The Major Impact of Obesity on the Development of Type 2 Diabetes (T2D) in Women With PCOS: A Systematic Review and Meta-Analysis of Observational Studies

Sarantis Livadas. Athens Medical Center

Panagiotis Anagnostis, MD, PhD¹, Rodis Paparodis, MD², Julia Bosdou, MD, PhD³, Christina Bothou, MD Msc⁴, Dimitrios G. Goulis, MD, PhD¹, Djuro P. Macut, MD, PhD⁵, Andrea Elizabeth Dunaif, MD⁶, Sarantis Livadas, MD, PhD⁷.

¹Unit of Reproductive Endocrinology, 1st Department of Obstetrics and Gynecology, Medical School, Aristotle University of Thessaloniki, Thessaloniki, Greece, ²Center for Diabetes and Endocrine Research, University of Toledo College of Medicine and Life Sciences, Toledo, OH, USA, ³Unit for Human Reproduction, 1st Department of Obstetrics and Gynecology, Medical School, Aristotle University of Thessaloniki, Thessaloniki, Greece, ⁴Klinik für Endokrinologie, Diabetologie und Klinische Ernährung, Universitätsspital Zürich, Zürich, Switzerland, ⁵Clinic for Endocrinology, Diabetes and Metabolic Diseases, Faculty of Medicine, University of Belgrade, Belgrade, Serbia, ⁶Icahn School of Medicine at Mount Sinai, New York, NY, USA, ⁷Endocrine Unit, Athens Medical Center, Athens, Greece.

Background/Aims: Polycystic ovary syndrome (PCOS) is associated with disordered carbohydrate metabolism and an increased risk for T2D. However, there are limited data on the magnitude of this risk. Furthermore, 50-80% of women with PCOS are obese and obesity is known to have a synergistic deleterious effect on glucose tolerance in affected women. We systematically reviewed the literature regarding the association between PCOS, obesity and T2D risk.

Methods: A comprehensive search was conducted in PubMed, CENTRAL and Scopus databases. Data are expressed as relative risk (RR) with 95% confidence intervals (CI). The I2 index was employed for heterogeneity. The available data, did not allow us to analyze the impact of weight status as normal, overweight and obese and as a consequence the studied subjects were stratified as obese (BMI>30 kg/m2) and non-obese (BMI<30kg/m2).

Results: Twelve studies fulfilled eligibility criteria, yielding a total of 224,284 participants (45,361 PCOS and 5,717 T2DM cases). Women with PCOS had a higher risk of T2D compared with to unaffected women (RR 3.13, 95% CI, 2.83-3.47, p<0.001; I2 40.1%). When women with PCOS were stratified according to the presence or absence of obesity, the RR for developing T2D in obese compared with non-obese women with PCOS was 4.20 (95% CI 1.97-9.10; p<0.001). Moreover, compared to control women, the RR for developing T2D was significantly increased only in obese PCOS, RR 4.06 (95% CI 2.75-5.98; p<0.001). There was a trend toward significantly increased risk in non-obese PCOS women [RR 2.68 (95% CI 0.97-7.49; p=0.06).

Conclusion: Women with PCOS have a >3-fold increased risk of T2D compared to women without PCOS, but this risk is substantially increased by the presence of obesity. Accordingly, weight reduction should be pursued in these women.