


Research Article

Correlation of Anti Mullerian Hormone and Antral Follicular Count in Ovarian Reserve Testing

Anuradha K^{1*}, Partha Majumder S², Shiffin R³

Abstract

Background: Ovarian reserve defines a woman's reproductive ability and the number and quality of oocytes she possesses. It is a complex clinical state dependent on age, genetics, and environmental issues. It can reflect women's endocrine function and fertility which may gradually decrease with increasing age.

Methods: This study was a cross-sectional study conducted at the outpatient department of Obstetrics and Gynae, in Enam Medical College Hospital, Savar and a local private hospital at Savar. The study was conducted during the period of July 2019-December 2019. The sample size for this study was 120.

Result: The most respondent 44 (36.7%) were in between 35-40 years. The mean \pm SD of BMI was 26.61 ± 1.96 and followed by duration of infertility (years) was 3.75 ± 1.64 , total ovarian volume (ml) was 7.66 ± 1.32 . Tubal factor was found in 27 (21.7%) cases and followed by male factor was in 24 (20%), PCOS was in 20 (16.7%), endometriosis was in 6 (5%), unexplained infertility was in 22 (18.3%). In low group AFC (mean \pm SD) was 07.15 ± 4.82 where AMH (mean \pm SD) was 6.66 ± 5.34 and followed by normal was 09.38 ± 3.59 and 9.48 ± 3.91 and high was 15.45 ± 5.46 and 16.08 ± 5.23 . There was no significance correlation found in these two predictors.

Conclusion: AMH is considered as most reliable investigation for ovarian reserve testing. Serum AMH level has strong correlation with comparatively low cost Antral follicular count. Antral follicular count can be done in poor patients for ovarian reserve test.

Keywords: Anti Mullerian Hormone (AMH); Antral Follicular Count (AFC); Ovarian Reserve Testing

Introduction

Ovarian reserve defines a woman's reproductive ability and the number and quality of oocytes she possesses [1]. It is a complex clinical state dependent on age, genetics, and environmental issues [2]. It can reflect women's endocrine function and fertility which may gradually decrease with increasing age [3-5]. This decline is unavoidable but the rate of primordial follicles lose varies significantly along with variation on the onset of barrenness and time of menopausal transition [2]. Ovarian reserve tests (ORT) helps in distinguishing and treating infertility and in evaluating prior to in vitro fertilization [6]. ORT needs to be easy to perform and followed up and reliable for making decision [7]. There are two best ovarian reserve markers to forecast ovarian response to FSH are mean antral follicle count (AFC) and anti-Mullerian

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hormone (AMH) [8, 9]. Both these markers considered to be accurate in predicting response to control ovarian stimulation in the in-vitro fertilization setting and have higher reliable predictive value for poor ovarian response (POR) comparing to other indicators [10-14]. However, AMH and AFC may show varied results, especially in where AMH and AFC level could be at odds with each other [8, 15]. AMH is formed by the granulosa cells of pre-antral and small antral follicles, and the menstrual cycle don't affect its level or exogenous hormonal supplementation [16, 17]. Hence, AMH levels can better represent the number of primordial follicles and reflect ovarian reserve function. The AFC denotes to the number of follicles with diameters of 2 mm to 9 mm and these follicles tends to grow after enrollment in the luteal phase of the previous cycle and mostly reflect the number of follicles that will continue to mature at the time of ovulation treatment cycle [18]. Some studies had emphasis that AMH can reflect both the number of antral follicles and the quality of oocytes [19, 20]. Usually, it is thought that AMH is maintained throughout the menstrual cycle and it is stable as well [21-23]. Hence, AMH is measured to be the best indicator to assess ovarian reserve. The objective of this study was to find out the correlation between the anti-Mullerian hormone (AMH) and antral follicular count (AFC) in ovarian reserve testing.

Objective of the study

The objective of this study was to find out the correlation of anti-Mullerian hormone (AMH) and antral follicular count (AFC) in ovarian reserve testing.

