

Wasniewska Małgorzata¹, Morabito Letteria Anna¹, Baronio Federico², Einaudi Silvia³, Salerno Maria Carolina⁴, Bizzarri Carla⁵, Russo Gianni⁶, Chiarito Mariangela⁷, Grandone Anna⁸, Guazzarotti Laura⁹, Spinuzza Antonietta¹⁰, Di Carlo Silvia¹, Ortolano Rita², Balsamo Antonio², Abrigo Enrica³, Baldini-Ferrolì Barbara⁵, Alibrandi Angela¹⁰, Capalbo Donatella⁴, Faienza Maria Felicia⁷

- ¹Department of Human Pathology of Adulthood and Childhood 'Gaetano Barresi', University of Messina, Messina, Italy
²Department of Women, Children and Urological Diseases, S. Orsola Malpighi University Hospital, Bologna, Italy.
³Department of Pediatric Endocrinology and Diabetology, Regina Margherita Children Hospital, University of Turin, Turin, Italy.
⁴Department of Pediatrics, University "Federico II", Naples, Italy
⁵Unit of Endocrinology and Diabetes, 'Bambino Gesù Children's Hospital, Rome, Italy
⁶Department of Pediatrics, Endocrine Unit, Scientific Institute San Raffaele, Milan, Italy
⁷Department of Biomedical Sciences and Human Oncology, Pediatric Section, University of Bari 'A. Moro', Bari, Italy
⁸Department of Woman, Child and General and Specialized Surgery, Univeristà degli Studi della Campania "Luigi Vanvitelli", Naples, Italy.
⁹Department of Pediatrics, University of Padua, Padua, Italy
¹⁰Department of Economics, Unit of Statistical and Mathematical Sciences, University of Messina, Messina, Italy.

Background

Subjects with non classical congenital adrenal hyperplasia (NCCAH) often present an increased growth velocity secondary to elevation of adrenal androgens that promote early bone maturation and compromise final height (FH). The aim of the study was to analyze prognostic factors affecting growth trajectory and FH in children with NCCAH.

DESIGN:

retrospective, multicentric study

Study population:

192 (140 females) NCCAH children with confirmed molecular diagnosis followed from diagnosis up to FH.

Methods:

clinical records were collected and analyzed. The study population was divided for **gender**, with or without **hydrocortisone treatment** (171 treated with hydrocortisone) and type of the **mutation of CYP21A2 gene** (V281L homozygosis in 55, compound heterozygosis with V281L in 85 and other mutations in 48 cases).

FH (SDS), pubertal growth (**PG**) (cm), growth trajectory (**GT**) since diagnosis to FH (SDS) and FH adjusted to target (TH) (**FH-TH**)(SDS) were evaluated as outcomes using stepwise linear regression models.

Results:

FH SDS and **FH-TH** were not significantly different in both gender (**-0.34 vs -0.36, p=0.98** and **-0.05 vs 0.05, p=0.65**, respectively).

At stepwise linear regression analysis, **FH** and **FH-TH** resulted significantly related to **chronological age (CA)** ($p=0.008$ and 0.016), **bone age (BA) / CA ratio** ($p=0.004$ and 0.001), **height (H)** ($p=0.000$ for both parameters) at NCCAH diagnosis and TH ($p=0.013$ and 0.000).

