ($p < 0.001$ vs baseline for all ICs). C/A-reported and P/C-observed improvements were generally greater with demanding tasks than at rest and more pronounced with increasing incobotulinumtoxinA ICs.

Conclusion: In addition to muscle tone regulation, incobotulinumtoxinA provided sustained pain relief across multiple ICs for children with CP and UL SRP, even when they were engaged in demanding tasks.

P-250 | Muscle stem cell characteristics across contracted muscles and functional levels in children with cerebral palsy

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Introduction: Satellite cells (SC) are muscle-specific stem cells that facilitate postnatal development and repair. Children with CP have impaired muscle growth and develop contractures. We investigated if SC characteristics differs between contracted and non-contracted muscles and across GMFCS functional levels in children with CP.

Patients and methods: Thirty-six children participated in this study (CP: mean age 11.32 ± 0.57 , sex: 22 male, 12 female, GMFCS: I-V, TD: mean age 13.50 ± 0.76 , sex: 3 male, 3 female). Muscle biopsies ($n = 40$) were obtained from vastus lateralis (TD and CP), adductors and gastrocnemius (CP) during surgery for contractures (CP) or for ACL repair (TD). Muscle cross-sections were immunohistochemically labeled for Pax7, Dystrophin, MHC-1, DAPI, Ki67, MyoD in 2–3 serial sections.

Results: Non-contracted muscles (VL) in children with CP have overall lower SC abundance and a higher percentage of activated SCs compared to TD children (10.1 ± 2.0 vs. 19.7 ± 8.5 SC/100 fibers, $p < 0.05$) (3.9 ± 2.5 vs. $1.3 \pm 2.01\%$ activated SCs, $p < 0.05$). Contractured muscles exhibited higher SC abundance with lower percentage of activated SCs compared to non-contracted muscles (16 ± 4.4 vs. 10.1 ± 2 SC/100 fibers, $p < 0.05$) (0.6 ± 0.8 vs. $3.9 \pm 2.5\%$ activated SCs $p < 0.05$). SC abundance was similar across GMFCS levels.

Conclusion: SCs of CP contracted muscle appear to be abundant but have lower myogenic potential compared to non-contracted muscles. These SCs may contain an exasperated intrinsic cell deficiency disallowing them from progressing into proliferative/differentiative stages in attempts to facilitate growth.

P-251 | Prevalence of spasticity-related pain in children/adolescents with cerebral palsy

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