

# DEMOGRAPHIC, CLINICAL AND BIOCHEMICAL CHARACTERISTICS OF PEDIATRIC OBESITY: INTERIM ANALYSIS OF A LARGER PROSPECTIVE STUDY



Maja TANKOSKA<sup>1</sup>, Dejan JAKIMOVSKI<sup>2</sup>, Ana STAMATOVA<sup>1</sup>, Elita MANEVA<sup>1</sup>, Avdi MURTEZANI<sup>1</sup>, Elena SHUKAREVA-ANGELOVSKA<sup>1</sup>, Beti GJURKOVA-ANGELOVSKA<sup>1</sup>, Svetlana KOCEVA<sup>1</sup>, Konstadina KUZEVSKA-MANEVA<sup>3</sup> and Marina KRSTEVSKA-KONSTANTINOVA<sup>1</sup>

<sup>1</sup>Department of Endocrinology and Genetics, University Pediatric Clinic Skopje, North Macedonia; <sup>2</sup>Buffalo Neuroimaging Analysis Center, Department of Neurology, Jacobs Medical School and Biomedical Sciences, University at Buffalo, State University of New York, NY, USA; <sup>3</sup>Department of Cardiology, University Pediatric Clinic, Skopje, North Macedonia

**Introduction:** Pediatric obesity is the most common nutritional disorder that affects more than a third of the young population and predisposes individuals to greater future morbidity and mortality. Therefore, rising obesity epidemics is becoming one the most important healthcare problems.

Our study aimed at determining the demographic, clinical, and biochemical characteristics of patients referred from primary physicians due to concerns of obesity or prediabetes. Determining early signs of premorbid abnormalities would substantially help with creating preventive strategies regarding reduction in pediatric obesity and future obesity-related complications.

**Materials and Methods:** In the period of 2017-2018, 62 consecutive pediatric patients referred to the University Pediatric clinic were recruited. Demographic and clinical information for both the patients and their parents were collected using in-person interview and standardized questionnaires. Specific data regarding weight, height, systolic (SP) and diastolic (DP) blood pressure, lipid metabolic profile, thyroid hormone levels, and glucose and insulin levels before and after oral glucose tolerance test (OGTT with 75g glucose dose) were collected. Body mass index was determined and patients were classified based on the International Obesity Task Force (IOTF) criteria. Appropriate descriptive, comparative parametric and non-parametric tests and Spearman's ranked correlations were used for statistical analyses.

**Results:** The population was consisted of 34 males and 28 females with respective age of 11.6 and 11.8 years old (p=0.781) were recruited. The mean BMI was 30.5 (SD 5.5), of which 8 were with normal weight ( $\leq 25$  BMI), 22 were overweight (25-30 BMI) and 32 were obese ( $\geq 30$  BMI). Patients BMI was significantly associated with the BMI of their parents (Spearman's ranked  $r=0.395$ ,  $p=0.004$ ). Both SP and DP were significantly different between the BMI subgroups (one-way ANOVA  $p=0.005$  and  $p=0.001$ , respectively) with the obese group having the highest BP values (post-hoc Benjamini  $p=0.004$ ).

Obese children had lower average T4 levels when compared to the comparators (7.5  $\mu\text{g/dL}$  vs. 9.9  $\mu\text{g/dL}$ , t-test  $p=0.021$ ). The obese group of children has significantly lower baseline glucose levels and higher insulin levels when compared to the overweight/normal BMI groups (4.1 mmol/L vs. 4.8 mmol/L, t-test  $p<0.001$  and 21.8  $\mu\text{gU/mL}$  vs. 12.9  $\mu\text{gU/mL}$ , Mann-Whitney U test  $p=0.003$ ). Although not significantly different at follow-up, the obese children had greatest numerical increase in glucose levels during the OGTT ( $\Delta 3.5$  mmol/L vs.  $\Delta 2.4$  mmol/L,  $p=0.063$ ) and smaller absolute insulin response ( $\Delta 86.1$   $\mu\text{IU/mL}$  vs.  $\Delta 125.7$   $\mu\text{IU/mL}$ ).

**Table 1.** Demographic characteristics of the total study population and for male and female pediatric patients separately

Demographic and clinical characteristics	Total cohort (n=62)	Female (n=28)	Male (n=34)	Female vs. Male p-value
Female, n (%)	28 (45.2)	-	-	-
Age, mean (SD)	11.6 (2.5)	11.8 (2.7)	11.4 (2.3)	0.538
Weight, mean (SD)	73.4 (20.7)	72.2 (13.1)	74.4 (23.9)	0.678
Height, mean (SD)	153.7 (13.5)	153.4 (13.1)	153.9 (14.0)	0.897
BMI, mean (SD)	30.5 (5.5)	30.4 (4.6)	30.7 (6.2)	0.834
Obese/overweight/normal, n	32/22/8	16/10/2	16/12/6	0.446
Waist circumference, mean (SD)	98.3 (14.7)	97.4 (10.3)	99.0 (17.7)	0.684
Systolic BP, mean (SD)	117.4 (12.9)	117.3 (14.1)	117.5 (12.1)	0.961
Diastolic BP, mean (SD)	76.3 (10.2)	76.1 (10.4)	76.4 (10.4)	0.913
Ejection fraction, mean (SD)	68.4 (7.2)	68.1 (6.9)	68.5 (7.4)	0.835
Maternal BMI, mean (SD)	28.7 (6.2)	28.1 (4.7)	29.5 (7.4)	0.662
Paternal BMI, mean (SD)	30.9 (5.6)	31.3 (6.3)	30.7 (4.9)	0.385

Legend: BMI – body mass index, BP – blood pressure, SD – standard deviation.

