

# **Chapter 4**

## **NAUSEA AND VOMITING OF PREGNANCY**

**İlknur ÇÖL MADENDAĞ<sup>1</sup>**

Nausea and vomiting is one of the most common complaints seen in pregnancy. These symptoms are generally defined as physiological changes in pregnancy. However, sometimes these may be a harbinger of some disease such as thyroid diseases, cholecystitis, pancreatitis, hepatitis, and psychiatric disorders. These symptoms can affect both the patient and her family, can decrease quality of life, especially when persistent and/or severe (Lacasse, Rey, Ferreira, Morin, & Berard, 2008a). Severe symptoms can negatively affect work performance and daily functioning, and may cause anxiety, and lead to some women considering termination of pregnancy or avoiding a future pregnancy (Tan, Lowe, & Henry, 2018).

### **DEFINITION AND INCIDENCE AND RISK FACTORS**

Nausea and vomiting of pregnancy can be categorized into degrees of severity by assessing the duration of these symptoms and the amount of vomiting per day (Attard et al., 2002). Hyperemesis gravidarum (HEG) is defined as the severe end of the symptom spectrum (including weight loss exceeding 5 percent of pre-pregnancy body weight), whereas morning sickness defined as mild to moderate disease (Koot et al., 2018). The incidence of HEG is about 3% and of all pregnancies have prevalence rates for nausea of 50–80% and for vomiting of 50% (Matthews, Haas, O’Mathuna, & Dowswell, 2015). Ethnic differences and differences in the definition of the disease may account, in part, for the variability. The probability of emergence of these symptoms in the next pregnancy is 15% to 81% (Trogstad, Stoltenberg, Magnus, Skjaerven, & Irgens, 2005). Some clinical conditions such as multiple gestation (Basso & Olsen, 2001), hydatidiform molar pregnancy (Hou, Wan, Xiang, Qi, & Yang, 2008), daughters of women with hyperemesis (Vikanes et al., 2010), and acid reflux or other gastrointestinal disorders may increase the risk of nausea and vomiting (Gill, Maltepe, & Koren, 2009). Using alcohol and cigarettes may decrease the risk of these symptoms (Weigel & Weigel, 1988).

---

<sup>1</sup> MD, Kayseri City Hospital, ilknurmadendag@gmail.com

intravenously every eight hours) (Ramin et al., 2018). Ondansetron can cause some adverse effects such as headache, somnolence, tiredness, and constipation. Also, it may prolong the QT interval. In addition several congenital defects such as cardiac septum defect and cleft palate were found in some studies (Parker, Van Bennekom, Anderka, Mitchell, & Stu, 2018). There are no pregnant women data on the security of granisetron and dolasetron in pregnancy.

### **Steroids**

There is a deficiency of evidence that glucocorticoids are effective. For treatment of refractory to other medications in pregnant women with severe HEG, glucocorticoids may be tried. However, use of glucocorticoids should be avoided in the first trimester, since it may cause oral cleft for fetus (Pradat et al., 2003). Methylprednisolone 16 mg every 8 hours, orally or intravenously, for 3 days (Ramin et al., 2018).

### **PARENTERAL NUTRITION**

Intravenous hydration should be given for the patient when she cannot tolerate oral liquids for a long time or if clinical signs of dehydration are present. Correction of ketosis and vitamin deficiency should be strongly considered. Dextrose and vitamins should be given to the patient. Enteral tube feeding should be admitted as the first-line treatment to provide nutritional support to the woman with hyperemesis gravidarum who is refractory to other medical therapy and cannot maintain her weight (Erick et al., 2018).

### **Hospitalization**

When a woman cannot tolerate fluids due to hyperemesis and cannot respond to outpatient treatment, hospitalization is suggested for evaluation and treatment of dehydration and electrolyte imbalance (McCarthy et al., 2014).

### **REFERENCES**

- Abas, M. N., Tan, P. C., Azmi, N., & Omar, S. Z. (2014). Ondansetron Compared With Metoclopramide for Hyperemesis Gravidarum A Randomized Controlled Trial. *Obstetrics and Gynecology*, 123 (6), 1272-1279. doi:10.1097/AOG.0000000000000242
- Atmaca, U., Atalay, M. A., Ozcimen, N., Sidal, B., & Ates, U. (2015). Investigating Maternal Serum Thyroid Hormone, Beta-Human Chorionic Gonadotropin (Beta-HCG) and Free Beta-HCG Levels in Hyperemesis Gravidarum. *Erciyes Medical Journal*, 37 (4), 128-132. doi:10.5152/etd.2015.6577
- Attard, C. L., Kohli, M. A., Coleman, S., Bradley, C., Hux, M., Atanackovic, G., & Torrance, G. W. (2002). The burden of illness of severe nausea and vomiting of pregnancy in the United States. *Am J Obstet Gynecol*, 186 (5), S220-S227. doi:10.1067/mob.2002.122605

