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**Original Research Article** 

# An Evaluation of the Relationship between Hyperprolactinemia and Abnormalities in Seminal Fluid Analysis in Male Partners of Infertile Couples Undergoing Infertility Treatment in Southern Nigeria

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#### Abstract

Background: Male infertility is a significant yet under-reported public health concern. It affects 30% of couples who regularly engage in unprotected sexual activity. In Nigeria, the male partner is sometimes unwilling to have a fertility evaluation, which results in the underreporting of male infertility. The female partner is frequently held responsible for infertility. Hyperprolacting has been identified as a reversible cause of male infertility, which is amenable to treatment, but studies to evaluate the concept are sparse. Aim: To determine if there is an association between high prolactin levels and abnormal seminal fluid analysis parameters. *Methods*: The study is a cross-sectional analytical study involving male partners of infertile couples. Two hundred and thirty-three (233) patients that met the inclusion criteria and gave informed consent were enrolled on the study at the Human Reproduction and Research Program Unit (HRRP) of the University of Benin Teaching Hospital, Benin-city, Nigeria. A detailed history was taken, and a physical examination was done for all subjects. Seminal fluid analysis was done for all subjects, and blood was also collected for serum prolactin assay. Information was obtained from sociodemographic data and medical history. Data obtained using interviewer-administered questionnaires were analyzed with SPSS Package version 20.0. Results: The study included 233 patients, with a mean age of 40.77.1 and a majority (57.7%) of the subjects with aberrant SFA between the ages of 40 and 49. Participants with normal and abnormal SFA had hyperprolactinemia at 4.3% and 14.2%, respectively. There was a significant negative correlation between prolactin level, motility (r=-0.010, p=0.001), morphology (r=-0.077, p=0.001) and sperm count (r=-0.082, p=0.003). Obesity, alcohol consumption and smoking were significant predictors of abnormal sperm parameters (p=0.011, p=0.001, and p=0.001, respectively). Conclusion: This study indicated a relationship between hyperprolactinaemia, sperm count, motility, and morphology, which suggests that increased prolactin may negatively affect semen quality if left untreated.

Keywords: Hyperprolactinaemia, seminal fluid analysis, male infidelity.

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# **INTRODUCTION**

Infertility is a global public health and social problem, especially in developing countries like Nigeria, where childbirth is highly valued (Osaikhuwuomwan James and Osemwenkha Abieyuwa 2015). Around 17.5% of the adult population, approximately 1 in 6 worldwide, will experience infertility at some stage(Organization 2023). A report from Africa reveals that infertility is a significant issue in reproductive health and a frequent cause of gynaecological presentation(Madziyire, Magwali *et al.*, 2021). Due to socio-cultural beliefs, it is common in Nigeria to blame women for infertility, even though there is scientific data to suggest that in 40 - 50% of infertile couples, the male is the only factor contributing to infertility (Kumar and Singh 2015, Uadia and Emokpae 2015). In Sub-Saharan Africa, little has been done to address the causes of male infertility, even though research on the causes of infertility in females has been extensively conducted (Uadia and Emokpae 2015, Sharma 2017).

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The causes of infertility could either be male, female or combined. Male infertility becomes important when identifiable female causes of infertility are excluded, especially when there is abnormal semen quantity and quality based on World Health Organization criteria 2021(Organization 2023). However, no associated risk factors are identifiable in about 40-50% of males (Oladosu, Biliaminu *et al.*, 2017, Onyebuchi, Ifeoma *et al.*, 2018, Juneja, Phukan *et al.*, 2019). This is worse in third-world countries with a paucity of facilities for detailed investigations (Kaiser 2012, Juneja, Phukan *et al.*, 2019).

The capability of a man to impregnate a woman depends on his ability to produce and ejaculate semen with optimal parameters, and this depends on the proper functioning of the hypothalamic-pituitarygonadal axis, just as is the case in females. Any hormonal imbalance can potentially impact spermatogenesis and gonad genesis in general (Tritos and Klibanski 2019, Babakhanzadeh, Nazari et al., 2020). Elevated prolactin levels affect folliculogenesis by altering the normal pulsatile release of gonadotrophin-releasing hormone from the hypothalamus, consequently resulting in anovulatory infertility in the female population (Balen, Morley et al., 2016).