Materials and Methodology

This study was a cross-sectional study conducted at the out-patient department of Obstetrics and Gynae, in Enam Medical College Hospital, Savar and a local private hospital at Savar. The study was conducted during the period of July 2019-December 2019. The sample size for this study was 120.

Inclusion criteria

- The adult patients who were aged more than 20 years were included in this study.
- The patients who came for infertility treatment.
- The patients having the normal sonographic texture of ovaries.
- Patients with no signs of hyper- androgenetic.
- The patients who were willing to give their consent after knowing the study purpose.

Exclusion criteria

- The patients who had the history of ovarian surgery, ovarian cyst, or endocrine disease.
- The patients who were not willing to give their consent after knowing the study purpose.

The AMH and AFC measurements were done on the second or third day of the menstrual cycle and this was done consistently. The clinical history of all the respondents was recorded with due consents from the hospital authority. Besides, all the respondents were given a consent from where they agreed to give their consent after knowing the study purpose. All participants were assured of high confidentiality. The baseline and demographic data of all the study patients was also recorded which further used in this study. For statistical analysis, the SPSS version 21 was used as the statistical tool.

Result

Figure 1, shows the age distribution of the respondents. A few of the respondents 10 (8.3%) were aged between 20-24 years and followed by 37 (30.8%) were 25-29 years, 29 (24.2%) were 30-34 years and the most 44 (36.7%) were 35-40 years. Table 1 represents the baseline characteristics of the respondents where the mean \pm SD of BMI was 26.61 ± 1.96 and followed by duration of infertility (years) was 3.75 ± 1.64 , total ovarian volume (ml) was 7.66 ± 1.32 , number of oocytes was 6.72 ± 3.68 and Number of embryos was 5.36 ± 2.40 . Figure 2 shows the etiological factors of the respondents where tubal factor was found in 27 (21.7%) cases and followed by male factor was in 24 (20%), PCOS was in 20 (16.7%), endometriosis was in 6 (5%), unexplained infertility was in 22 (18.3%) and more than 1 factor was in 22 (18.3%) cases. Table 2 represents the distribution of the study patients according to serum AMH level. Serum AMH

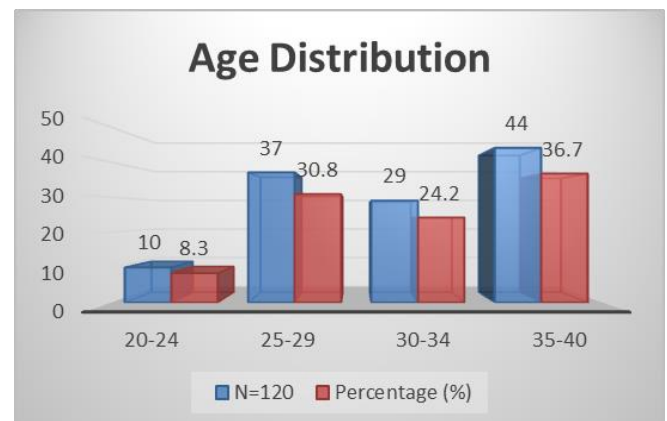


Figure 1: Age Distribution of the Respondents.

Baseline Characteristics	Mean \pm SD	P value
BMI	26.61 ± 1.96	0.724 (NS)
Duration of infertility, years	3.75 ± 1.64	0.672 (NS)
Total ovarian volume (ml)	7.66 ± 1.32	0.067 (NS)
Number of oocytes	6.72 ± 3.68	0.000 (HS)
Number of embryos	5.36 ± 2.40	0.000 (HS)

Table 1: Baseline Characteristics of the Respondents.

level <1.0 (Low) was seen in 2 (8.7%) cases of 25-29 years, 8 (27.6%) cases of 30-34 years and 21 (47.7%) cases of 35-40 years and followed by level 1.0-3.5 (Normal) was in 9 (90%), 35 (91.3%), 20 (69%) and 23 (52.3%) and level >3.5 (High) was seen in 1 (10%) and 1 (3.4%) of these age groups. The Mean ± SD of Serum AMH of these age groups was 2.67 ± 0.80, 2.24 ± 0.77, 1.57 ± 1.10, 1.17 ± 1.06 where the range

(min-max) was in between (1.50-3.50), (0.46-3.50), (0.18-3.30) and (0.02-3.48).