## *Obstetrics and Gynecology I*

- Basso, O., & Olsen, J. (2001). Sex ratio and twinning in women with hyperemesis or pre-eclampsia. *Epidemiology*, 12 (6), 747-749. doi:Doi 10.1097/00001648-200111000-00026
- Bischoff, S. C., & Renzer, C. (2006). Nausea and nutrition. *Autonomic Neuroscience-Basic & Clinical*, 129 (1-2), 22-27. doi:10.1016/j.autneu.2006.07.011
- Bustos, M., Venkataraman, R., & Caritis, S. (2017). Nausea and vomiting of pregnancy - What's new? *Auton Neurosci*, 202, 62-72. doi:10.1016/j.autneu.2016.05.002
- Czeizel, A. E., Dudas, I., Fritz, G., Tecsoi, A., Hanck, A., & Kunovits, G. (1992). The Effect of Periconceptional Multivitamin-Mineral Supplementation on Vertigo, Nausea and Vomiting in the 1st Trimester of Pregnancy. *Archives of Gynecology and Obstetrics*, 251 (4), 181-185. doi:Doi 10.1007/Bf02718384
- Davis, M. (2004). Nausea and vomiting of pregnancy - An evidence-based review. *Journal of Perinatal & Neonatal Nursing*, 18 (4), 312-328. doi:Doi 10.1097/00005237-200410000-00002
- Depue, R. H., Bernstein, L., Ross, R. K., Judd, H. L., & Henderson, B. E. (1987). Hyperemesis Gravidarum in Relation to Estradiol Levels, Pregnancy Outcome, and Other Maternal Factors - a Seroepidemiologic Study. *Am J Obstet Gynecol*, 156 (5), 1137-1141. doi:Doi 10.1016/0002-9378 (87)90126-8
- Dochez, V., Dimet, J., David-Gruselle, A., Le Thuaut, A., & Ducarme, G. (2016). Validation of specific questionnaires to assess nausea and vomiting of pregnancy in a French population. *International Journal of Gynecology & Obstetrics*, 134 (3), 294-298. doi:10.1016/j.ijgo.2016.01.023
- Einarson, A., Koren, G., & Bergman, U. (1998). Nausea and vomiting in pregnancy: A comparative European study. *European Journal of Obstetrics Gynecology and Reproductive Biology*, 76 (1), 1-3. doi:Doi 10.1016/S0301-2115 (97)00129-2
- Erick, M., Cox, J. T., & Mogensen, K. M. (2018). ACOG Practice Bulletin 189: Nausea and Vomiting of Pregnancy. *Obstetrics and Gynecology*, 131 (5), 935-935. doi:10.1097/AOG.0000000000002604
- Furneaux, E. C., Langley-Evans, A. J., & Langley-Evans, S. C. (2001). Nausea and vomiting of pregnancy: Endocrine basis and contribution to pregnancy outcome. *Obstetrical & Gynecological Survey*, 56 (12), 775-782. doi:Doi 10.1097/00006254-200112000-00004
- Gill, S. K., Maltepe, C., & Koren, G. (2009). The effect of heartburn and acid reflux on the severity of nausea and vomiting of pregnancy. *Canadian Journal of Gastroenterology and Hepatology*, 23 (4), 270-272. doi:Doi 10.1155/2009/678514
- Goldman, A. M., & Mestman, J. H. (2011). Transient non-autoimmune hyperthyroidism of early pregnancy. *J Thyroid Res*, 2011, 142413. doi:10.4061/2011/142413
- Goodwin, T. M. (2008). Hyperemesis gravidarum. *Obstet Gynecol Clin North Am*, 35 (3), 401-417, viii. doi:10.1016/j.ogc.2008.04.002
- Heitmann, K., Nordeng, H., Havnen, G. C., Solheimsnes, A., & Holst, L. (2017). The burden of nausea and vomiting during pregnancy: severe impacts on quality of life, daily life functioning and willingness to become pregnant again - results from a cross-sectional study. *Bmc Pregnancy and Childbirth*, 17. doi:ARTN 75 10.1186/s12884-017-1249-0
- Hinkle, S. N., Mumford, S. L., Grantz, K. L., Silver, R. M., Mitchell, E. M., Sjaarda, L. A., . . . Schisterman, E. F. (2016). Association of Nausea and Vomiting During Pregnancy With Pregnancy Loss A Secondary Analysis of a Randomized Clinical Trial. *Jama Internal Medicine*, 176 (11), 1621-1627. doi:10.1001/jamainternmed.2016.5641
- Hou, J. L., Wan, X. R., Xiang, Y., Qi, Q. W., & Yang, X. Y. (2008). Changes of Clinical Features in Hydatidiform Mole Analysis of 113 Cases. *Journal of Reproductive Me-*

