

Evaluation of prepubertal patients with suspected neurosecretory dysfunction for growth hormone secretion (NSD): diagnostic steps and treatment response

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Conclusion: According to our results, analysing overnight GH-secretion remains to only method to identify children with neurosecretory dysfunction for GH-secretion (NSD). Age, auxologic data, delay of bone age and IGF-I/IGFBP3-levels are not different between patients and controls. Yet, as response to GH-treatment is comparable to results in idiopathic GH-deficiency (GHD), it is worth to consider this diagnosis.

Background / Aims: Existence and diagnostic procedures of neurosecretory dysfunction (NSD) are still a matter of debate. The aim of our study was to analyse prediagnostic data of short-statured children with pathologic and normal spontaneous GH-secretion and to evaluate the effect of GH-therapy in NSD-patients.

Methods: In 90 children aged 3 to 16 years 12-hour night profiles for GH-secretion (samples every 30 minutes) were performed (unicentric). Children fulfilling 2 of the 3 following criteria were diagnosed having NSD: number of peaks \leq 3, maximal GH-secretion < 8,0 ng/ml, mean secretion < 3,2 ng/ml. By this, 49 children were classified having NSD and treatment with recombinant GH was started. Their auxologic data, IGF-I-/IGFBP3-levels, GH-stimulation tests as well as spontaneous overnight GH-secretion were analysed and compared to the data of the 41 children with normal spontaneous GH-secretion. Additionally, follow-up auxological data of the NSD-patients during GH-treatment were collected.

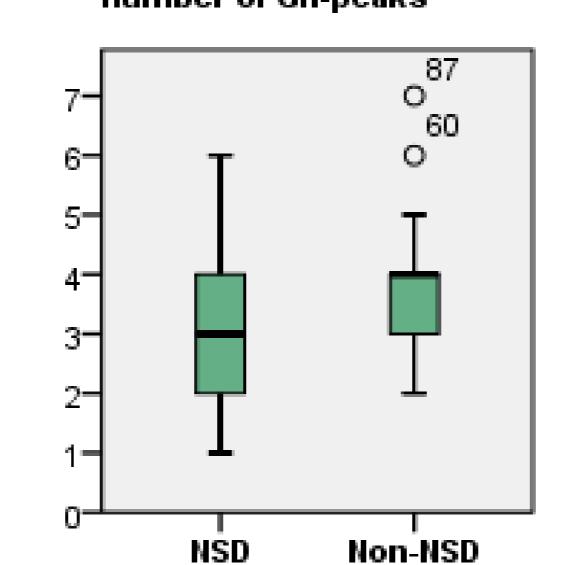
Results:

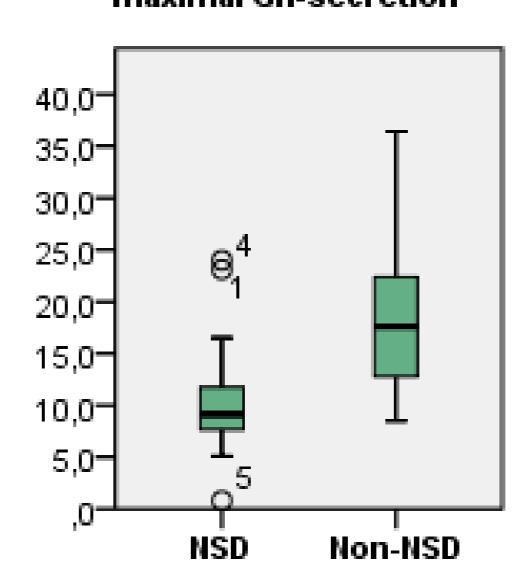
Age, prediagnostic auxologic data (height, weight, HSDS, HV, HV-SDS), delay of bone age and IGF-I/IGFBP3-levels did not differ between the two groups.

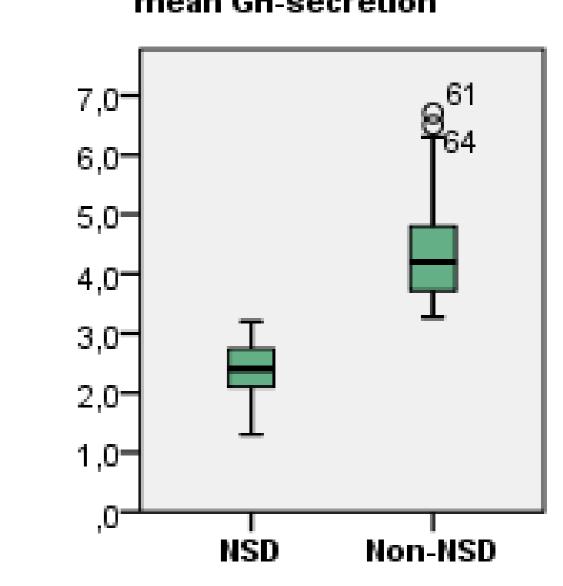
| | | age (years) | weight (kg) | height (cm) | H-SDS | HV (cm/year) | HV-SDS | dBA (years) | IGF- SDS | IGFBP3- SDS |
|---------------------------|---------|----------------|----------------|----------------|-------|-----------------|--------|----------------|-------------|----------------|
| NSD- | n | 49 | 47 | 49 | 49 | 49 | 48 | 49 | 47 | 45 |
| group | mean | 9,73 | 22,94 | 120,85 | -2,82 | 4,32 | -1,03 | 1,6 | -1,81 | -0,62 |
| | median | 10,33 | 22 | 124,4 | -2,75 | 4,4 | -1,36 | 1,67 | -2 | -0,71 |
| | SDS | 2,97 | 8,43 | 15,83 | 0,64 | 1,69 | 2,04 | 1,21 | 0,67 | 0,9 |
| Non-NSD | n | 41 | 40 | 41 | 41 | 41 | 41 | 40 | 41 | 39 |
| group | mean | 10,42 | 24,39 | 125,18 | -2,91 | 4,76 | -0,06 | 1,82 | -1,79 | -0,73 |
| | median | 10,33 | 21,9 | 122 | -2,92 | 4,6 | -0,87 | 2,21 | -1,78 | -0,7 |
| | SDS | 3 | 8,47 | 16,33 | 0,98 | 2,13 | 2,76 | 1,68 | 0,85 | 0,76 |
| statistical difference | p-value | 0,275 | 0,429 | 0,206 | 0,612 | 0,281 | 0,06 | 0,468 | 0,894 | 0,552 |