Table 3 shows Distribution of the study patients according to total AFC level. AFC level <5 (Low) was seen in 2 (8.7%) cases of 25-29 years, 2 (6.9%) cases of 30-34 years and 10 (22.7%) cases of 35-40 years and followed by level 5-15 (Normal) was in 10 (100%), 31 (83.8%), 27 (93.1%) and 33 (75%) and level >15 (High) was seen in 4 (10.8%) and 1 (2.3%). The Mean ± SD of AFC level of these age groups was 12.5 ± 1.7, 12.3 ± 3.2, 8.8 ± 3.1, 7.5 ± 3.3 where the range (min-max) was in between (11.0-15.0), (4.0-18.0), (4.0-14.0) and (4.0-16.0). Table 4 explains the comparison of average follicle number among AMH and AFC groups. In low group AFC (mean ± SD) was 07.15 ± 4.82 where AMH (mean ± SD) was 6.66 ± 5.34 and followed by normal was 09.38 ± 3.59 and 9.48 ± 3.91 and high was 15.45 ± 5.46 and 16.08 ± 5.23. There was no significance correlation found in these two predictors. Table 5 shows the cost of ovarian reserve testing. The Mullerian Hormone test costs Tk 7000 where Antral Follicular test around Tk 1500.

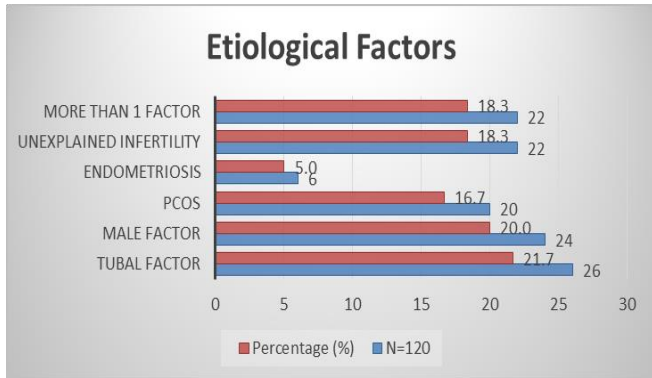


Figure 2: Etiological Factors.

Table 2: Distribution of the study patients according to Serum AMH Level.

Serum AMH (ng/ml)	20-24 years		25-29 years		30-34 years		35-40 years		P-value
	N=10	(%)	N=37	(%)	N=29	(%)	N=44	(%)	
<1.0 (Low)	0	0.0	2	8.7	8	27.6	21	47.7	
1.0-3.5 (Normal)	9	90.0	35	91.3	20	69.0	23	52.3	
>3.5 (High)	1	10.0	0	0	1	3.4	0	0.0	
Mean ±SD	2.67 ± 0.80		2.24 ± 0.77		1.57 ± 1.10		1.17 ± 1.06		0.001
Range (min-max)	(1.50-3.50)		(0.46-3.50)		(0.18-3.30)		(0.02-3.48)		

Table 3: Distribution of the study patients according to total AFC Level.

Total AFC (Number)	20-24 years		25-29 years		30-34 years		35-40 years		P-value
	N=10	(%)	N=37	(%)	N=29	(%)	N=44	(%)	
<5 (Low)	0	0	2	8.7	2	6.9	10	22.7	
5-15 (Normal)	10	100	31	83.8	27	93.1	33	75.0	
>15 (High)	0	0	4	10.8	0	0.0	1	2.3	
Mean ±SD	12.5 ± 1.7		12.3 ± 3.2		8.8 ± 3.1		7.5 ± 3.3		0.001
Range (min-max)	(11.0-15.0)		(4.0-18.0)		(4.0-14.0)		(4.0-16.0)		

Table 4: Comparison of average follicle number among AMH and AFC groups.

