



Effects of GLP-1 Receptor Agonists on Oocytes and Female Fertility

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The global rise in obesity among reproductive-age women has increased the need to understand how excess adiposity influences fertility and oocyte quality. Recent studies suggest that glucagon-like peptide-1 receptor agonists (GLP-1 RAs), which have been broadly utilized for metabolic control and weight reduction, may have direct favorable effects on the female reproductive system. This editorial aims to draw attention to the current evidence on the molecular and clinical mechanisms by which GLP-1 RAs may improve oocyte quality, follicular function, and fertility outcomes.

The increasing number of obese or overweight women looking for reproductive assistance has intensified research into the effects of excess weight on oocyte competence and fertility. A recent meta-analysis and systematic review reported that obesity exerts a weak but statistically significant negative impact on assisted reproductive technology (ART) outcomes, including ovarian stimulation duration, the required total gonadotropin dose, and the number of mature oocytes retrieved (1).

Preclinical studies have demonstrated that diet-induced obesity leads to profound ovarian alterations, including depletion of primordial follicles, follicular apoptosis, increased meiotic aneuploidy, reduced follicle number and size, and increased mitochondrial dysfunction within oocytes (2-6). Although dietary interventions can restore systemic metabolic parameters, persistent meiotic and mitochondrial defects suggest that obesity may cause irreversible ovarian injury (7).

Recent evidence suggests that the receptors of GLP-1 are expressed in several components of the human reproductive system (8). Beyond their established metabolic roles, GLP-1 and its analogs may exert anti-fibrotic and anti-inflammatory effects on gonadal and endometrial tissues affected by polycystic ovary syndrome (PCOS), obesity, and diabetes. In PCOS women, GLP-1 RAs have been shown to reverse polycystic ovarian

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morphology and reduce serum androgen concentrations and bioavailability (8).

In a clinical study among obese and PCOS women, combination therapy with metformin and low-dose liraglutide significantly ameliorated both in vitro fertilization (IVF) and spontaneous pregnancy rates compared with metformin monotherapy (9, 10). In the combination group, the higher pregnancy rate per embryo transfer implies that GLP-1 RAs may enhance fertility not only through weight reduction but also via direct reproductive benefits.

At the cellular level, liraglutide has been shown to ameliorate dehydroepiandrosterone (DHEA) -induced follicular dysfunction by decreasing CXCL10 secretion from granulosa cells and suppressing the JAK signaling pathway in PCOS patients (11). CXCL10 (IP-10, interferon- γ -inducible protein 10) is a pro-inflammatory chemokine implicated in follicular dysfunction, oxidative stress, and impaired oocyte maturation. By reducing CXCL10 levels, liraglutide may restore granulosa cell homeostasis and improve oocyte quality. Furthermore, semaglutide has been shown to attenuate autophagy and oxidative stress in oocytes by modulating the pathway of PI3K/AKT/mTOR signaling (12).

Collectively, current evidence suggests that GLP-1 receptor agonists exert positive effects on oocyte competence, follicular function, and fertility outcomes.

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These benefits likely arise from both systemic metabolic improvements and direct anti-inflammatory actions within the ovary. Nonetheless, additional clinical and mechanistic study is needed to elucidate the safety and long-term effects of GLP-1 RAs on female reproductive health.

Competing Interests

None declared.

Ethical Issues

Not applicable.

Authors' Contribution

Conceptualization: Zehra Kurdoğlu.

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Formal analysis: Canan Tapkan.

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References

- Ribeiro LM, Sasaki LMP, Silva AA, et al. Overweight, obesity and assisted reproduction: A systematic review and meta-analysis. *Eur J Obstet Gynecol Reprod Biol* 2022;271:117–127. doi:10.1016/j.ejogrb.2022.01.019
- Jungheim ES, Schoeller EL, Marquard KL, Loudon ED, Schaffer JE, Moley KH. Diet-induced obesity model: abnormal oocytes and persistent growth abnormalities in the offspring. *Endocrinology* 2010;151:4039–4046. doi:10.1210/en.2010-0098
- Luzzo KM, Wang Q, Purcell SH, et al. High fat diet induced developmental defects in the mouse: oocyte meiotic aneuploidy and fetal growth retardation/brain defects. *Plos One*. 2012;7:e49217. doi:10.1371/journal.pone.0049217
- Igosheva N, Abramov AY, Poston L, et al. Maternal diet-induced obesity alters mitochondrial activity and redox status in mouse oocytes and zygotes. *Plos One*. 2010;5:e10074. doi:10.1371/journal.pone.0010074
- Boots CE, Boudoures A, Zhang W, Drury A, Moley KH. Obesity-induced oocyte mitochondrial defects are partially prevented and rescued by supplementation with co-enzyme Q10 in a mouse model. *Hum Reprod*. 2016;31:2090–2097. doi:10.1093/humrep/dew181
- Zhou J, Lin L, Liu L, Wang J, Xia G, Wang C. The transcriptome reveals the molecular regulatory network of primordial follicle depletion in obese mice. *Fertil Steril*. 2023;120:899–910. doi:10.1016/j.fertnstert.2023.05.165
- Reynolds KA, Boudoures AL, Chi MM, Wang Q, Moley KH. Adverse effects of obesity and/or high-fat diet on oocyte quality and metabolism are not reversible with resumption of regular diet in mice. *Reprod Fertil Dev*. 2015;27:716–724. doi:10.1071/RD14251
- Bednarsz K, Kowalczyk K, Cwynar M, et al. The Role of Glp-1 Receptor Agonists in Insulin Resistance with Concomitant Obesity Treatment in Polycystic Ovary Syndrome. *Int J Mol Sci*. 2022;23:4334. doi:10.3390/ijms23084334
- Salamun V, Jensterle M, Janez A, Vrtacnik Bokal E. Liraglutide increases IVF pregnancy rates in obese PCOS women with poor response to first-line reproductive treatments: a pilot randomized study. *Eur J Endocrinol*. 2018;179:1–11. doi:10.1530/EJE-18-0175
- Jensterle M, Herman R, Janez A. Therapeutic Potential of Glucagon-like Peptide-1 Agonists in Polycystic Ovary Syndrome: From Current Clinical Evidence to Future Perspectives. *Biomedicines* 2022;10:1989. doi:10.3390/biomedicines10081989
- Zhao M, Liao B, Yun C, Qi X, Pang Y. Liraglutide improves follicle development in polycystic ovary syndrome by inhibiting CXCL10 secretion. *Reprod Biol Endocrinol*. 2024;22:98. doi:10.1186/s12958-024-01269-9
- Guo S, Li X, Liu M, et al. Semaglutide Alleviates Ovarian Oxidative Stress and Autophagy via the PI3K/AKT/mTOR Pathway in Mice with Polycystic Ovary Syndrome. *Drug Des Devel Ther*. 2025;19:4297–4310. doi:10.2147/DDDT.S522730

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