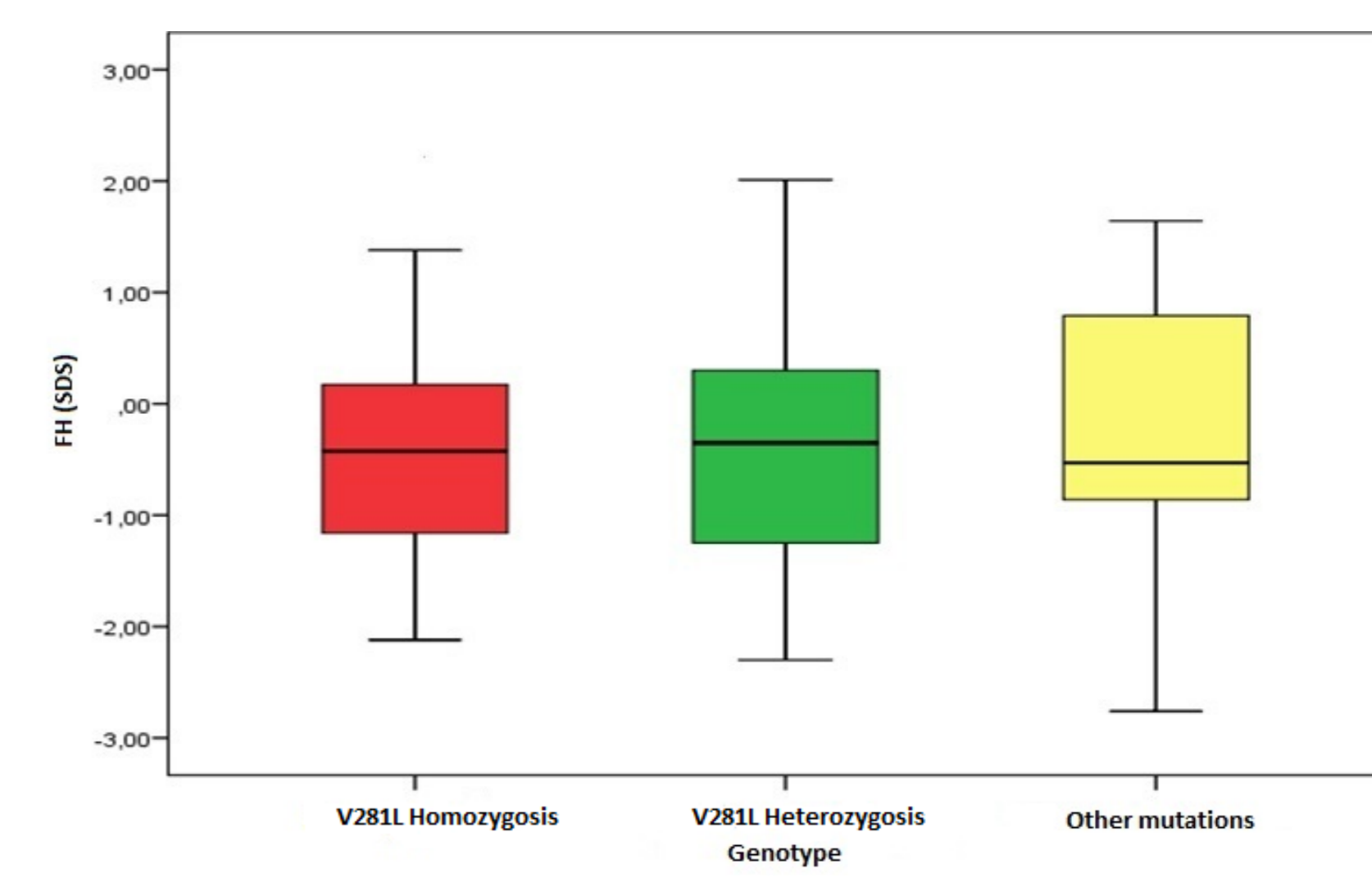
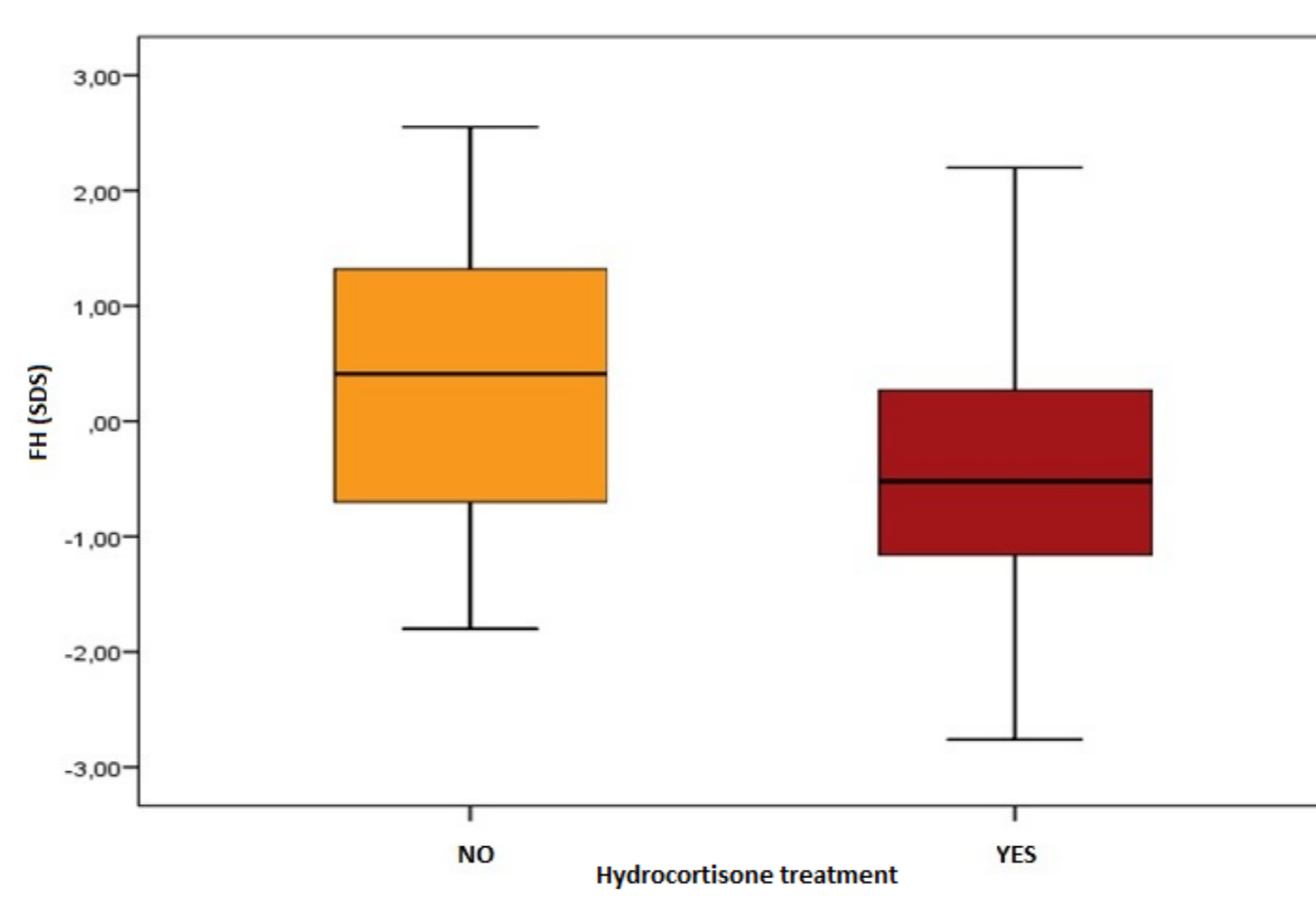
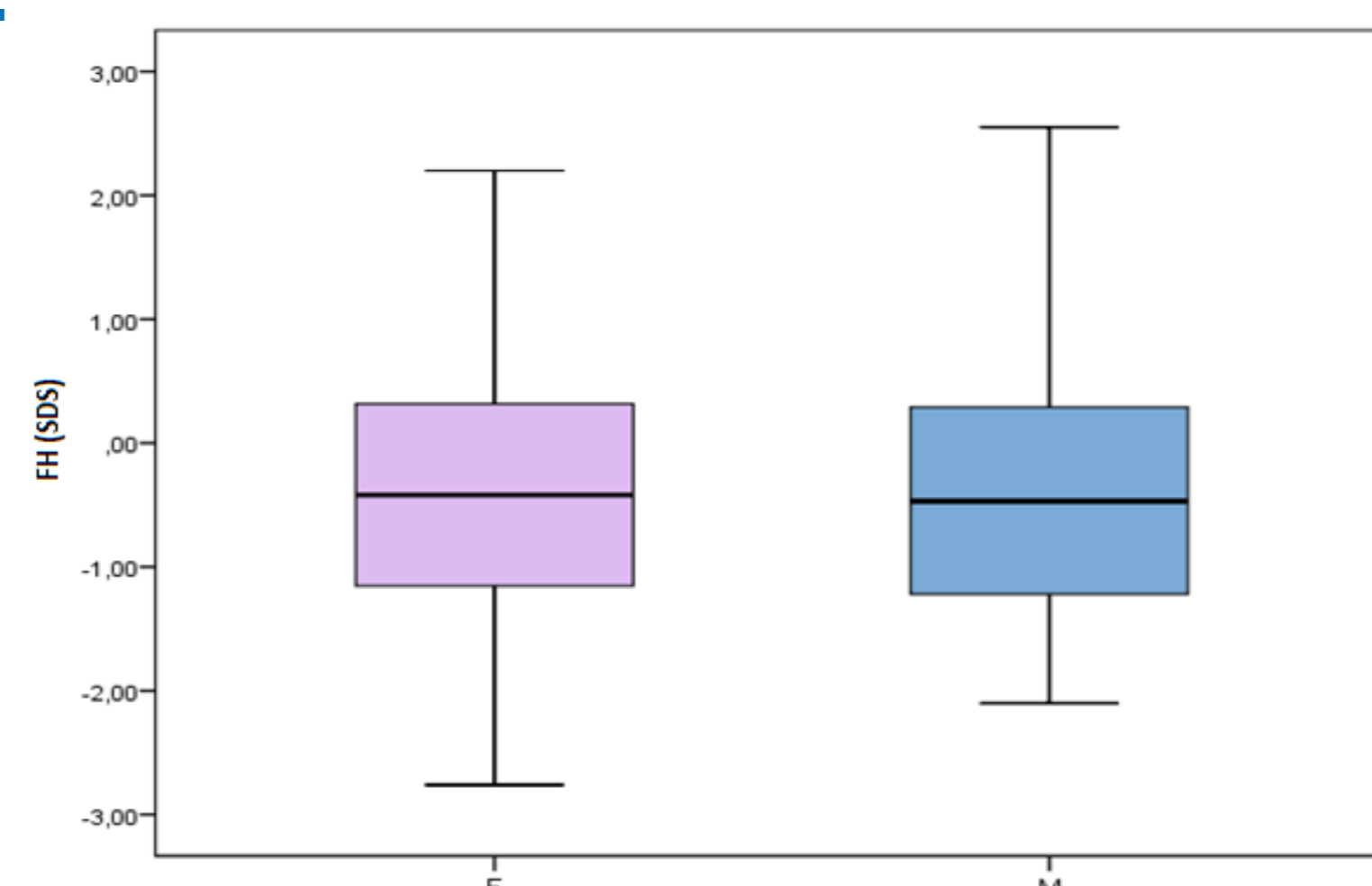
PG was **higher in males** (22.59 ± 5.74 vs 20.72 ± 17.4 cm in females) ($p=0.002$), as physiologically observed, and was **positively related to H** ($p=0.027$), **negatively to BMI** ($p=0.001$) and **BA/CA ratio** ($p=0.001$) at NCCAH diagnosis.