The influence of elevated prolactin on the hypothalamic-pituitary-gonadal axis and, by extension, abnormal sperm parameters is the subject of debate in the literature (Shahroona, Fatima *et al.*, 2007, Balen, Morley *et al.*, 2016, Kutlešić, Popović *et al.*, 2016, Ahmed and Ahmed 2017, Samperi, Lithgow *et al.*, 2019).

Prolactin is thought to work by suppressing the production gonadotropin-releasing pulsatile of hormone, which lowers follicle-stimulating, luteinizing, and testosterone (Hasan and Wijesinghe 2016, O'Leary 2020). Additionally, it may halt spermatogenesis and affect the quality of the semen (Anawalt 2013). Prolactin is synthesized in the anterior lobe of the hypophyseal gland. It is a hormone that affects sexual and metabolic functions and stimulates testicular function in men (Anawalt 2013, Hasan and Wijesinghe 2016). Less than 20ng/ml of serum prolactin is normal (Wong, Eloy et al., 2015). A myriad of factors other than disease conditions can affect serum prolactin levels (Benjamin, Akhere et al., 2014, Wong, Eloy et al., 2015). Men's prolactin levels are affected by stress, coitus, the use of oestrogens, progesterone, and androgens, as well as some medications like risperidone, metoclopramide, phenothiazines, and thyrotrophin-releasing hormones (Anawalt 2013, Balen, Morley et al., 2016, Tritos and Klibanski 2019). L-dopa and bromocriptine can lower prolactin levels (Wong, Eloy et al., 2015, Samperi, Lithgow et al., 2019).

Emerging evidence (Benjamin, Akhere *et al.*, 2014, Ajah, Ozumba *et al.*, 2016) revealed that 35% of male factors infertility do not have any identifiable cause at presentation. It is apt to investigate the cause of infertility, including abnormal parameters, on a semen analysis (Benjamin, Akhere *et al.*, 2014, Ajah, Ozumba *et al.*, 2016, Abdulhadi, Kawo *et al.*, 2021). Apart from patients with varicocele, a potential cause of infertility in Nigeria, most patients lack a definite cause, with poor semen quality accounting for 20–48% (Abdulhadi, Kawo *et al.*, 2021).

Current management aims to increase the couple's fertility potential. One of the ways to achieve this in males is to identify obvious risk factors and some less apparent links but whose treatment may be advantageous, like hyperprolactinemia (Ajah, Ozumba *et al.*, 2016, Abdulhadi, Kawo *et al.*, 2021). Therefore, it becomes imperative to determine the association between hyperprolactinaemia and seminal fluid analysis abnormalities of male partners of infertile couples attending the infertility clinic at the University of Benin Teaching Hospital, Benin City.

The study was carried out to determine the association between high prolactin levels (hyperprolactinaemia) and abnormal seminal fluid analysis parameters among males attending the infertility clinic at the University of Benin Teaching Hospital.

## **MATERIALS AND METHODS**

The prospective cross-sectional analytical study was conducted at the Human Reproduction and Research Programme (HRRP) unit of the Obstetrics and Gynaecology department of the University of Benin Teaching Hospital, Benin City, amongst male partners of infertile couples attending the clinic. The study was conducted from Oct 1 2021, to Mar 31 2023. The participants were voluntarily enrolled after counselling on; the nature of the study, the amount of blood/semen to be used, the benefits of the study, maintaining anonymity and their decision or not to participate in the study would not affect their care. The interview was conducted privately in the serene environment of HRRP. The social classification was according to the revised scheme by Ibadin and Akpede (Ibadin and Akpede 2021). All consenting infertile males were recruited until the sample size was completed. Data was obtained with an interviewer-administered structured questionnaire. Data was entered using Statistical Package for scientific solution (SPSS) Statistical Software Version 20. A statistical significance test was done at p < 0.05, while the confidence interval was constructed at a 95% confidence limit. The result of the analysis was presented as frequency distribution tables and figures.

# RESULTS

There were 58 (24.9%) participants with normal seminal fluid analysis (SFA), 175 (75.1%) had abnormal SFA and 43 (18.5%) cases of hyperprolactinaemia among the study population.

