

Obesity and cardio-metabolic risk factors among children and adolescents with non classical 21-Hydroxylase deficiency



מרכז שניידר לרפואת ילדים בישראל אקאל سוויבר לאור ועקלפון פא ושעווייל Schneider Children's Medical Center of Israel

Member of Clalit Health Services

Liat de Vries, Yael Lebenthal, Ariel Tenenbaum, Moshe Phillip, Rachel Bello

The Jesse Z and Sara Lea Shafer Institute for Endocrinology and Diabetes

Schneider Children's Medical Center of Israel, Petah Tikva, and Sackler Faculty of Medicine, Tel Aviv University, Tel Aviv, Israel

## Introduction

Increased risk of obesity and cardio-metabolic risk factors were reported in patients with classical

# Clinical characteristics of patients with NCCAH at diagnosis

F:M	91:23
Age at diagnosis ( yrs)	7.9±4.2
Age at treatment initiation (yrs)	8.6±3.4
1 <sup>st</sup> year Hydrocortisone dose (mg/m <sup>2</sup> )	10.6±4.9
Height-SDS	0.10±1.22
Weight-SDS	0.22±1.23
BMI-SDS	0.35±1.13
Bone age – chronological age (yrs)	1.06±1.26
Basal 170HP (nmol/l)	28.7±27.6
	(2.4-174)
Peak 17OHP (nmol/l)	129.5±96.8
Stimulated cortisol level (nmol/l)	567±165
Paternal BMI	26.7±3.9
Maternal BMI	26.0±5.1

### Rates of overweight and obesity

	n	<b>Overweight (%)</b>	Obesity (%)
All patients	114	21.9	11.4
National health survey	3,443	26.5	15.1

congenital adrenal hyperplasia but little is known

about adiposity among patients with non-classical congenital adrenal hyperplasia (NCCAH).

## Objectives

To assess the prevalence of overweight, obesity and cardio-metabolic risk factors among NCCAH patients

diagnosed in childhood.

# Study design and methods

#### Design

• A cross-sectional retrospective study conducted in a tertiary pediatric endocrinology unit.

#### Study cohort

# Clinical characteristics by current glucocorticoid treatment

	Treated (n=76)	Untreated (n=38)	<i>P</i> -value
Current age (yrs)	$17.4 \pm 7.1$	$16.6 \pm 6.5$	0.5
F:M	63:13	28:10	0.2
Tanner (1, 2-4, 5)	8, 14, 54	6, 6, 26	0.3
Treatment duration (yrs)	$7.3 \pm 6.3$	6.5 ± 5.2	0.5
Current Weight-SDS	$0.39 \pm 0.99$	0.06 ± 1.29	0.1
Current Height-SDS	$-0.44 \pm 0.99$	$-0.42 \pm 1.16$	0.9
Current BMI-SDS	$0.65 \pm 0.89$	$0.34 \pm 1.00$	0.09
Current daily HC dose (mg/m <sup>2</sup> )	9.2 ± 4.5		
Systolic blood pressure (mm Hg)	$113 \pm 11$	$116 \pm 11$	0.3
Systolic BP percentile	$61.2 \pm 27.1$	68.9 ± 24.4	0.2
Diastolic blood pressure (mm Hg)	66.7 ± 9.5	67.9 ± 7.8	0.6
Diastolic BP percentile	$56.1 \pm 26$	$60.4 \pm 22.5$	0.5
17OHP (nmol/l)	$22.0 \pm 29.5$	$19.5 \pm 17.6$	0.7
Androstenedione (nmol/l)	$6.4 \pm 4.1$	9.2 ± 8.3	0.4
Testosterone (nmol/l)	2.5 ± 4.8	$1.9 \pm 3.1$	0.8

Study cohort (7-12th grade)	76	23.5	10.5
<i>P</i> -value		0.24	0.18
Mothers	100	36.8	13.6
<i>P</i> -value		0.01	0.4
Fathers	86	52.5	14.5
<i>P</i> -value		<0.001	0.3