The **type of the mutation** of CYP21A gene and **hydrocortisone doses** did not influence significantly the parameters of growth of our NCCAH patients.

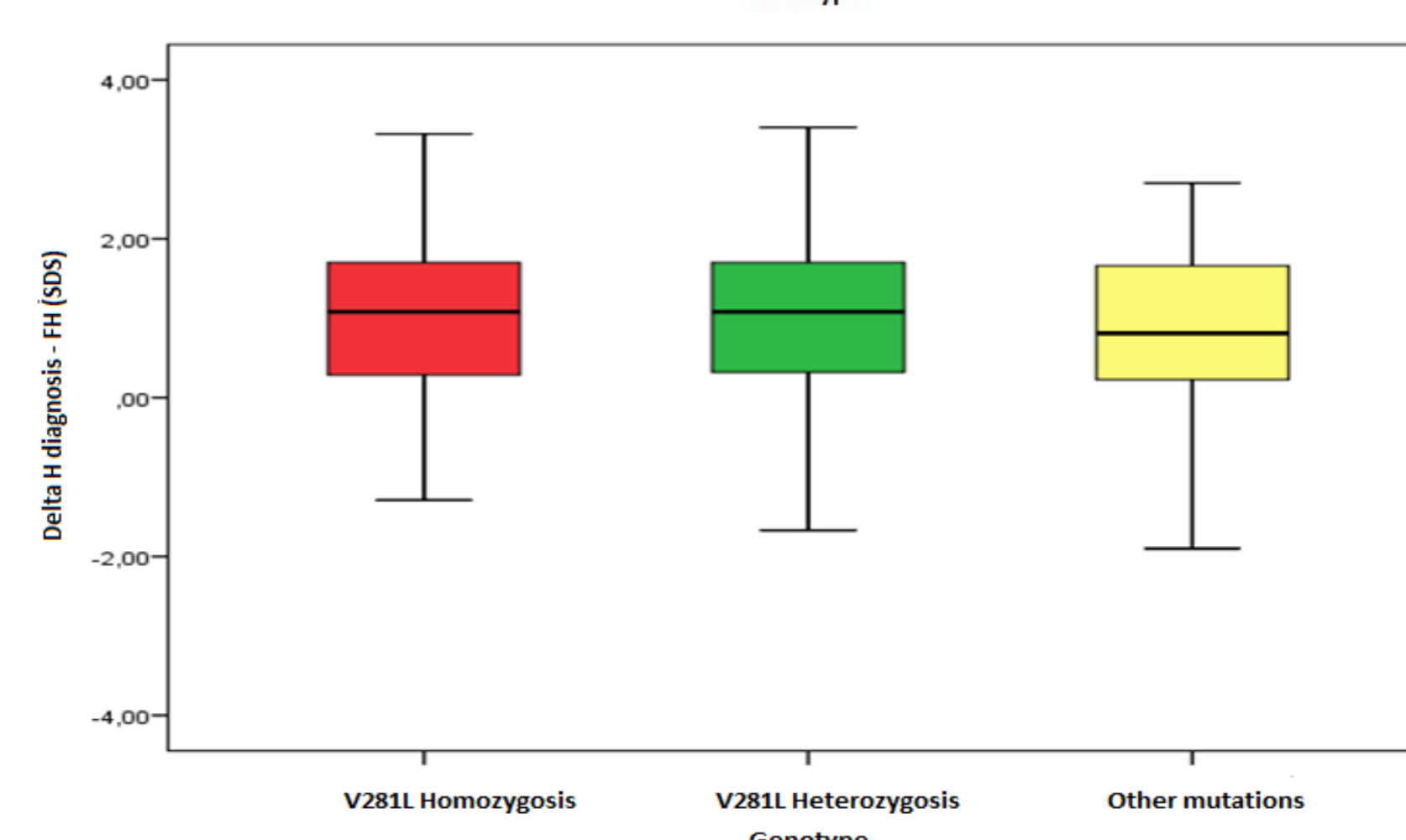
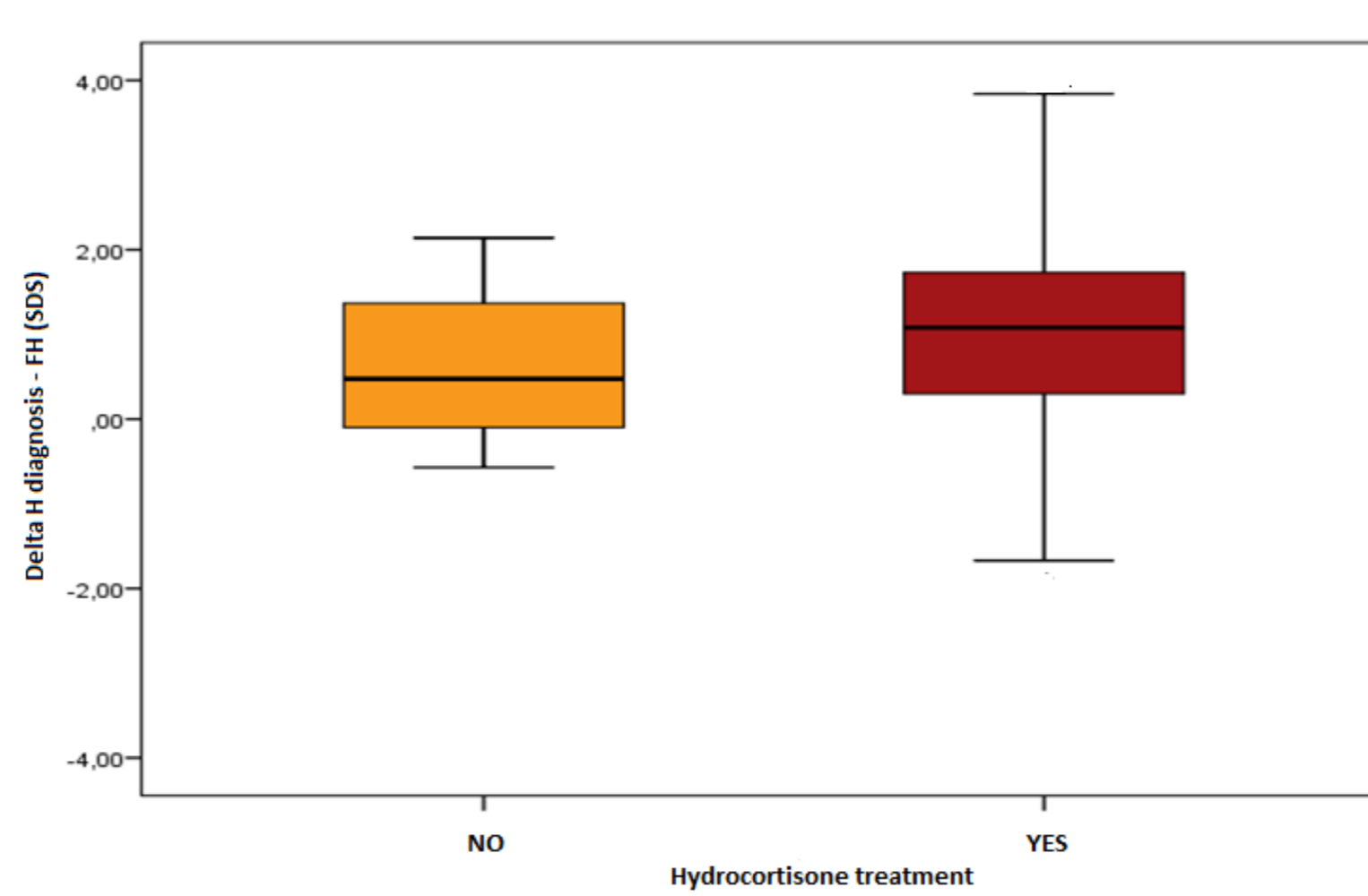