The mean age of participants in the study was  $40.7 \pm 7.1$  years. The difference between the mean ages of the participants with normal and abnormal SFA (38.8  $\pm$  6.9 vs 42.5  $\pm$  8.2 years, p-value= 0.305) was not statistically significant. The other socio-demographic characteristic, such as level of education, social class and duration of infertility among the participants with normal and abnormal SFA, were similar with no statistically significant difference—table 1a.

		phic Characteristics			
	Study Population		Abnormal SFA n=175	χ² Test	P –value
	N = 233 (%)	(%)			
Age Group (Years)				1	1
30-39	92 (39.5)	32 (55.2)	60 (34.3)		
40-49	120 (51.5)	19 (32.8)	101 (57.7)		
50-59	14 (6.0)	5 ( 8.6)	9 ( 5.1)	1.642	0.801
60-69	5 (2.1)	2 ( 3.4)	3 ( 1.7)		
≥70	2 (0.9)	0 ( 0.0)	2(1.1)		
Mean ± SD	40.7±7.1	38.8±6.9	42.5 ±8.2	1.033	0.305
Educational Status					
No formal education	0 (0.0)	0(0.0)	0(0.0)		
Completed Primary	21 (9.0)	6(10.3)	15(8.6)	0.020	0.00
Completed Secondary	55 (23.6)	9(15.5)	46(26.3)	0.832	0.660
Tertiary	157 (67.4)	43(74.1)	114(65.1)		
Social class					
Ι	116 (49.8)	34(58.6)	82(46.8)		
II	39 (16.7)	4(6.9)	35(20.0)		
III	51 (21.9)	11(18.9)	40(22.9)	7.655	0.105
IV	11 (4.7)	6(10.3)	5(2.9)		
V	16 (6.9)	3(5.2)	13(7.4)		
<b>Duration of Infertility</b>	(Yrs)				
1-5	97(41.6)	17(29.3)	80(45.7)		
6-10	105 (45.1)	35(60.3)	70(40.0)	2.007	0.502
11 – 15	27 (11.6)	4(6.9)	23(13.1)	2.007	0.503
16 - 20	4 (1.7)	2(3.4)	2(1.1)	1	

Table 1 a. Dama survey his d	Characteristics of Study Population
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	Table 1b: Clinical Cha Study Population	Study Population Normal SFA n= 58 Abnormal SFA $\chi^2$ P –						
	N = 233 (%)	(%)	n=175(%)	Test	value			
Smoking status	1							
Yes	44 (18.9)	7(12.1)	37(21.1)	13.890	0.001			
No	189 (81.1)	51(87.9)	138(78.9)					
Alcohol								
Yes	85 (36.5)	12(20.7)	73(41.7)	40.258	0.001			
No	148 (63.5)	46(79.3)	102(58.3)					
Body Mass Index	•							
Underweight	0 (0.0)	0(0.0)	0(0.0)	6.420 0.011				
Normal	67 (28.8)	22(37.9)	45(25.7)					
Overweight	109 (46.8)	31(53.4)	78(44.6)	_				
Obese	57 (24.4)	5(8.6)	52(29.7)					
Previous pelvic or groin Surg	gery							
Yes	9 (3.9)	0(0.0)	9(5.1)	0.036 0.850				
No	224 (96.1)	58(100.0)	166(94.9)					
Previous history of Varicoco	ele		•					
Yes	3 (1.3)	0(0.0)	3(1.7)	1.007 0	.316			
No	230 (98.7)	58(100.0)	172(98.3)					
<b>Previous Infections like uret</b>	hritis	•						
Yes	2 (0.9)	0(0.0)	2(1.1)	0.669 0	.414			
No	231 (99.1)	58(100.0)	173(98.9)	-				
Chronic medical illness								
Yes	27 (11.6)	5(8.6)	22(12.6)	0.052 0	.820			
No	206 (88.4)	53(91.4)	153(87.4)					

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When compared to persons with normal SFA, a substantially more significant percentage of participants who smoked cigarettes, drank alcohol and were obese had aberrant SFA parameters (21.1%, 41.7%, and 29.7% vs 12.1%, 20.7%, and 8.6%, with p-values of 0.001, 0.001, and 0.011, respectively).

Other clinical traits, such as prior groin surgery, previous pelvic infection, history of varicocele, and prior chronic medical illness, were comparable and did not statistically differ between people with normal and abnormal SFA (Figure 1b).

