

International Journal of Reproductive Medicine & Gynecology

Research Article

Dyslipidemia and Preeclampsia: Pathogenesis or Co-Incidence? - 👌

Ramin Kheirjou^{1*} and Sahar Soltani²

¹Departament of Medicine, Tehran University of Medical Sciences, Tehran, Iran ²Departament of Medicine, Zanjan University of Medical Sciences, Zanjan, Iran

*Address for Correspondence: Ramin Kheirjou, Tehran University of Medical sciences, Enqelab Square, Tehran, Iran, Tel: +98-912-444-096-02; E-mail: kheirjou.ramin@gmail.com

Submitted: 24 February 2021; Approved: 06 April 2021; Published: 07 April 2021

Cite this article: Kheirjou R, Soltani S. Dyslipidemia and Preeclampsia: Pathogenesis or Co-Incidence? Int J Reprod Med Gynecol. 2021 April 07;7(1): 006-09.

Copyright: © 2021 Kheirjou R, et al. This is an open access article distributed under the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

ISSN: 2640-0944

ABSTRACT

Background: Recently the role of lipids in the pathogenesis of preeclampsia is considered. We aimed to assess the association between maternal and cord blood lipid and preeclampsia, especially in pregnant women without any risk factors.

Methods: This cross-sectional study was conducted from 2014 to 2016 on 116 terms pregnant women in an educational hospital in Zanjan- Iran. All participants were 18-35 years of age. The study was approved by the Ethics Committee of Zanjan University of Medical Sciences, and written informed consent was obtained from the entire participant. A total of 58 pregnant women with preeclampsia and 58 healthy pregnant women were included in the study. Cord blood samples were collected immediately after the caesarian section and were taken from the umbilical vein. Controls, which were selected from the study population, tried to be matched to cases by age, sex of a fetus, and gestational week. We tried to reduce confounding factors as possible by selecting our participants carefully.

Results: There were no significant differences in maternal age, birth weight, and neonatal gender between the two groups. Serum total cholesterol, Triglycerides (TG), Low-Density Lipoprotein Cholesterol (LDL-C), and High-Density Lipoprotein Cholesterol (HDL-C) concentrations were significantly lower in preeclamptic women than in the healthy pregnant women, but no difference was observed in cord blood sample between groups.

Conclusion: Our findings suggest a relationship between low lipid value and preeclampsia. We think prepregnancy factors might be involved in dyslipidemia associated with Preeclampsia. Further studies are needed to explore potential mechanisms.

Keywords: Hyperlipidemia; Preeclampsia; Pregnancy

INTRODUCTION

Preeclampsia is a new onset of hypertension diagnosed after 20 weeks of gestation and is estimated to complicate 5% - 8% of pregnancies. It has been defined by a blood pressure greater than 140/90 mmHg and proteinuria [1-3]. Preeclampsia is a major cause of maternal mortality and morbidity and also strongly associated with 500,000 infant mortality annually [4]. Impaired trophoblast differentiation and invasion is the characteristic point of preeclampsia that leads to high resistance placental arterial and more vasoconstriction and ischemia and oxidative stress [5,6]. The exact etiology of this disease remains unknown [7]. However, risk factors include Antiphospholipid syndrome, Previous preeclampsia, Insulindependent diabetes, Multiple pregnancies, Null parity, Family history of preeclampsia, Age >40 years and Preexisting hypertension [8,9]. Recently the role of lipids in the pathogenesis of preeclampsia is considered [10-12]. Lipoprotein metabolism disorders are known as the main cause of endothelial damage that leads to hypertension [13] for example plasma high triglycerides could damage vascular function by prothrombotic activity [14]. We think that the possibility of anticipating preeclampsia could enable the physician to plan better management of this disease which still represents a major challenge in prenatal medicine. This study aimed to evaluate maternal and umbilical cord serum lipid profiles between healthy pregnant women and preeclamptic patients.