Sub group	AFC (mean ± SD)	AMH (mean ± SD)	p value
Low	07.15 ± 4.82	6.66 ± 5.34	0.54
Normal	09.38 ± 3.59	9.48 ± 3.91	0.99
High	15.45 ± 5.46	16.08 ± 5.23	0.76

Table 5: Cost of Ovarian Reserve Testing.

Name of Test	Cost in Tk
Mullerian Hormone	7000
Antral Follicular	1500

Discussion

A few of the respondents 8.3% were aged between 20-24 years and followed by 30.8% were 25-29 years, 24.2% were 30-34 years and the most 36.7% were 35-40 years [Figure 1]. Juthi Bhowmik et al. in their study showed a few of the respondents 8.1% were aged between 21-25 years and followed by 31.1% were 26-30 years, 24.3% were 30-35 years and the most 36.5% were 36-40 years [24]. The mean \pm SD of BMI was 25.61 ± 1.96 and followed by duration of infertility (years) was 3.65 ± 1.64 , total ovarian volume (ml) was 7.56 ± 1.32 , number of oocytes was 6.62 ± 3.68 and Number of embryos was 5.26 ± 2.40 . [table I] Shahinaz H. El-Shorbagy found the mean \pm SD of BMI was 26.61 ± 1.96 and followed by duration of infertility (years) was 3.75 ± 1.64 , total ovarian volume (ml) was 7.66 ± 1.32 , number of oocytes was 6.72 ± 3.68 and Number of embryos was 5.36 ± 2.40 [25]. Tubal factor was found in 21.7% cases and followed by male factor was in 20%, PCOS was in 16.7%, endometriosis was in 5%, unexplained infertility was in 18.3% and more than 1 factor was in 18.3% cases [Figure 2]. Shembekar CA et al in their study found the tubal factor was present in 22% cases and followed by male factor was in 20%, PCOS was in 17%, endometriosis was in 5%, unexplained infertility was in 18% and more than 1 factor was in 18% cases [26]. Serum AMH level <1.0 (Low) was seen in 8.7% cases of 25-29 years, 27.6% cases of 30-34 years and 47.7% cases of 35-40 years and followed by level 1.0-3.5 (Normal) was in 90%, 91.3%, 69% and 52.3% and level >3.5 (High) was seen in 10% and 3.4% of these age groups. The Mean \pm SD of serum AMH of these age groups was 2.67 ± 0.80 , 2.24 ± 0.77 , 1.57 ± 1.10 , 1.17 ± 1.06 where the range (min-max) was in between (1.50-3.50), (0.46-3.50), (0.18-3.30) and (0.02-3.48) [Table 2]. In a related study, the serum AMH level <1.0 (Low) was seen in 4.3% cases of 26-30 years, 27.8% cases of 31-35 years and 48.1% cases of 36-40 years and followed by level 1.0-3.5 (Normal) was in 100%, 95.7%, 72.2% and 51.9% and the Mean \pm SD of serum AMH of these age groups was 2.87 ± 0.80 , 2.44 ± 0.77 , 1.77 ± 1.10 , 1.37 ± 1.06 where the range (min-max) was in between (1.50-3.50), (0.46-3.50), (0.18-3.30) and (0.02-3.48).²⁴ AFC level <5 (Low) was seen in 8.7% cases of 25-29 years, 6.9% cases of 30-34 years and 22.7% cases of 35-40 years and followed by level 5-15 (Normal) was in 100%, 83.8%, 93.1% and 75% and level >15 (High) was seen in 10.8% and 2.3%. The Mean \pm SD of AFC level of these age groups was 12.5 ± 1.7 , 12.3 ± 3.2 , 8.8 ± 3.1 , 7.5 ± 3.3 where the range (min-max) was in between (11.0-15.0), (4.0-18.0), (4.0-14.0) and (4.0-16.0) [Table 3]. Juthi Bhowmik et al. in their study found the AFC level <5 (Low) was in 4.3% cases of 26-30 years, 5.6% cases of 31-35 years and 22.2% cases of 36-40 years and followed by level 5-15 (Normal) was in 100%, 82.6%, 94.4% and 74.1% and level >15 (High) was seen in 13% and 3.7%. The Mean \pm SD