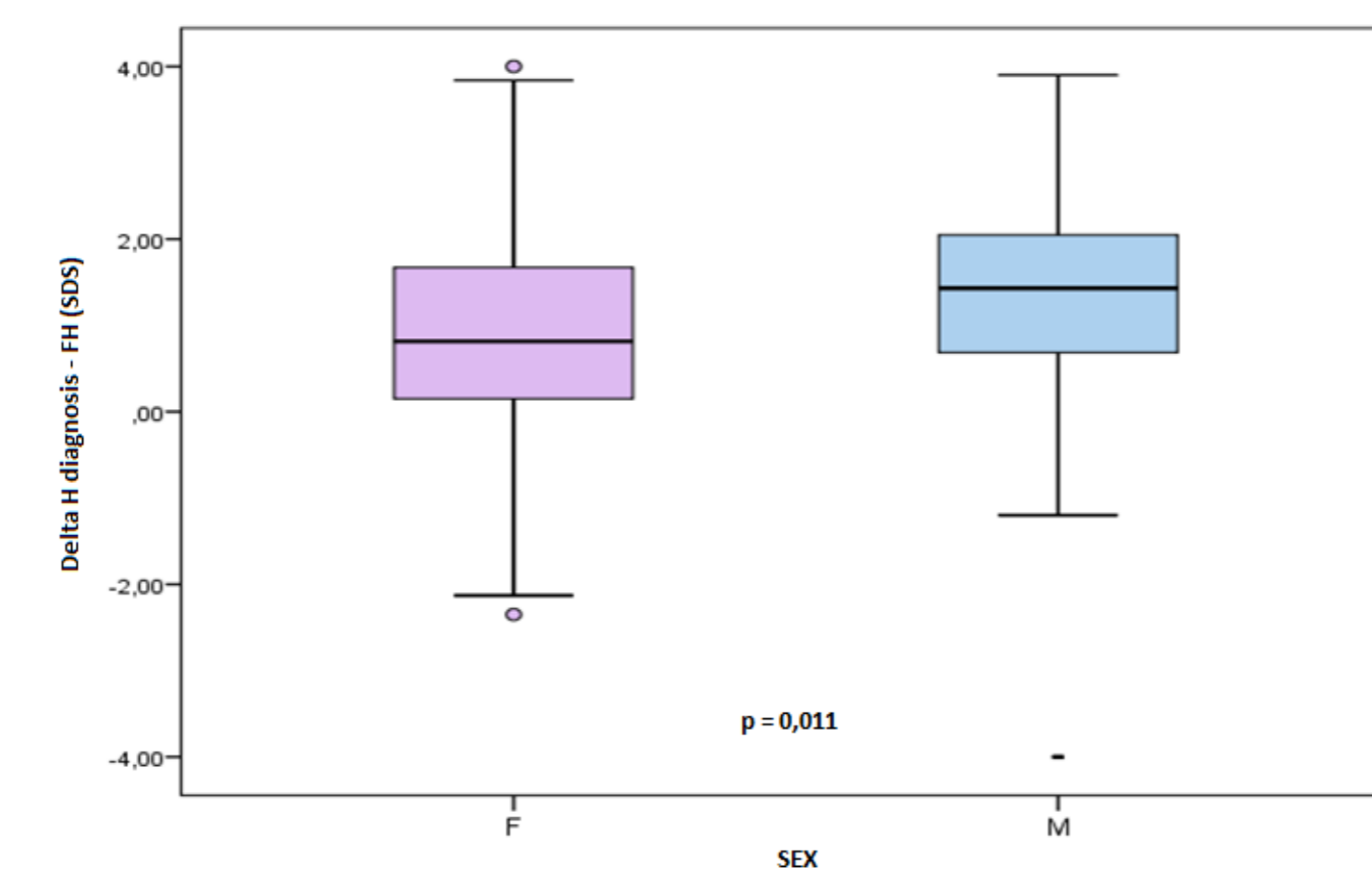
The comparison between treated with hydrocortisone and untreated patients did not evidence significant differences on GT, but the statistic value of these results is limited by the small number of untreated group