# Cardio-metabolic risk factors by current treatment

	Treated (n=22)	Untreated (n=16)	<i>P</i> -value
Age (years)	$15.7\pm5.4$	$17.1\pm5.7$	0.5
%fat by Bioelectrical impedance	$\textbf{27.8} \pm \textbf{6.8}$	$\textbf{21.4} \pm \textbf{8.3}$	0.02
lliac skin fold (mm)	$16.1\pm8.9$	$14.2\pm7.0$	0.5
Triceps skin fold (mm)	$18.4\pm8.7$	$14.1\pm6.6$	0.1
Scapular skin fold (mm)	$13.4\pm9.6$	9.3 ± 6.0	0.2
Waist circumference (cm)	74.1 ± 13.2	67 ± 11.9	0.1
Patients with waist circumference percentile >50 (%)	55.5	33	0.2
Hip circumference (cm)	86 ± 16	79 ± 18	0.28
Waist/Hip ratio	$\textbf{0.88} \pm \textbf{0.13}$	$0.87\pm0.09$	0.8
HDL (mg/dl)	$59\pm19$	$57\pm16$	0.6
LDL (mg/dl)	$107\pm33$	$106\pm28$	0.9
TG (mg/dl)	89 ± 32	$118\pm 66$	0.07
Cholesterol (mg/dl)	$187\pm48$	$191\pm44$	0.8
Glucose (mg/dl)	85 ± 14	$82\pm11$	0.6
Insulin (micU/ml)	$5.6\pm6.1$	3.9 ± 2.7	0.5
HOMA-IR	$1.1 \pm 1.2$	$0.8\pm0.4$	0.48

 114 patients with NCCAH (93 females) aged <18 years at diagnosis.

### **Clinical evaluation**

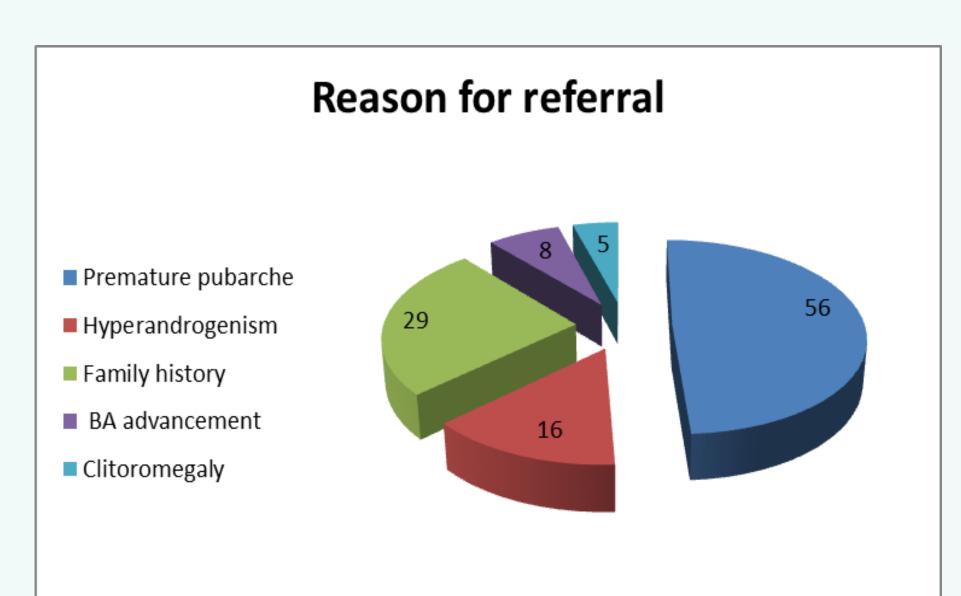
- Weight, height, blood pressure, bio-impedance, waist and hip circumference, skin folds thickness.
- Fasting glucose, insulin, lipid profile and androgens.

### Analyses

- Anthropometric data were compared to their parental measurements.
- Data of patients in grades 7-12 (n=76) were compared to those of the National Health and Nutrition Survey (grades 7-12).
- Currently treated patients were compared to untreated patients (26 previously treated, 11 never treated).

None of the patients were defined as having the metabolic syndrome using IDF consensus criteria, for children and adolescents.

### Results



### Effects of dose and duration of therapy

- Treatment duration was not associated total-, HDL-, and LDL-cholesterol, triglycerides, fasting glucose, HOMA-IR, skin folds thickness or body fat mass.
- Hydrocortisone dose during the 1st year of therapy was associated with slightly higher systolic and diastolic blood pressure.
- Current HC dose was associated only with higher LDL-cholesterol levels (*r*=0.46, *p*<0.05).

## Conclusions

- NCCAH diagnosed in childhood (treated or untreated) is not associated with increased risk of overweight, obesity or metabolic derangements.
- Hydrocortisone dose but not therapy duration may be associated with adverse metabolic effect.
- Larger, long-term studies are needed to confirm our results.

#### Authors have nothing to disclose

P1-P002