of AFC level of these age groups was 13.5 ± 1.7 , 13.3 ± 3.2 , 9.8 ± 3.1 , 8.5 ± 3.3 where the range (min-max) was in between (11.0-15.0), (4.0-18.0), (4.0-14.0) and (4.0-16.0) [24]. In low group AFC (mean \pm SD) was 07.15 ± 4.82 where AMH (mean \pm SD) was 6.66 ± 5.34 and followed by normal was 09.38 ± 3.59 and 9.48 ± 3.91 and high was 15.45 ± 5.46 and 16.08 ± 5.23 . There was no significance correlation found in these two predictors [Table 4]. In the study of Parvathy T et al, the low group AFC (mean \pm SD) was seen 07.25 ± 4.82 where AMH (mean \pm SD) was 6.76 ± 5.34 and followed by normal was 09.48 ± 3.59 and 9.58 ± 3.91 and high was 15.55 ± 5.46 and 16.08 ± 5.23 . There was no significance correlation found in these two predictors of ovarian reserve [27]. The Mullerian Hormone test costs Tk 7000 where Antral Follicular test around Tk 1500 [Table 5].

Conclusion

Although there is a lack of data to accomplish which of the two markers served better to predict ovarian reserve, but most of the studies claimed for the two best ovarian reserve markers to forecast ovarian response to FSH are mean antral follicle count (AFC) and anti-Mullerian hormone (AMH). Although, AMH is considered to be more effective in predicting the ovarian response but many authors thought that AMH and AFC are having the same level of accuracy and clinical value in prediction of ovarian response. So, those authors emphasis that AFC can be considered as a substitute of expensive AMH estimation in predicting the ovarian response. However, better understanding of patients AMH and AFC level, the physician can make a better treatment plan which will bring better treatment outcome.

References

1. Practice Committee of the American Society for Reproductive Medicine. Testing and interpreting measures of ovarian reserve: A committee opinion. Fertil Steril 103 (2015): 9-17.
2. Tal R, Seifer DB. Ovarian reserve testing: A user's guide. American Journal of Obstetrics Gynaecology 217 (2017): 129-140.
3. Fábregues F, Iraola A, Casals G, et al. Evaluation of Two Doses of Recombinant Human Luteinizing Hormone Supplementation in Down-Regulated Women of Advanced Reproductive Age Undergoing Follicular Stimulation for IVF: A Randomized Clinical Study. Eur J Obstet Gynecol Reprod Biol 158 (2011): 56-61.
4. Azhar E, Seifer DB, Melzer K, et al. Knowledge of Ovarian Reserve and Reproductive Choices. J Assist Reprod Genet 32 (2015): 409-415.
5. Kirshenbaum M, Orvieto R. Premature Ovarian Insufficiency (POI) and Autoimmunity-An Update Appraisal. J Assist Reprod Genet 36 (2019): 2207-2215.