MATERIALS AND METHODS

A cross-sectional study was designed and 116 women with pregnancies between 38 and 41 weeks of gestation were investigated. The patient group consisted of 58 women with preeclampsia. The control group consisted of 58 pregnant healthy women. Preeclampsia was diagnosed by blood pressure more than 140/90 mmHg on 2 measurements by 6 hours interval and with proteinuria of 100 mg/ dl by urine analysis or 300 mg in a 24-h urine collection after 20 weeks of gestation [15]. Inclusion criteria were women aged 18-35 years with pregnancies between 38 and 41 weeks of gestation and normal prepregnancy BMI without confounding factors like smoking, multiparity, use of alcohol, intensive work, PROM, diabetes mellitus, chronic hypertension, heart failure, inflammatory or infective disorders, renal disease, pregnant with donor eggs, history of previous preeclampsia in participants or first-degree relative

females and women who in labor. All of the participants were being supplemented with multivitamins during pregnancy according to the doctor's advice. All women delivered *via* elective cesarean section. Maternal samples were obtained after 8 hours of fasting and exactly before going to the operating room. Umbilical cord blood was collected at the time of delivery. The samples were analyzed within four hours of collection, using the enzymatic method.

Statistical analyses

Independent samples T-tests were conducted to analyze differences in lipid profile levels between cases and controls using SPSS version 16. The results were expressed as the mean \pm standard deviation. The value was assumed to be significant at a p < 0.05.

Ethical approval

Informed consent was obtained from the participants in the study, and ethical approval was granted by the Zanjan University of Medical Science Ethics Committee.

RESULTS

A total of 116 women enrolled in the study including 58 patients with preeclampsia as group 1 and 58 controls as group 2. The mean maternal age was 28.3 ± 5.5 and 26.8 ± 5.3 for group 1 and group 2 respectively 34.5 percent of group 1 and 53.4 percent of group 2 was primigravid that was a significant difference (p = 0.04).

According to birth sex, 62 % of group 1 was male while 55% of group 2 was male. The mean birth weight in case and control were 3190 ± 354.3 gr, 3236.3 ± 377.2 gr. In terms of maternal age, birth sex, and birth weight there were no significant differences between the two groups, while in terms of gravid, there were significant differences between two groups; 21% of healthy mothers and 34% of preeclamptic mothers was multigravida (p = 0.01) Maternal plasma lipid concentrations were measured and presented as mean \pm SD in table 1.

According to table 2, all variables in healthy women were higher than preeclamptic women but there were no significant differences regarding plasma TG and HDL level between two groups, whereas serum total cholesterol and LDL was found to be significantly higher in healthy pregnant women (p < 0.05)

ISSN: 2640-0944

On the other hand in the umbilical cord blood samples exposed to preeclampsia, lipid levels were more than the control group except for HDL which was almost equal in the two groups, however, there was no statistically significant difference.

Table 1: Comparing mean ± SD of lipid profile in the maternal blood (mg/dl).				
Variable	Preeclamptic	Healthy	p value	
TG	188.5 ± 68.9	199.6 ± 64.2	0.26	
TC	222.3 ± 48.5	253.8 ± 74.4	0.03	
HDL	48.9 ± 16.7	49.3 ± 11.3	0.17	
LDL	105.9 ± 34.5	129.4 ± 51.4	0.02	
TG: Trialyceride:]	C: Total Cholesterol	I DI · Low-Density I	inoprotein: HDI ·	

Friglyceride; IC: Total Cholesterol; LDL: Low High-Density Lipoprotein.

Variable	Preeclamptic	Healthy	<i>p</i> value	
TG	38.5 ± 24.8	34 ± 22.1	0.34	
TC	84 ± 27	77.5 ± 26.7	0.19	
HDL	27 ± 8.6	27.9 ± 11.2	0.78	
LDL	26.1 ± 11.5	26.1 ± 11	0.91	
TG: Triglyceride: TC: Total Cholesterol: I DI : Low-Density Lipoprotein: HDI :				

G: Triglyceride; TC: Total Cholesterol; LDL: Low-Density Lipoprotein; HDL: High-Density Lipoprotein.

