Does increased Body Mass Index increase the risk of recurrent pregnancy loss?

Saleemah Abdul Majeed Omar Alya Abdul-Rahman Sharef Awara Ahmed Rashid1 A

Gynecology and Obstetrics Department / Azadi Teaching Hospital, Kirkuk governorate, Iraq

Corresponding author:

Saleemah Abdul Majeed Omar Gynaecology and Obstetrics Department, Azadi Teaching Hospital, Kirkuk governorate, Iraq **Email:** saleemahomar@yahoo.com

Received: January 2020; Accepted: February 2020; Published: March 1, 2020. Citation: Saleemah Abdul Majeed Omar, Alya Abdul-Rahman Sharef, Awara Ahmed Rashid. Does increased Body mass index increase the risk of recurrent pregnancy loss?. World Family Medicine. 2020; 18(3): 57-61 DOI: 10.5742MEWFM.2020.93774

Abstract

Background: Obesity is a worldwide health problem. It is also associated with maternal and fetal outcome, such as sudden and unexplained intrauterine death, and diabetes, and polycystic ovary syndrome (PCOS). Little is known about the relation of obesity and recurrent pregnancy loss. This study aimed to assess the relation and impact of obesity on the risk of recurrent pregnancy loss.

Patient and methods: A cross sectional study was done in gynecology and obstetrics department in Azadi Teaching Hospital during 1st May 2019-1st August 2019. We randomly selected 402 patients of reproductive age. Data was obtained through standardized questionnaire including obstetrical, medical and surgical history information. BMI was divided according to the the WHO criteria: underweight <19, normal (19±24.9), over weight (25-29.9), and obese women (BMI >30).

Results: The prevalence of recurrent miscarriage was 24 (6%); those who had 1-2 miscarriages was 165 (41%). Recurrent miscarriage was significantly common among those aged \geq 31 years 15 (9.9%), followed by 25-30 years 7(5.4%), and < 18 years 1 (3.4%). It was significantly common among obese 10 (12.7%), overweight 8 (4.5%) women, and underweight 1 (4.3). Recurrent pregnancy loss was not significantly more among those who had \geq 5 children 2 (12.5%) than those had 0-1 children14 (6.1%). Risk to have 1-2 miscarriages was significantly 0.3 times among those aged <18 years than those aged 19-30 years, and those who had 2-4 children 1.6 times than those had 0-1 children . Recurrent miscarriage was significantly 3.6 times more among obese women as compared with normal weight women, and 3.2 times among those aged \geq 31, as shown in Table 4.

Conclusion and recommendation: Obesity was significantly associated with increased risk of RPL. It is important to recommend females with recurrent pregnancy loss to decrease weight in order to get better results.

Key words: recurrent miscarriage, recurrent pregnancy loss, obesity, overweight, BMI

Introduction

The definition of recurrent miscarriage is the manifestation of three or above sequential miscarriages and is one of the great distressing untoward reproductive outcomes influencing about 1% of fertile couples [1, 2]. Fifty percent of cases of recurrent miscarriage are of unknown etiologies, adding to the associated distress. The other 50% of cases are due to a number of identifiable causes: chromosomal, uterine, coagulation, and immunological causes [2, 3]. Increased BMI (obesity), which has been associated with a number of adverse pregnancy outcomes, is of increasing prevalence [4,5]. Several studies revealed an association between obesity and miscarriage [6-8], which may be due to adverse influences on the embryo, the endometrium, or both [9]. From the assessment of evidence from several studies that discuss the relation between miscarriage and obesity, we found that obese females have a significant increased risk of miscarriage regardless of the conception method [10]. Unfortunately, still the influence of obesity on the risk of recurrent miscarriage, and the association between the various degrees of obesity and miscarriage rates has not so far been studied. The aim of this study is to assess the relation and impact of obesity on the risk of miscarriage.

Patients and Methods

Cross sectional study was done in gynecology and obstetrics department in Azadi Teaching Hospital during the period 1st May 2019-1st August 2019. Detailed information was taken from the patient through comprehensive questionnaire including information about obstetrical, medical and surgical history obtained from the patients.

Randomly selected 402 patients in reproductive age were interviewed in the obstetrics and gynecology department.