Table 2: Association between Protoctin levels and seminal fluid analysis abnormalStudy populationAbnormal SFANormal SFA $\chi^2$ TestN=233n = 175, N (%)n = 58, N (%)					
Prolactin level					
Hyperprolactinaemia	43 (18.5)	33 (18.9)	10 (17.2)	9.053	0.003
Normal Prolactin	190 (81.5)	142 (81.1)	48 (82.8)		

There were 43 cases of hyperprolactinemia, with a prevalence rate of 18.5 per cent overall. Hyperprolactinaemia was more in men with abnormal SFA—14.2% (33/233) versus 4.3% (10/233) in those with normal SFA. Compared to 10 (17.2%) patients with normal SFA and hyperprolactinaemia, 33 (18.9%) participants had abnormal SFA and

hyperprolactinaemia. The proportional difference was statistically significant at a p-value of 0.003 (Table 2).

In comparison to people with normal prolactin levels, a substantially more significant percentage of those with hyperprolactinaemia had abnormal sperm count (93% vs. 71%; p-value = 0.003), sperm morphology (90.7 vs. 60.5; p-value = 0.001), and sperm motility (93% vs. 58.9%; p-value = 0.001).

Table 3: Relationship between	prolactin level	and sperm parame	ters
Prolactin groups Number (Per	Unadjusted	95% CI for	
Hyperprolactinaemia	Normal	Odds ratio	Odds ratio

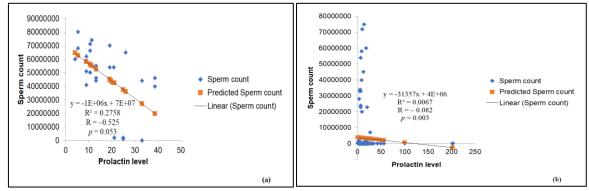
Variable	Prolactin groups Number (Percentage)		Unadjusted	95% CI for the	<i>p</i> -
	Hyperprolactinaemia	Normal	Odds ratio	Odds ratio	value
Sperm count					
Abnormal	40 (93.0)	135 (71.1)	0.184	0.055 - 0.620	0.003
Normal	3 (7.0)	55 (28.9)			
Sperm morphology					
Abnormal	39 (90.7)	115 (60.5)			
Normal	4 (9.3)	75 (39.5)	0.157	0.054 - 0.458	0.001
Sperm motility					
Abnormal	40 (93.0)	112 (58.9)	0.108	0.032 - 0.361	0.001
Normal	3 (7.0)	78 (41.1)			

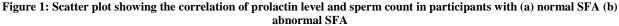
Participants with hyperprolactinaemia were likelier to have abnormal sperm count, morphology, and motility by an odds ratio of 0.184, 0.157, and 0.108, respectively, and statistically significant (Table 3).

Prolactin level, sperm count, sperm motility, and sperm morphology were all found to have non-significant negative correlations in individuals with normal SFA parameters (r=-0.525, p=0.053, -0.009, and

-0.209, respectively). Conversely, a significant negative correlation was seen between prolactin level and sperm count (r = -0.082, p = 0.003), sperm motility (r = -0.010, p=0.001), and sperm morphology (r = -0.077, p=0.001) in contrast, among individuals with abnormal SFA.

#### Table 4 and Figures 1 to 3





SFA			Prolactin level	Sperm count	Sperm motility	Sperm morphology
Normal SFA	Prolactin level	r	1	525*	.009	209
		p		.003	.964	.277
		Ń	58	58	58	58
	Sperm count	r	525*	1	.029	.270
		p	.053		.881	.157
		Ń	58	58	58	58
	Sperm motility	r	009	.029	1	303
		p	.964	.881		.111
		Ň	58	58	58	58
	Sperm	r	209	.270	303	1
	morphology	p	.100	.157	.111	
		Ń	58	58	58	58
Abnormal	Prolactin level	r	1	082	.010	077
SFA		p		.244	.884	.276
		Ń	175	175	175	175
	Sperm count	r	082	1	.183*	.008
		p	.003		.009	.911
		Ń	175	175	175	175
	Sperm motility	r	010	.183*	1	.071
		p	.001	.884		.313
		Ń	175	175	175	175
	Sperm	r	077	.008	.071	1
	morphology	p	.001	.911	.313	
		Ń	175	175	175	175