DISCUSSION

In the present study, we evaluated lipid levels in cord and plasma of women with normal pregnancy and preeclampsia. The results of our study showed that serum total cholesterol and low-density lipoprotein cholesterol level was significantly different between the two groups. Also lipid level in the cord of preeclamptic mothers was higher than in the healthy group but this difference was not significant. These findings are consistent with results from some previous studies [16,17] Furthermore, lipid level in the multigravid group was lower than primigravid group but that was not significant. Our data showed that lipid values in pregnant women without preeclampsia were significantly lower than those found in pregnant women with preeclampsia.

Use of vitamin C and vitamin E can cause a reduction in lipid values, and some studies suggest that this may be one of the reasons for the lower lipid levels [18-20] observed during pregnancy in our study, however, we think vitamins could not explain these differences, since women with and without preeclampsia both take same vitamins. We tried to reduce confounding factor and select our participants carefully so by considering that diet and nutrition can affect lipid level [21,22]. We believe that such differences in lipid profile values in our research are more likely to be caused by other unexpected reasons like being more careful about nutrition and diet in preeclamptic mothers. On the other hand, many previous studies documented high lipid level in preeclamptic cases but unlike our study, almost any of them didn't exclude patients with risk factors from their study and as we know some of these risk factors for example prepregnancy BMI could be one of reasons of dyslipidemia in that studies [23,24] so we thought this difference between our results and their's could be a cause of unknown predisposing factors that were excluded in our study. Moreover in our study participation's delivery was done by elective cesarean and as some studies show elective cesarean delivery's stress is lower than vaginal delivery's and it could reduce lipid level especially in cord samples [25,26].

Our study did have a limitation. Firstly, it was a cross-sectional study and the association between lipid profile and the maternal and fetal adverse consequence could not be evaluated. Secondly, our sample of Iranian pregnant women may differ from other populations, although our results are in accordance with those of other studies done in different countries [27-29] third, this study was performed at a single center and uses a very small sample population. Selective bias may be unavoidable. The existence of maternal predisposing factors like diet seems to be necessary to find out why some preeclamptic women have dyslipidemia and why others have not.

CONCLUSION

Based on these results, we conclude that maternal lipid levels don't increase in preeclampsia always; this appears it depends on predisposing factors. Further molecular case-control studies are needed to evaluate the value of lipid level for the detection of preeclampsia.

REFERENCES

- 1. Steegers EA, von Dadelszen P, Duvekot JJ, Pijnenborg R. Pre-eclampsia. Lancet. 2010 Aug 21;376(9741):631-44. doi: 10.1016/S0140-6736(10)60279-6. Epub 2010 Jul 2. PMID: 20598363.
- 2. Report of the National High Blood Pressure Education Program Working Group on High Blood Pressure in Pregnancy. Am J Obstet Gynecol. 2000 Jul;183(1):S1-S22. PMID: 10920346.
- 3. Obstetricians ACo, Gynecologists, editors. Task Force on Hypertension in Pregnancy. Hypertension in pregancy. LIBRARY OF CONGRESS CATALOGINGaINa PUBLICATION DATA; 2013.
- 4. Khan KS, Wojdyla D, Say L, Gülmezoglu AM, Van Look PF. WHO analysis of causes of maternal death: a systematic review. Lancet. 2006 Apr 1;367(9516):1066-1074. doi: 10.1016/S0140-6736(06)68397-9. PMID: 16581405.
- 5. Pijnenborg R, Vercruysse L, Hanssens M. The uterine spiral arteries in human pregnancy: facts and controversies. Placenta. 2006 Sep-Oct;27(9-10):939-58. doi: 10.1016/j.placenta.2005.12.006. Epub 2006 Feb 20. PMID: 16490251.
- 6. Roberts JM, editor. Endothelial dysfunction in preeclampsia. Seminars in reproductive endocrinology;1998: Copyright© 1998 by Thieme Medical Publishers Inc.
- 7. Staff AC, Benton SJ, von Dadelszen P, Roberts JM, Taylor RN, Powers RW, Charnock-Jones DS, Redman CW. Redefining preeclampsia using placentaderived biomarkers. Hypertension. 2013 May;61(5):932-42. doi: 10.1161/ HYPERTENSIONAHA.111.00250. Epub 2013 Mar 4. PMID: 23460278.
- 8. English FA, Kenny LC, McCarthy FP. Risk factors and effective management of preeclampsia. Integr Blood Press Control. 2015 Mar 3;8:7-12. doi: 10.2147/ IBPC.S50641. PMID: 25767405; PMCID: PMC4354613.
- 9. Kaaja R, editor Insulin resistance syndrome in preeclampsia. Seminars in reproductive endocrinology; 1998: Copyright© 1998 by Thieme Medical Publishers. Inc.
- 10. Enquobahrie DA, Williams MA, Butler CL, Frederick IO, Miller RS, Luthy DA. Maternal plasma lipid concentrations in early pregnancy and risk of preeclampsia. Am J Hypertens. 2004 Jul;17(7):574-81. doi: 10.1016/j. amjhyper.2004.03.666. PMID: 15233976.
- 11. Brown SH, Eather SR, Freeman DJ, Meyer BJ, Mitchell TW. A Lipidomic Analysis of Placenta in Preeclampsia: Evidence for Lipid Storage. PLoS One. 2016 Sep 29;11(9):e0163972. doi: 10.1371/journal.pone.0163972. PMID: 27685997: PMCID: PMC5042456.
- 12. Gratacós E. Lipid-mediated endothelial dysfunction: a common factor to preeclampsia and chronic vascular disease. Eur J Obstet Gynecol Reprod Biol. 2000 Sep;92(1):63-6. doi: 10.1016/s0301-2115(00)00427-9. PMID: 10986436.