Differences were derived by  $\chi^2$  and Student's t-test as appropriate.

**Table 2.** Biochemical and OGTT analysis of the study population

Biochemical analyses and OGTT	Total cohort (n=46)	Normal or overweight (n=22)	Obese (n=24)	Obese vs. other p-value
Cholesterol, mean (SD)	4.1 (0.7)	4.1 (0.6)	4.2 (0.9)	0.54
Total triglycerides, mean (SD)	1.4 (0.7)	1.3 (0.6)	1.4 (0.8)	0.467
T4, mean (SD)	8.5 (3.2)	9.9 (3.1)	7.5 (2.9)	<b>0.021</b>
TSH, mean (SD)	2.9 (1.3)	3.3 (1.1)	2.6 (1.3)	0.155
<i>Oral glucose tolerance test</i>				
Glucose at baseline, mean (SD)	4.4 (0.7)	4.8 (0.6)	4.1 (0.5)	<b>&lt;0.001</b>
Glucose during OGTT, mean (SD)	7.5 (1.8)	7.3 (1.9)	7.6 (1.6)	0.586
Insulin at baseline, mean (SD)	17.7 (9.2)	12.9 (5.5)	21.8 (9.8)	<b>0.003*</b>
Insulin during OGTT, mean (SD)	120.0 (119.7)	134.2 (153.5)	107.9 (82.9)	0.885*

Legend: OGTT – oral glucose tolerance test, TSH – thyroid stimulating hormone, SD – standard deviation, IQR – interquartile range

Differences between the obese and the remaining BMI groups (overweight and normal BMI) were derived with Student's t-test. \* - due to non-parametric nature of the variable Mann-Whitney U test was used. P-value lower than 0.05 was considered statistically significant and shown in bold.

Blood pressure measures are shown as mmHg, T4 levels as  $\mu\text{g/dL}$ , TSH levels as mIU/L, glucose levels as mmol/L, and insulin levels as  $\mu\text{IU/mL}$ .

**Conclusion:** Pediatric patients in our clinic demonstrate familial type of obesity which is characterized with premorbid asymptomatic endocrine impairments. In order to maintain normal glucose levels, obese pediatric patients demonstrate high levels of resting insulin levels and diminished response after OGTT load. Failure of these compensatory mechanisms may lead to early development of diabetes type 2.

## References

1. Arslanian S, Kim JY, Nasr A, Bacha F, Tfayli H, Lee S, et al. Insulin sensitivity across the lifespan from obese adolescents to obese adults with impaired glucose tolerance: Who is worse off? *Pediatr Diabetes*. 2018;19(2):205-11.
2. Atkin AJ, Foley L, Corder K, Ekelund U, van Sluijs EM. Determinants of Three-Year Change in Children's Objectively Measured Sedentary Time. *PLoS One*. 2016;11(12):e0167826.
3. Ferrannini E, Natali A, Bell P, Cavallo-Perin P, Lalic N, Mingrone G. Insulin resistance and hypersecretion in obesity. European Group for the Study of Insulin Resistance (EGIR). *J Clin Invest*. 1997;100(5):1166-73.
4. Gauthier KI, Krajicek MJ. Obesogenic environment: a concept analysis and pediatric perspective. *J Spec Pediatr Nurs*. 2013;18(3):202-10.
5. Goran MI, Gower BA. Longitudinal study on pubertal insulin resistance. *Diabetes*. 2001;50(11):2444-50.
6. Gortmaker SL, Must A, Perrin JM, Sobol AM, Dietz WH. Social and economic consequences of overweight in adolescence and young adulthood. *N Engl J Med*. 1993;329(14):1008-12.
7. Juonala M, Magnussen CG, Berenson GS, Venn A, Burns TL, Sabin MA, et al. Childhood adiposity, adult adiposity, and cardiovascular risk factors. *N Engl J Med*. 2011;365(20):1876-85.
8. Lightwood J, Bibbins-Domingo K, Coxson P, Wang YC, Williams L, Goldman L. Forecasting the future economic burden of current adolescent overweight: an estimate of the coronary heart disease policy model. *Am J Public Health*. 2009;99(12):2230-7.
9. May AL, Kuklina EV, Yoon PW. Prevalence of cardiovascular disease risk factors among US adolescents, 1999-2008. *Pediatrics*. 2012;129(6):1035-41.
10. Ward ZJ, Long MW, Resch SC, Giles CM, Craddock AL, Gortmaker SL. Simulation of Growth Trajectories of Childhood Obesity into Adulthood. *N Engl J Med*. 2017;377(22):2145-53.