# Table 4: Correlation matrix of prolactin level vs SFA parameters in participants with normal SFA and those with abnormal SFA

r = Correlation coefficient \* p < 0.05 p = probability value N = Frequency

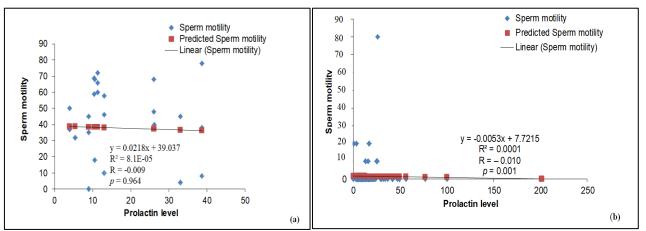


Figure 2: Scatter plot showing the correlation of prolactin level and sperm motility in participants with (a) normal SFA and (b) abnormal SFA

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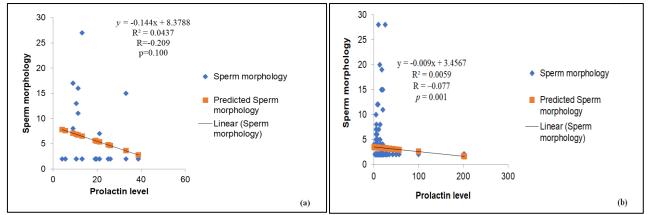


Figure 3: Scatter plot showing the correlation of prolactin level and sperm morphology in participants with (a) normal SFA and (b) abnormal SFA

 Table 5: Binary logistic regression analysis of factors associated with abnormal sperm count among the study population

Variable	B-regression Coefficient	Odds ratio	95% CI for the Odds ratio	p-value
Prolactin levels		o dub rutio		p fuide
Hyperprolactinaemia	-1.692	0.184	0.055 - 0.620	0.006
Normal prolactin				
Obesity	l .			
Yes	1.070	2.914	1.239 - 6.855	0.014
No*				
Previous groin surge	ry	•	•	•
Yes	0.154	1.167	0.235 - 5.780	0.850
No*				
Age >50years	•	•		•
Yes	-0.099	0.905	0.277 - 2.961	0.870
No <sup>*</sup>				
Alcohol				
Yes	3.963	52.615	7.126 - 388.467	0.001
No <sup>*</sup>				
Infections like urethi				
Yes	20.110	5416.047	0.001	0.999
No <sup>*</sup>				
Smoking				_
Yes	2.301	9.984	2.342 - 42.567	0.002
No <sup>*</sup>				
Chronic medical illne				
Yes	0.112	1.118	0.426 - 2.935	0.820
No <sup>*</sup>				
Previous varicocele	Γ	1		1
Yes	20.116	5447.536	0.001	0.999
No <sup>*</sup>		1*2		

CI= Confidence interval <sup>\*</sup>Reference category

Binary logistics regression analysis of factors associated with abnormal sperm morphology showed that hyperprolactinaemia, obesity, alcohol consumption and smoking were significant predictors of abnormal sperm morphology (p-value=0.001, 0.001, 0.001, 0.001) and were more likely to cause abnormality in sperm morphology by odds of 0.157, 0.284, 28.852 and 16.398 respectively. Furthermore, there was a non-significant trend for other predictors of abnormal sperm morphology (previous groin surgery, pelvic infection, age, varicocele and chronic medical illness) (Table 5).

Table 6:Binary logistic regression analysis of factors associated with abnormal sperm morphology among the study population

	study	population		
	<b>B-regression Coefficient</b>	Odd ratio	95% CI for the Odds ratio	<i>p</i> -value
Prolactin levels	•	•	•	
Hyperprolactinaemia	-1.850	0.157	0.054 -0.458	0.001
Normal Prolactin				
Obesity	•		•	
Yes	-1.259	0.284	0.131 - 0.615	0.001
No <sup>*</sup>				
Previous groin surge	ry	•	•	
Yes	0.606	1.833	0.372 - 9.040	0.457
No*				
Age > 50 years	•	•	•	
Yes	0.366	1.442	0.444 - 4.684	0.542
No <sup>*</sup>				
Alcohol	·		·	
Yes	8.362	28.852	8.722 –95.442	0.001
No <sup>*</sup>				
Infections like urethi	itis		·	
Yes	20.548	8396.23	0.001	0.999
No <sup>*</sup>				
Smoking	·		·	
Yes	2.797	16.398	3.864 - 69.599	0.001
No <sup>*</sup>				
Chronic medical illne	ess (Htn, DM)	•	•	
Yes	0.370	1.448	0.581 - 3.606	0.427
No <sup>*</sup>				
Previous varicocele		•	•	•
Yes	20.555	8451.829	0.001	0.999
No <sup>*</sup>				
	CI: Confi	dence interval	•	

<sup>\*</sup>Reference category

Binary logistics regression analysis of factors associated with abnormal sperm count revealed that hyperprolactinaemia, obesity, alcohol consumption and smoking were significant predictors of abnormal sperm count (p- value=0.006, 0.014, 0.001 and 0.002) and were more likely by odds of 0.184, 2.914, 52.615, and 9.984 respectively to lead to an abnormality in sperm count.