Recurrent pregnancy loss (RPL), is defined as 3 consecutive pregnancy losses prior to 20 weeks from the last menstrual period [11].

The body mass index (BMI) of all mothers was calculated using the formula (weight in kg/(height in meters)2.

BMI was divided according to the the WHO criteria: underweight <19, normal (19±24.9), over weight (25-29.9), and obese women (BMI >30).

Data entry and analysis was done using SPSS -25. The significance of differences between the groups was analyzed using Chi-square test; and odds ratio used to calculate the risk using logistic regression. p<0.05 was considered statistically significant.

Results

The prevalence of recurrent miscarriage was 24(6%), those whohad 1-2miscarriages was 165(41%), as shown in Figure 1.

Recurrent miscarriage was common among those aged \geq 31 years 15 (9.9%), followed by 25-30 years 7(5.4%), and < 18 years 1 (3.4%), This relation was statistically significant as shown in Table 1.

Recurrent miscarriage was common among those obese women 10(12.7%), followed by overweight 8(4.5%), and underweight1 (4.3%), respectively. This relation was statistically significant as shown in Table 2.

Recurrent miscarriage was commonly found among those who had \geq 5 children 2 (12.5%) than those had 0-1 children 14(6.1) This relation was statistically not significant as shown in Table 3.

Logistic regression of number of abortions shows that risk to have 1-2 miscarriages was significantly 0.3 times more among those aged <18 years than those aged 19-30 years, and those who had 2-4 children 1.6 times than those who had 0-1 child.

Recurrent miscarriage was significantly 3.6 times more among obese women as compared with normal weight women, and 3.2 times more among those aged \geq 31, as shown in Table 4.

Table 1: The distribution of patients according to number of miscarriages and age.

	0.000			
Age	0	1-2	>3	Total
<18 year	23	5	1	29
	79.3%	17.2%	3.4%	100.0%
18-24	56	36	1	93
	60.2%	38.7%	1.1%	100.0%
25-30	64	58	7	129
	49.6%	45.0%	5.4%	100.0%
≥31	70	66	15	151
	46.4%	43.7%	9.9%	100.0%
Total	213	165	24	402
	53.0%	41.0%	6.0%	100.0%

X²=19.188, df= 6, P value =0.0

Figure 1. The prevalence of abortion



BMI	0	1-2	>3	Total
19-24.9	61	55	5	121
	50.4%	45.5%	4.1%	100.0%
<19	10	12	1	23
	43.5%	52.2%	4.3%	100.0%
25-29.9	108	63	8	179
	60.3%	35.2%	4.5%	100.0%
>30	34	35	10	79
	43.0%	44.3%	12.7%	100.0%
Total	213	165	24	402
	53.0%	41.0%	6.0%	100.0%

Table 2: The distribution of patients according to number of miscarriages and BMI

X²=14.188, df=6, P value =0.028 Significant

П

Table 3:	The distribution of	of patients accordin	a to number o	of miscarriages and	d number of live births
			.g		

in and the second	Miscarriage			
No. of live births	0	1-2	>3	Total
0-1	131	83	14	228
	57.5%	36.4%	6.1%	100.0%
2-4	74	76	8	158
	46.8%	48.1%	5.1%	100.0%
≥5	8	6	2	16
	50.0%	37.5%	12.5%	100.0%
Total	213	165	24	402
	53.0%	41.0%	6.0%	100.0%

X²=6.5, df=4, P value =0.162 Non Significant

Variables	1_2		≥3			
	Sig.	OR	Sig.	OR		
	BMI					
19-24.9	S.	1		1		
<19	0.54	1.331	0.862	1.22		
25-29.9	0.075	0.647	0.864	0.9		
≥30	0.663	1.142	0.03	3.6		
14	Age					
<18 year	0.012	0.3	0.694	0.7		
19-30	2	1	0	1		
≥31	0.4	1.204	0.012	3.2		
No. of live births						
0-1		1		1		
2_4	0.025	1.6	0.980	1.012		
≥5	0.762	1.2	0.311	2.34		

Table 4: The logistic regression of different characteristics and number of miscarriages

Discussion

The prevalence of recurrent miscarriage was 24(6%), with significant association with OR (3.2) for those aged \geq 31. This was higher than found by Jauniaux E et al (2.3%) [12].