## *Obstetrics and Gynecology I*

- dicine, 53 (8), 629-633.*
- Kimura, M., Amino, N., Tamaki, H., Ito, E., Mitsuda, N., Miyai, K., & Tanizawa, O. (1993). Gestational Thyrotoxicosis and Hyperemesis Gravidarum - Possible Role of Hcg with Higher Stimulating Activity. *Clinical Endocrinology, 38* (4), 345-350. doi:DOI 10.1111/j.1365-2265.1993.tb00512.x
- Kirshon, B., Lee, W., & Cotton, D. B. (1988). Prompt Resolution of Hyperthyroidism and Hyperemesis Gravidarum after Delivery. *Obstetrics and Gynecology, 71* (6), 1032-1034.
- Koot, M. H., Boelig, R., van't Hooft, J., Limpens, J., Roseboom, T. J., Painter, R. C., & Grooten, I. J. (2018). Variation in hyperemesis gravidarum definition and outcome reporting in randomised clinical trials: a systematic review. *Bjog-an International Journal of Obstetrics and Gynaecology, 125* (12), 1514-1521. doi:10.1111/1471-0528.15272
- Lacasse, A., Rey, E., Ferreira, E., Morin, C., & Berard, A. (2008a). Nausea and vomiting of pregnancy: what about quality of life? *Bjog-an International Journal of Obstetrics and Gynaecology, 115* (12), 1484-1493. doi:10.1111/j.1471-0528.2008.01891.x
- Lacasse, A., Rey, E., Ferreira, E., Morin, C., & Berard, A. (2008b). Validity of a modified Pregnancy-Unique Quantification of Emesis and Nausea (PUQE) scoring index to assess severity of nausea and vomiting of pregnancy. *Am J Obstet Gynecol, 198* (1). doi:ARTN 71.e1  
10.1016/j.ajog.2007.05.051
- Macfie, A. G., Magides, A. D., Richmond, M. N., & Reilly, C. S. (1991). Gastric-Emptrying in Pregnancy. *British Journal of Anaesthesia, 67* (1), 54-57. doi:DOI 10.1093/bja/67.1.54
- Madjunkova, S., Maltepe, C., & Koren, G. (2014). The Delayed-Release Combination of Doxylamine and Pyridoxine (Diclegis (A (R))/Diclectin (A (R))) for the Treatment of Nausea and Vomiting of Pregnancy. *Pediatric Drugs, 16* (3), 199-211. doi:10.1007/s40272-014-0065-5
- Magee, L. A., Mazzotta, P., & Koren, G. (2002). Evidence-based view of safety and effectiveness of pharmacologic therapy for nausea and vomiting of pregnancy (NVP). *Am J Obstet Gynecol, 186* (5), S256-S261. doi:10.1067/mob.2002.122596
- Matok, I., Gorodischer, R., Koren, G., Sheiner, E., Wiznitzer, A., & Levy, A. (2009). The Safety of Metoclopramide Use in the First Trimester of Pregnancy EDITORIAL COMMENT. *Obstetrical & Gynecological Survey, 64* (10), 637-638. doi:DOI 10.1097/01.ogx.0000359282.75337.83
- Matthews, A., Haas, D. M., O'Mathuna, D. P., & Dowswell, T. (2015). Interventions for nausea and vomiting in early pregnancy. *Cochrane Database of Systematic Reviews* (9). doi:ARTN CD007575  
10.1002/14651858.CD007575.pub4
- McCarthy, F. P., Murphy, A., Khashan, A. S., McElroy, B., Spillane, N., Marchocki, Z., . . Higgins, J. R. (2014). Day Care Compared With Inpatient Management of Nausea and Vomiting of Pregnancy A Randomized Controlled Trial. *Obstetrics and Gynecology, 124* (4), 743-748. doi:10.1097/AOG.0000000000000449
- Niemeijer, M. N., Grooten, I. J., Vos, N., Bais, J. M. J., van der Post, J. A., Mol, B. W., . . Painter, R. C. (2014). Diagnostic markers for hyperemesis gravidarum: a systematic review and metaanalysis. *Am J Obstet Gynecol, 211* (2). doi:ARTN 150.e1  
10.1016/j.ajog.2014.02.012
- Parker, S. E., Van Bennekom, C., Anderka, M., Mitchell, A. A., & Stu, N. B. D. P. (2018). Ondansetron for Treatment of Nausea and Vomiting of Pregnancy and the Risk of Specific Birth Defects. *Obstetrics and Gynecology, 132* (2), 385-394. doi:10.1097/