International Journal of Reproductive Medicine & Gynecology

- Winkler K, Wetzka B, Hoffmann MM, Friedrich I, Kinner M, Baumstark MW, Zahradnik HP, Wieland H, März W. Triglyceride-rich lipoproteins are associated with hypertension in preeclampsia. J Clin Endocrinol Metab. 2003 Mar;88(3):1162-6. doi: 10.1210/jc.2002-021160. PMID: 12629100.
- 14. Lewis GF, Steiner G. Hypertriglyceridemia and its metabolic consequences as a risk factor for atherosclerotic cardiovascular disease in non-insulindependent diabetes mellitus. Diabetes Metab Rev. 1996 Apr;12(1):37-56. doi: 10.1002/(SICI)1099-0895(199603)12:1<37::AID-DMR154>3.0.CO;2-Q. PMID: 8861500.
- ACOG Committee on Obstetric Practice. ACOG practice bulletin. Diagnosis and management of preeclampsia and eclampsia. Number 33, January 2002. American College of Obstetricians and Gynecologists. Int J Gynaecol Obstet. 2002 Apr;77(1):67-75. PMID: 12094777.
- Barat S, Basirat Z. Association of Preeclampsia with Lipid Concentration of Maternal Plasma and Umbilical Cord. Journal of Mazandaran University of Medical Sciences. 2012;22(88):96-101. https://bit.ly/3cTikrp
- Rodie VA, Caslake MJ, Stewart F, Sattar N, Ramsay JE, Greer IA, Freeman DJ. Fetal cord plasma lipoprotein status in uncomplicated human pregnancies and in pregnancies complicated by pre-eclampsia and intrauterine growth restriction. Atherosclerosis. 2004 Sep;176(1):181-7. doi: 10.1016/j. atherosclerosis.2004.04.026. PMID: 15306192.
- Catalgol B, Ozer NK. Protective effects of vitamin E against hypercholesterolemia-induced age-related diseases. Genes Nutr. 2012 Jan;7(1):91-8. doi: 10.1007/s12263-011-0235-9. Epub 2011 May 18. PMID: 21590435; PMCID: PMC3250520.
- Porkkala-Sarataho E, Salonen JT, Nyyssönen K, Kaikkonen J, Salonen R, Ristonmaa U, Diczfalusy U, Brigelius-Flohe R, Loft S, Poulsen HE. Long-term effects of vitamin E, vitamin C, and combined supplementation on urinary 7-hydro-8-oxo-2'-deoxyguanosine, serum cholesterol oxidation products, and oxidation resistance of lipids in nondepleted men. Arterioscler Thromb Vasc Biol. 2000 Sep;20(9):2087-93. doi: 10.1161/01.atv.20.9.2087. PMID: 10978253.
- Rafighi Z. Arab S, Yusof RM, Shiva A. The Effect of vitamin C and E on lipid profile in people with type 2 Diabetes Mellitus. Global Journal of Health Science. 2011;3(2):69. https://bit.ly/3sVB0MK
- Khoury J, Henriksen T, Christophersen B, Tonstad S. Effect of a cholesterollowering diet on maternal, cord, and neonatal lipids, and pregnancy outcome: a randomized clinical trial. Am J Obstet Gynecol. 2005 Oct;193(4):1292-301. doi: 10.1016/j.ajog.2005.05.016. PMID: 16202717.