Other predictors of inadequate sperm count (past groin surgery, prior pelvic infection, age, prior varicocele, and chronic disease) also showed a non-significant trend) (Table 5).

Table 7: Binary logistic regression analysis of factors associated with abnormal sperm motility among the study
population

population						
	<b>B-regression</b> Coefficient	Odd ratio	95% CI for the Odds ratio	<i>p</i> -value		
Prolactin levels						
Hyperprolactinaemia	-2.228	0.108	0.320 - 0.361	0.001		
Normal Prolactin						
Obesity						
Yes	-1.306	0.271	0.125 - 0.587	0.001		
No <sup>*</sup>						
Previous groin surge	ry					
Yes	0.645	1.907	0.387 - 9.400	0.428		
No <sup>*</sup>						
Age > 50 years						
Yes	0.068	1.070	0.353 - 3.245	0.904		
No <sup>*</sup>						
Alcohol						
Yes	3.861	47.514	11.266 - 200.385	0.001		

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No <sup>*</sup>				
Infections like ure	thritis			-
Yes	20.587	8723.548	0.0001	0.999
No <sup>*</sup>				
Smoking				
Yes	2.842	17.142	4.040 - 72.739	0.001
No <sup>*</sup>				
Chronic medical i	llness (Htn, DM)			
Yes	0.412	1.510	0.607 - 3.759	0.376
No <sup>*</sup>				
Previous varicocel	e			
Yes	20.593	8782.119	0.0001	0.999
No <sup>*</sup>				

CI: Confidence interval <sup>\*</sup>Reference category

# DISCUSSION

Participants' average age was 40.7 years, similar to the studies' average ages of 40.3(Benjamin, Akhere et al., 2014), 40.5(Onyebuchi, Ifeoma et al., 2018), and 41.0(Omokanye, Olatinwo et al., 2016) years reported by other studies. Other researchers reported higher than 37.0 years(Ahmed and Ahmed 2017) and 38.6 years(Oladosu, Biliaminu et al., 2017). This observed discrepancy might be due to the different study settings and designs in these investigations. The higher mean age may be due to delayed presentation at the HRRP because it is a specialized infertility clinic/referral centre. Some clients have had initial consultations at other general gynaecological clinics before coming to the HRRP. Furthermore, most men do not readily admit their role in infertility in our culture, which always leads to the late presentation (Osaikhuwuomwan James and Osemwenkha Abieyuwa 2015).

The prevalence of hyperprolactinaemia in participants with normal seminal fluid analysis at 4.3% in this study was comparable to findings from other studies(Oladosu, Biliaminu et al., 2017), higher than the (2.1%) prevalence reported from Chicago (Ambulkar, Darves-Bornoz et al., 2022) However, it is lower in frequencies of 5.1%, 9.4%, and 16.7%, had been reported by Ozoemena et al., (Ozoemena, Ezugworie et al., 2011)., Hasan and Wijesingle (Hasan and Wijesinghe 2016), and Green and Amadi (Green and Amadi 2020). The observed difference may be due to regional disparity, smaller sample size and subject selection criteria used. Furthermore, this study used World Health Organisation 2021 criteria for seminal fluid analysis instead of the other studies that employed the older version. Evidence has shown that the prevalence of hyperprolactinaemia varies across the globe (Emokpae, Uadia et al., 2007, Ozoemena, Ezugworie *et al.*, 2011, Organization 2023)

In contrast, research from Sudan (Ahmed and Ahmed 2017) and Japan (Okada, Iwamoto *et al.*,1996) found a robust negative correlation between abnormal sperm parameters and hyperprolactinaemia. The study conducted in Sudan revealed a negative clinical correlation, which was insignificant. But it demonstrates that there might be an association between high sperm counts and elevated serum prolactin levels. Also, the study in Japan employed World Health Organization 1987 standards, which had a lower limit of 20 x  $10^6$ /ml, as opposed to our study, which used World Health Organisation 2021 criteria, which had a lower limit of 16 x  $10^6$ /ml.