## *Obstetrics and Gynecology I*

- Aog.0000000000002679
- Power, M. L., Holzman, G. B., & Schulkin, J. (2001). A survey on the management of nausea and vomiting in pregnancy by obstetrician/gynecologists. *Prim Care Update Ob Gyns*, 8 (2), 69-72.
- Practice Bulletin No. 187: Neural Tube Defects. (2017). *Obstet Gynecol*, 130 (6), e279-e290. doi:10.1097/aog.0000000000002412
- Pradat, P., Robert-Gnansia, E., Di Tanna, G. L., Rosano, A., Lisi, A., Mastroiacovo, P., & Database, M. (2003). First trimester exposure to corticosteroids and oral clefts. *Birth Defects Research Part a-Clinical and Molecular Teratology*, 67 (12), 968-970. doi:10.1002/bdra.10134
- Ramin, S. M., Turrentine, M. A., & Zahn, C. M. (2018). ACOG Practice Bulletin 189: Nausea and Vomiting of Pregnancy Reply. *Obstetrics and Gynecology*, 131 (5), 935-936. doi:10.1097/AOG.0000000000002605
- Sherman, P. W., & Flaxman, S. M. (2002). Nausea and vomiting of pregnancy in an evolutionary perspective. *Am J Obstet Gynecol*, 186 (5), S190-S197. doi:10.1067/mob.2002.122593
- Slaughter, S. R., Hearn-Stokes, R., van der Vlugt, T., & Joffe, H. V. (2014). FDA Approval of Doxylamine-Pyridoxine Therapy for Use in Pregnancy. *New England Journal of Medicine*, 370 (12), 1081-1083. doi:10.1056/NEJMmp1316042
- Tan, A., Lowe, S., & Henry, A. (2018). Nausea and vomiting of pregnancy: Effects on quality of life and day-to-day function. *Australian & New Zealand Journal of Obstetrics & Gynaecology*, 58 (3), 278-290. doi:10.1111/ajo.12714
- Trogstad, L. I. S., Stoltenberg, C., Magnus, P., Skjaerven, R., & Irgens, L. M. (2005). Recurrence risk in hyperemesis gravidarum. *Bjog-an International Journal of Obstetrics and Gynaecology*, 112 (12), 1641-1645. doi:10.1111/j.1471-0528.2005.00765.x
- Tsuruta, E., Tada, H., Tamaki, H., Kashiwai, T., Asahi, K., Takeoka, K., . . . Amino, N. (1995). Pathogenic Role of Asialo Human Chorionic-Gonadotropin in Gestational Thyrotoxicosis. *Journal of Clinical Endocrinology & Metabolism*, 80 (2), 350-355. doi:DOI 10.1210/jc.80.2.350
- Vandraas, K. F., Vikanes, A. V., Vangen, S., Magnus, P., Stoer, N. C., & Grjibovski, A. M. (2013). Hyperemesis gravidarum and birth outcomes-a population-based cohort study of 2.2 million births in the Norwegian Birth Registry. *Bjog-an International Journal of Obstetrics and Gynaecology*, 120 (13), 1654-1660. doi:10.1111/1471-0528.12429
- Veenendaal, M. V. E., van Abeelen, A. F. M., Painter, R. C., van der Post, J. A. M., & Roseboom, T. J. (2011). Consequences of hyperemesis gravidarum for offspring: a systematic review and meta-analysis. *Bjog-an International Journal of Obstetrics and Gynaecology*, 118 (11), 1302-1313. doi:10.1111/j.1471-0528.2011.03023.x
- Vikanes, A., Skjaerven, R., Grjibovski, A. M., Gunnes, N., Vangen, S., & Magnus, P. (2010). Recurrence of hyperemesis gravidarum across generations: population based cohort study. *British Medical Journal*, 340. doi:ARTN c2050  
10.1136/bmj.c2050
- Vutyavanich, T., Supreeya, W., & Ruangsri, R. (1995). Pyridoxine for Nausea and Vomiting of Pregnancy - a Randomized, Double-Blind, Placebo-Controlled Trial. *Am J Obstet Gynecol*, 173 (3), 881-884. doi:Doi 10.1016/0002-9378 (95)90359-3
- Weigel, M. M., & Weigel, R. M. (1988). The Association of Reproductive History, Demographic-Factors, and Alcohol and Tobacco Consumption with the Risk of Developing Nausea and Vomiting in Early-Pregnancy. *American Journal of Epidemiology*, 127 (3), 562-570. doi:DOI 10.1093/oxfordjournals.aje.a114831