- Montoudis A, Simoneau L, Brissette L, Forest JC, Savard R, Lafond J. Impact of a cholesterol enriched diet on maternal and fetal plasma lipids and fetal deposition in pregnant rabbits. Life Sci. 1999;64(26):2439-50. doi: 10.1016/ s0024-3205(99)00201-5. PMID: 10403503.
- Samuels-Kalow ME, Funai EF, Buhimschi C, Norwitz E, Perrin M, Calderon-Margalit R, Deutsch L, Paltiel O, Friedlander Y, Manor O, Harlap S. Prepregnancy body mass index, hypertensive disorders of pregnancy, and long-term maternal mortality. Am J Obstet Gynecol. 2007 Nov;197(5):490. e1-6. doi: 10.1016/j.ajog.2007.04.043. Epub 2007 Aug 21. PMID: 17714679; PMCID: PMC2100395.
- Thadhani R, Stampfer MJ, Hunter DJ, Manson JE, Solomon CG, Curhan GC. High body mass index and hypercholesterolemia: risk of hypertensive disorders of pregnancy. Obstet Gynecol. 1999 Oct;94(4):543-50. doi: 10.1016/s0029-7844(99)00400-7. PMID: 10511356.
- 25. Paamoni-Keren O, Silberstein T, Burg A, Raz I, Mazor M, Saphier O, Weintraub AY. Oxidative stress as determined by glutathione (GSH) concentrations in venous cord blood in elective cesarean delivery versus uncomplicated vaginal delivery. Arch Gynecol Obstet. 2007 Jul;276(1):43-6. doi: 10.1007/s00404-006-0304-2. Epub 2007 Feb 28. Erratum in: Arch Gynecol Obstet. 2007 Dec;276(6):661. Weintraub, Adi Yekuda [added]. PMID: 17333227.
- Rogers MS, Mongelli JM, Tsang KH, Wang CC, Law KP. Lipid peroxidation in cord blood at birth: the effect of labour. Br J Obstet Gynaecol. 1998 Jul;105(7):739-44. doi: 10.1111/j.1471-0528.1998.tb10204.x. PMID: 9692414.
- Pardo IM, Geloneze B, Tambascia MA, Barros-Filho AA. Atherogenic lipid profile of Brazilian near-term newborns. Braz J Med Biol Res. 2005 May;38(5):755-60. doi: 10.1590/s0100-879x2005000500013. Epub 2005 May 25. PMID: 15917957.
- Ophir E, Oettinger M, Nisimov J, Hirsch Y, Fait V, Dourleshter G, Shnaider O, Snitkovsky T, Bornstein J. Cord blood lipids concentrations and their relation to body size at birth: possible link between intrauterine life and adult diseases. Am J Perinatol. 2004 Jan;21(1):35-40. doi: 10.1055/s-2004-820508. PMID: 15017481.
- Nayak CD, Agarwal V, Nayak DM. Correlation of cord blood lipid heterogeneity in neonates with their anthropometry at birth. Indian J Clin Biochem. 2013 Apr;28(2):152-7. doi: 10.1007/s12291-012-0252-5. Epub 2012 Aug 28. PMID: 24426201; PMCID: PMC3613497.

ISSN: 2640-0944