In Swedish women the incidence of recurrent miscarriage was (0.05%) in women aged 18-42 years and 650 per 100 000 (0.65%) in women who had achieved pregnancy [13].

Recurrent miscarriage was significantly increased with increasing age, mostly among those aged 31 years and more (9.9%). This goes with previous studies that found the risk of recurrent miscarriage increased with increasing age [14][15]. This may be due to fetal chromosomal abnormality [16,17].

Recurrent miscarriage was common among obese women 10 (12.7%), 3.6 times increased risk, than non-obese women compared with normal weight women. This was supported by previous studies that found obese women had higher percentage of recurrent miscarriage (59%), with OR 1.7 times more than normal and overweight patients [18].

Boots C E et al found similar results, (16.6%) of obese women had miscarriage, with risk of (1.3) times of having multiple pregnancy loss [19].

In 2004, Lashen et al found that obesity was associated with RPL with OR 3.5 of getting RPL [8].

In a meta-analysis of 16 studies, [10] researchers found significant relation between miscarriage and overweight and obesity, regardless of the method of conception (IVF or normal). The pooled OR of obese women to get RPL was (1.67, 95% CI 1.25–2.25).

In a retrospective study done in England, a significant increased risk of RPL among obese women was documented [20].

The explanation of the association between RPL and obesity may be due to the fact that many endocrine disorders are associated with obesity such as PCOS, diabetes and thyroid disease, all of which are risk factors of RPL [21].

The oocyte quality, embryo development, and endometrial receptivity, which are important in normal conception are controlled by the hypothalamic–pituitary– gonadal hormonal axis. The abnormality of this axis in obese women may be the cause of RPL [22].

In a meta-analysis study, the immunopathological pathways seem to be responsible for pregnancy loss among overweight and obese women. [23].

Many studies have linked the cause to the immunological factor depending on the fact that obesity is associated with high levels of C-reactive protein (CRP) and interleukin-6 (IL- 6) [24-26].

Leptin production stimulates the expression of matrix metalloproteinase by the cytotrophoblast. [27], and modulating the function of local T lymphocytes and protooncogenes [28].

Women with RPL had lower serum Leptin levels, and obesity is associated with leptin resistance and deficiency [29, 30].

Conclusion and recommendation

Obesity was significantly associated with increased risk of RPL, therefore it is important to assess the BMI of the patient with RPL and recommend them to decrease weight in order to get better results.

References

1. Bohlmann MK, Luedders DW, von Wolff M. Evidencebased guidelines for the investigation and medical treatment of recurrent miscarriage. Hum Reprod 2007;22:309; author reply 309–11.

2. Li TC, Makris M, Tomsu M, Tuckerman E, Laird S. Recurrent miscarriage: aetiology, management and prognosis. Hum Reprod Update 2002;8:463–81.

3. Cocksedge KA, Saravelos SH, Wang Q, Tuckerman E, Laird SM, Li TC. Does free androgen index predict subsequent pregnancy outcome in women with recurrent miscarriage? Hum Reprod 2008;23:797–802.

4. James PT. Obesity: the worldwide epidemic. Clin Dermatol 2004;22: 276–80.

5. Linne Y. Effects of obesity on women's reproduction and complications during pregnancy. Obes Rev 2004;5:137–43.

6. Hamilton-Fairley D, Kiddy D, Watson H, Paterson C, Franks S. Association of moderate obesity with a poor pregnancy outcome in women with polycystic ovary syndrome treated with low dose gonadotrophin. Br J Obstet Gynaecol 1992;99:128–31.

7. Bussen S, Sutterlin M, Steck T. Endocrine abnormalities during the follicular phase in women with recurrent spontaneous abortion. Hum Reprod 1999;14:18–20.

8. Lashen H, Fear K, Sturdee DW. Obesity is associated with increased risk of first trimester and recurrent miscarriage: matched case-control study. Hum Reprod 2004;19:1644–6.

9. Bellver J, Rossal LP, Bosch E, Zuniga A, C orona JT, Melendez F, et al. Obesity and the risk of spontaneous abortion after oocyte donation. Fertil Steril 2003;79:1136–40.

10. Metwally M, Ong KJ, Ledger WL, Li TC. Does high body mass index increase the risk of miscarriage after spontaneous and assisted conception? A meta-analysis of the evidence. Fertil Steril 2007.