Instead, for all 3 criteria used for evaluation of the 12-hour night profiles (number of peaks, maximal and mean GH-secretion) a significant difference between NSD- and Non-NSD-children was found (p-value < 0,005 for all parameters; maximal and mean GH-secretion shown in ng/ml).</p>
maximal and mean GH-secretion shown in ng/ml).
maximal GH-secretion

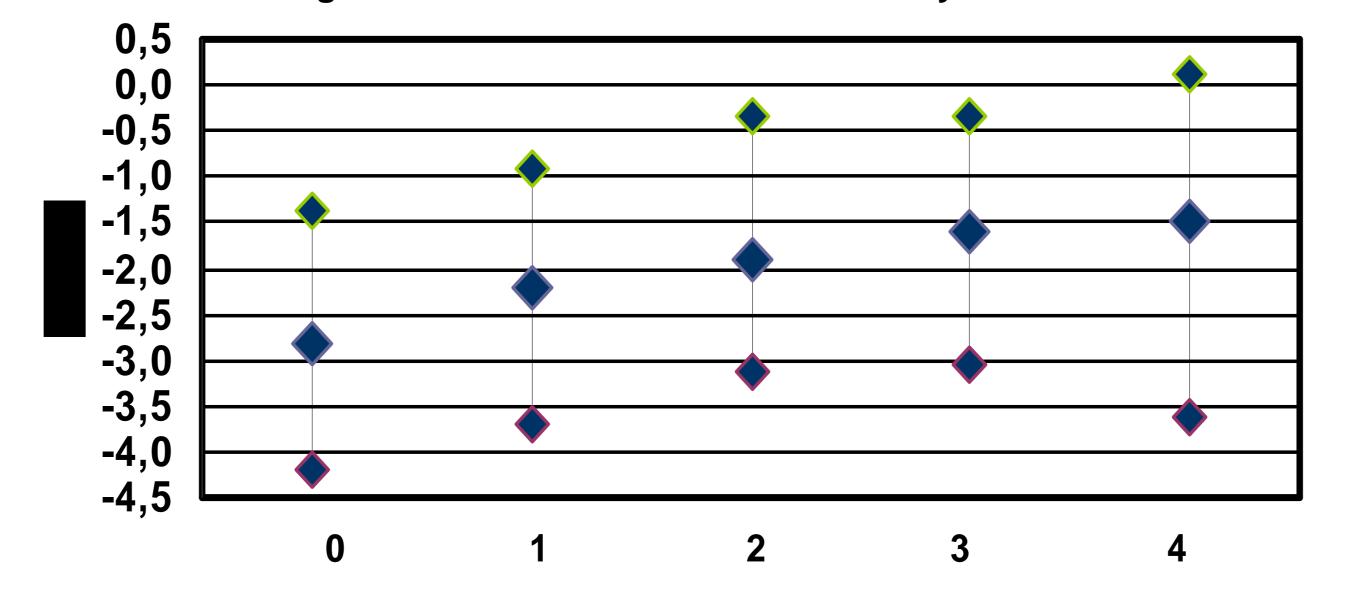
| 12-hour night profiles | | number of peaks | maximal secretion | mean secretion |
|---------------------------|--------|--------------------|-------------------|-------------------|
| cut-off | | 3,0 | 8 | 3,20 |
| NSD-group | mean | 2,9 | 10,2 | 2,40 |
| | median | 3 | 9,2 | 2,40 |
| | SDS | 1,1 | 4,1 | 0,50 |
| Non-NSD- | mean | 3,9 | 18,3 | 4,40 |
| group | median | 4 | 17,6 | 4,20 |
| | SDS | 1 | 6,6 | 0,90 |







Children with NSD showed a good response to GH-treatment after 1 year (DHSDS: +0,77 +/- 0,48, DHV-SDS: 4,4 +/- 3,51 cm/year) as well as until after 4 years (DHSDS: +1,51 +/- 0,75, DHV-SDS: +0,77 +/- 1,92 cm/year). These results are similar to those of children with idiopathic GHD.



Change in H-SDS from baseline until 4 years of treatment

This work was supported with an unrestricted research grant from Pfizer.