6. Scott RT, Hofmann GE. Prognostic assessment of ovarian reserve. *Fertil Steril* 63 (1995): 1-11.
7. Kwee J, Elting ME, Schats R, et al. Ovarian volume and Antral follicle count for the prediction of low and hyper-responders with in vitro fertilization. *Reprod Biol Endocrinol* 5 (2007): 9.
8. Zhang Y, Xu Y, Xue Q, et al. Discordance between antral follicle counts and anti- Müllerian hormone levels in women undergoing in vitro fertilization. *Reprod Biol Endocrinol* 17 (2019): 51.
9. Klenov V, Jungheim E. AntiMullerian hormone (AMH) is a better predictor of response to controlled ovarian hyperstimulation (COH) than antral follicle count (AFC) in women with discordant markers of ovarian reserve. *Fertil Steril* 106 (2016): e123-e124.
10. Depmann M, Broer SL, van der Schouw YT, et al. Can we predict age at natural menopause using ovarian reserve tests or mother's age at menopause? A systematic literature review. *Menopause* 23 (2016): 224-232.
11. Medicine PCotASfR. Testing and interpreting measures of ovarian reserve: a committee opinion. *Fertil Steril* 103 (2015): e9-e17
12. Broer SL, Dolleman M, Opmeer BC, et al. AMH and AFC as predictors of excessive response in controlled ovarian hyperstimulation: a meta-analysis. *Hum Reprod Update* 17 (2011): 46-54.
13. Vural B, Cakiroglu Y, Vural F, et al. Hormonal and functional biomarkers in ovarian response. *Arch Gynecol Obstet* 289 (2014): 1355-1361.
14. Broer SL, Mol BW, Hendriks D, et al. The role of anti-müllerian hormone in prediction of outcome after IVF: comparison with the antral follicle count. *Fertil Steril* 91 (2009): 705-714.
15. Tal R, Seifer DB. Ovarian reserve testing: a user's guide. *Am J Obstet Gynecol* 217 (2017): 129-140.
16. Cook CL, Siow Y, Taylor S, et al. Serum müllerian -inhibiting substance levels during normal menstrual cycles. *Fertil Steril* 73 (2000): 859-861.
17. Kelsey TW, Wright P, Nelson SM, et al. A validated model of serum anti-müllerian hormone from conception to menopause. *PLoS One* 6 (2011): e22024.
18. Rajpert-De Meyts E, Jorgensen N, Graem N, et al. Expression of anti-mullerian hormone during normal and pathological gonadal development: association with differentiation of Sertoli and granulosa cells. *J Clin Endocrinol Metab* 84 (1999): 3836- 3844.
19. Iliodromiti S, Kelsey TW, Wu O, et al. The predictive accuracy of anti-Müllerian hormone for live birth after assisted conception: a systematic review and meta-analysis of the literature. *Hum Reprod Update* 20 (2014): 560-570.
20. Pilsgaard F, Grynnerup AG, Løssl K, et al. The use of antiMullerian hormone for controlled ovarian stimulation in assisted reproductive technology, fertility assessment and -counseling. *Acta Obstet Gynecol Scand* 97 (2018): 1105-1113.
21. La Marca A, Stabile G, Artenisio AC, et al. Serum anti-Mullerian hormone throughout the human menstrual cycle. *Hum Reprod* 21 (2006): 3103-3107.
22. Massarotti C, La Pica V, Sozzi F, et al. Influence of age on response to controlled ovarian stimulation in women with low levels of serum anti-Müllerian hormone. *Gynecol Endocrinol* 9 (2020): 1-5.
23. Younis JS, Iskander R, Fauser BCJM, et al. Does an association exist between menstrual cycle length within the normal range and ovarian reserve biomarkers during the reproductive years? A systematic review and metaanalysis. *Hum Reprod Update* 26 (2020): 904-928.
24. Juthi Bhowmik, Parveen Fatima, Jesmine Banu, et al. Correlation Between Follicle Stimulating Hormone, Anti-Müllerian Hormone and Antral Follicle Count with Different Age Groups in Infertile Women, Chattogram Maa-O-Shishu Hospital Medical College Journal (2021).
25. El-Shorbagy SH. Comparison of the Predictive Value of Antral Follicle Count, Anti-Müllerian Hormone and Follicle- Stimulating Hormone in Women Following GnRH-Antagonist Protocol for Intracytoplasmic Sperm Injection. *Open Journal of Obstetrics and Gynecology* 7 (2017): 432-446.
26. Shembekar CA, Upadhye JJ, Shembekar MC, et al. Anti-Mullerian hormone (AMH) as predictor of ovarian reserve. *Int J Reprod Contracept Obstet Gynecol* 6 (2017): 4006- 4010.
27. Parvathy T, Fessy Louis T, Ramesh P, et al. Ante Mullerian Hormone versus Antral Follicle Count as a predictor of ovarian response to controlled ovarian hyper stimulation in Assisted Reproductive Technique- A prospective study (2009).