OUTCOMES

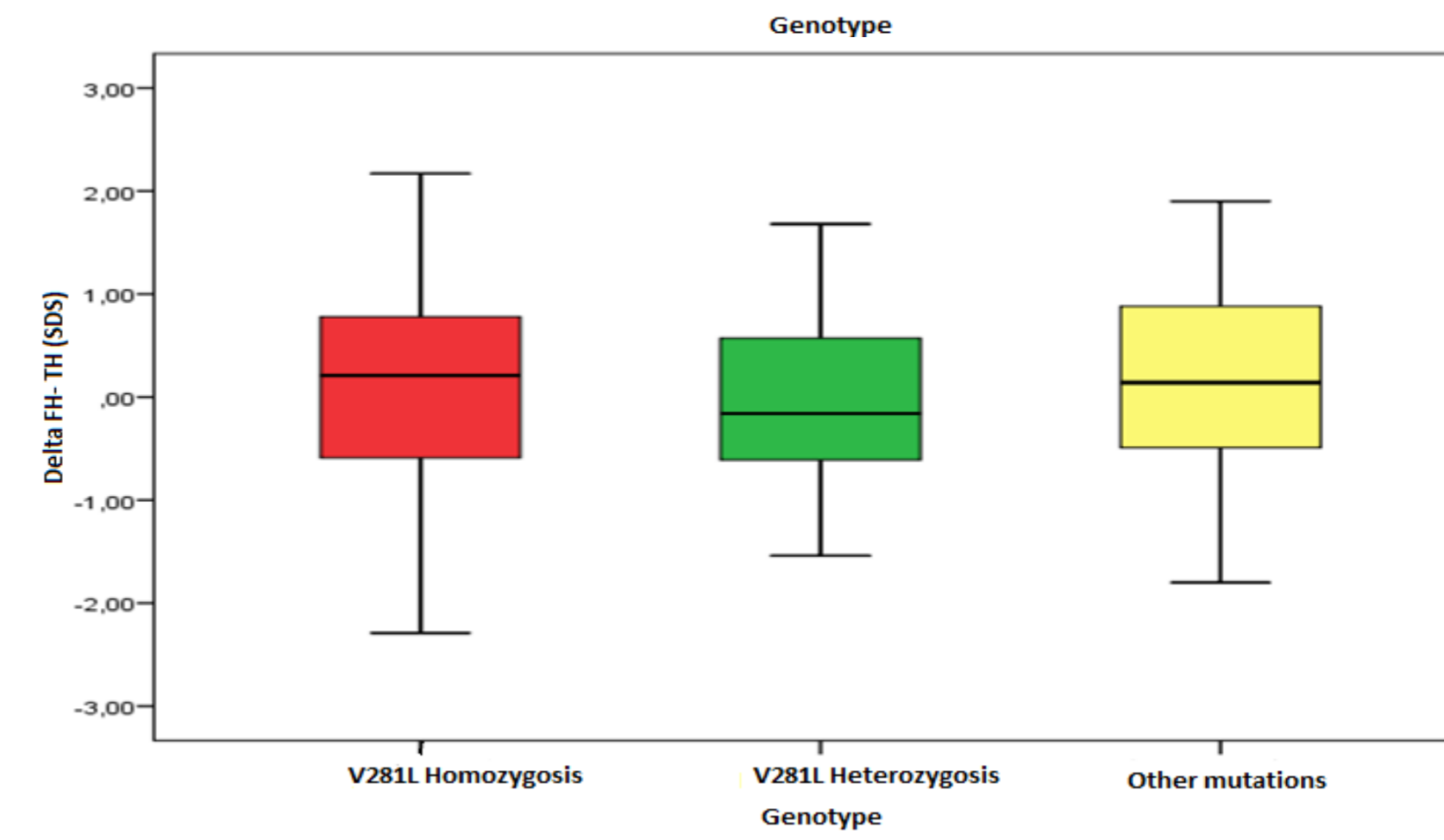
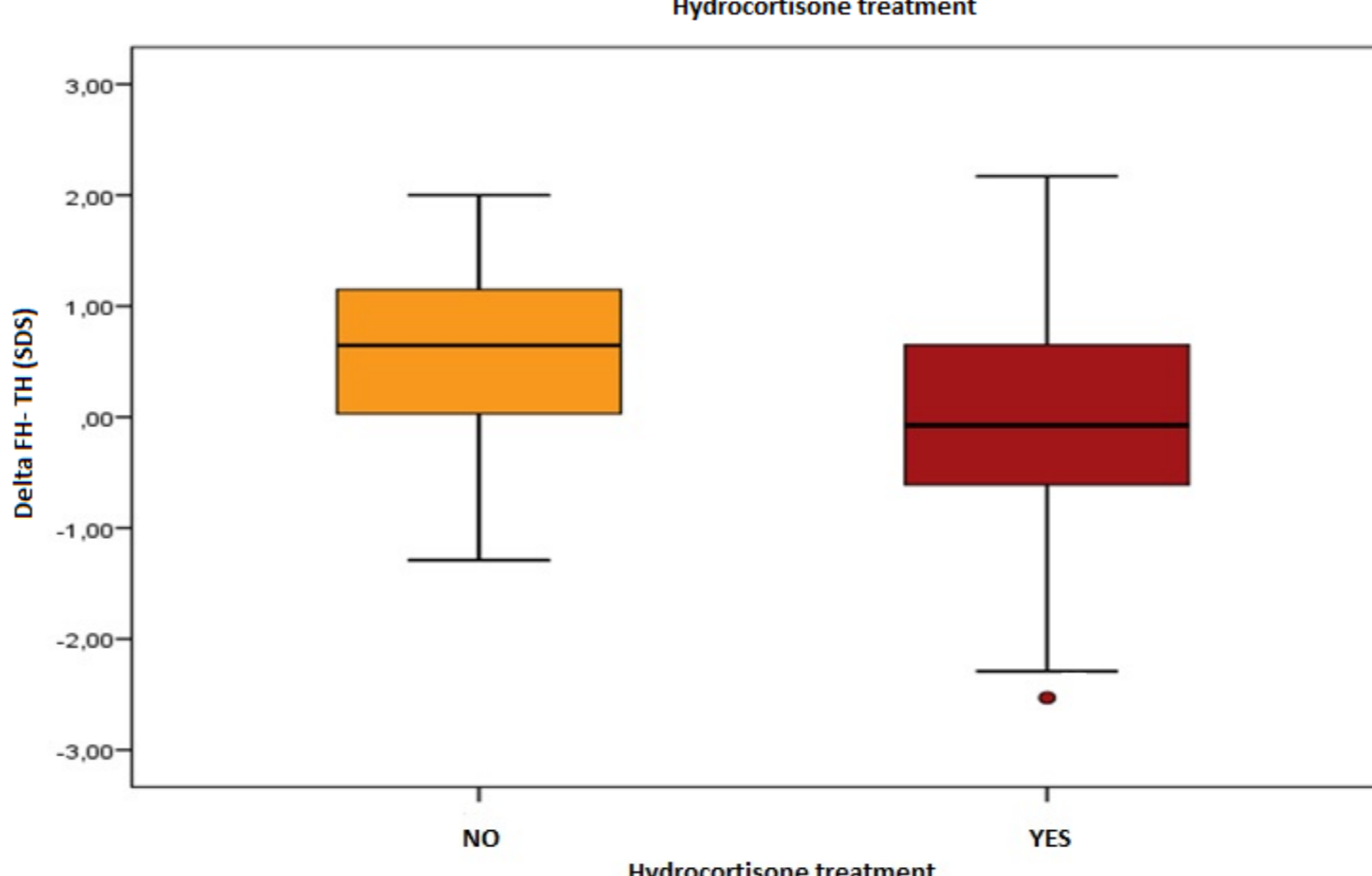
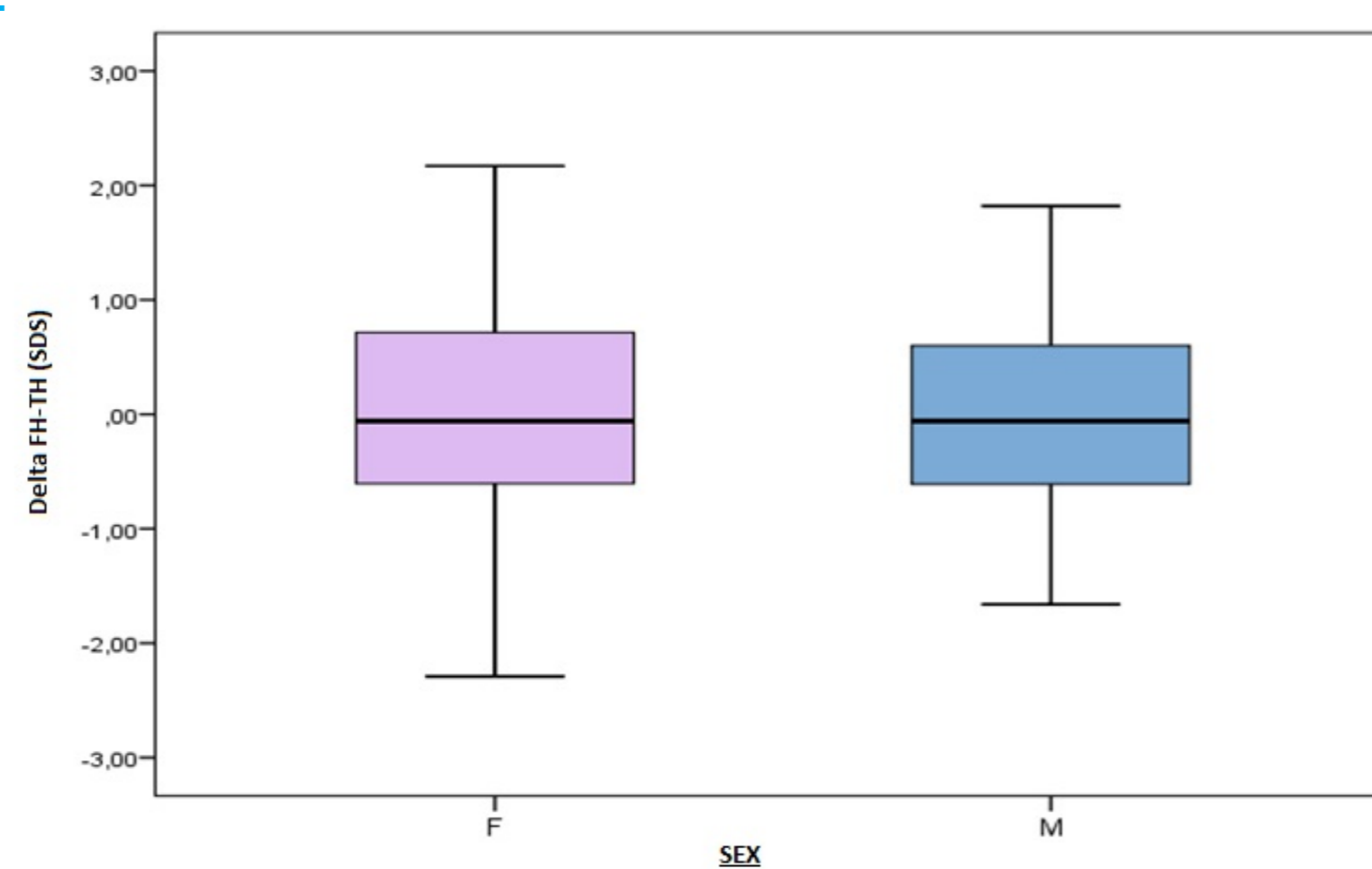
FH (SDS)



Delta H diagnosis - FH (SDS) = GT



Delta FH- TH (SDS)



CONCLUSION

FH and GT of NCCAH patients are significantly influenced by auxological parameters at diagnosis (CA, BA/CA ratio, H).

Gender, molecular alteration, biochemical picture and hydrocortisone doses seem to have no important influence on height outcome of these NCCAH children.