11. Stephenson MD. Frequency of factors associated with habitual abortion in 197 couples. Fertil Steril. 1996;66:24–29.

12. Jauniaux E, Farquharson RG, Christiansen OB, Exalto N. Evidence-based guidelines for the investigation and medical treatment of recurrent miscarriage. Hum Reprod. 2006;21:2216–22.

13. Rasmark Roepke E1, Matthiesen L, Rylance R, Christiansen OB. Is the incidence of recurrent pregnancy loss increasing? A retrospective register-based study in Sweden. Acta Obstet Gynecol Scand. 2017 Nov;96(11):1365-1372.

14. Practice Committee of the American Society for Reproductive Medicine. Evaluation and treatment of recurrent pregnancy loss: a committee opinion. Fertil Steril. 2012;98(5):1103–1111.

15. Wilcox AJ, Weinberg CR, O'Connor JF, et al. Incidence of early loss of pregnancy. N Eng J Med. 1988;319(4):189–194.

16. Marquard K, Westphal LM, Milki AA, Lathi RB. Etiology of recurrent pregnancy loss in women over the age of 35 years. Fertil Steril. 2010 Sep;94(4):1473-7.].

17. Bianco, K., Caughey, A.B., Shaffer, B.L., Davis, R., and Norton, M.E. History of miscarriage and increased incidence of fetal aneuploidy in subsequent pregnancy. Obstet Gynecol. 2006; 107: 1098–1102]

18. Winnie Lo, Raj Rai, Aisha Hameed, Susan R. Brailsford, Al-GhamdiAhlam A. Regan L.The effect of body mass index on the outcome of pregnancy in women with recurrent miscarriage. Journal of Family and Community Medicine | December 2012 | Vol 19 | Issue 3 | 167-171.

19.B oots C E, Bernardi LA., and Stephenson M D. Frequency of euploid miscarriage is increased in obese women with recurrent early pregnancy loss. Fertility and Sterility 2014; 102 (2):455-9.

20. Metwally M, Saravelos SH, Ledger WL, Li TC. Body mass index and risk of miscarriage in women with recurrent miscarriage. Fertil Steril 2010;94: 290–5.

21. Weiss JL, Malone FD, Emig D, Ball RH, Nyberg DA, Comstock CH, et al. Obesity, obstetric complications and cesarean delivery rate—a populationbased screening study. Am J Obstet Gynecol 2004;190:1091–7.

22. Broughton DE, Moley KH. Obesity and female infertility: Potential mediators of obesity's impact. Fertil Steril 2017;107: 840–847.

23. Cavalcante M B., Sarno M, Peixoto Alberto B., Júnior E A and Barini R. Obesity and recurrent miscarriage: A systematic review and meta-analysis. J. Obstet. Gynaecol. Res. 2019.;45(1): 30–38.

24. Giannini DT, Kuschnir MCC, de Oliveira CL, Szklo M. Waist-to-height ratio as a predictor of C-reactive protein levels. J Am Coll Nutr 2017; 36: 624–630.

25. Klein S, Fontana L, Young VL et al. Absence of an effect of liposuction on insulin action and risk factors for coronary heart disease. N Engl J Med 2004; 350: 2549–2557.

26. Sindhu S, Thomas R, Shihab P, Sriraman D, Behbehani K, Ahmad R. Obesity is a positive modulator of IL-6R and IL-6 expression in the subcutaneous adipose tissue: Significance for metabolic inflammation. PLoS One 2015; 10: e0133494.

27.Sagawa N, Yura S, Itoh H, Kakui K, Takemura M, Nuamah MA, et al. Possible role of placental leptin in pregnancy: a review. Endocrine 2002;19:65–71.

28. Bajoria R, Sooranna SR, Ward BS, Chatterjee R. Prospective function of placental leptin at maternal-fetal interface. Placenta 2002;23:103–15.

29. Laird SM, Quinton ND, Anstie B, Li TC, Blakemore AI. Leptin and leptinbinding activity in women with recurrent miscarriage: correlation with pregnancy outcome. Hum Reprod 2001;16:2008–13.

30.Moschos S, Chan JL, Mantzoros CS. Leptin and reproduction: a review. Fertil Steril 2002;77:433–44.