Similarly, Dagistani (Al-Daghistani and Abdel-Dayem 2002) and Adnan *et al.*, (Al-Janabi, Al-Mehdawi *et al.*, 2012) demonstrated a strong negative correlation between hyperprolactinaemia and sperm count. Since LH is crucial for stimulating steroidogenesis and spermatogenesis while FSH directly nourishes the spermatozoa, the negative correlation may indicate poor spermatogenesis in hyperprolactinaemic patients.

Evidence has shown that hyperprolactinaemia has a deleterious effect on ATP synthesis in the mitochondria, lowering the Ca+2 channels necessary for the acrosome reaction and reducing sperm motility(Al-Daghistani and Abdel-Dayem 2002). Other studies(Al-Daghistani and Abdel-Dayem 2002, Ahmed and Ahmed 2017) found a strong inverse relationship between hyperprolactinaemia and sperm motility. However, a study from Israel(Al-Janabi, Al-Mehdawi *et al.*, 2012) contradicts this.

This study demonstrates a statistically significant correlation between sperm morphology and high prolactin levels, comparable to the finding reported by *Al-Daghistani et al.*, (Al-Daghistani and Abdel-Dayem 2002). The association between prolactin levels and sperm morphology may result from prolactin's role in preserving the structural integrity of spermatozoa (Coutton, Fissore *et al.*, 2016). In proliferative cells, serum prolactin initiates the Ca<sup>2+</sup> pool movement, which is crucial for gene expression, cell development, and protein synthesis—all of which are components of the standard structure and morphology of the cell (Silvestroni and Menditto 1989,

Espino, Mediero *et al.*, 2009, Tritos and Klibanski 2019). These processes are disrupted, and an aberrant prolactin level impacts sperm morphology.

Other noteworthy study results revealed that smoking, alcohol use, and obesity were all significant predictors of aberrant sperm counts (After controlling for confounding variables). In other words, a greater BMI and more alcohol use are linked to a decreased sperm count. Other researchers reported similar findings (Muthusami and Chinnaswamy 2005, Geidam, Yawe *et al.*, 2008).

The regulation of the hypothalamic-pituitarygonadal axis is compromised by alcohol consumption and obesity. The pulsatile release of follicle-stimulating hormone and the luteinizing hormone is negatively impacted. These lower testosterone levels, reduce sperm generation and gradually damage the testicles (Asare-Anane, Bannison *et al.*, 2016).

Smoking also increases cadmium intake as tobacco plants absorb metal. As a result of its similar chemical properties to zinc, cadmium could replace zinc in DNA polymerase. The deficiency of zinc induces atrophy of seminiferous tubules, a secretory deficit of Leydig and Sertoli cells and subsequent failure of spermatogenesis(Lingappa, Govindashetty *et al.*, 2015, Asare-Anane, Bannison *et al.*, 2016).

This study has shown that hyper prolactin negatively affects the seminal fluid analysis parameters. Therefore, routine assay for serum prolactin levels for infertile men with abnormal SFA may help the couple, instead of the financially laden Assisted Reproduction Technology (ART), which is beyond the reach of most of the couples in our low-resource setting.

### ETHICAL CONSIDERATION

Approval for the study was obtained from the Ethical and Research Committee of the University of Benin Teaching Hospital.

#### **CONFLICT OF INTEREST**

There was no conflict of interest in the conduct of the study.

#### **REFERENCES**

- Abdulhadi, S., Kawo, A., Arzai, A., & Hamza, S. (2021). Semen Parameters of Infertile Out Patients Attending Murtala Muhammed Specialist Hospital, Kano, Nigeria. *International Journal of Biomedical and Health Sciences*, *3*(1).
- Ahmed, N. D., & Ahmed, H. S. (2017). The relationships between serum prolactin level and semen analysis parameters among patients attended the reproductive health centre, Khartoum 2016–2017. *Sudan Medical Monitor, 12*(2), 41.

- Ajah, L. O., Ozumba, B. C., Onoh, R. C., Ezeonu, P. O., Iyoke, C. A., & Ezeome, I. V. (2016). Seminal fluid features of male partners of infertility patients in southeast Nigeria. *African J Medic Health Sci*, 5(2), 85-91.
- Al-Daghistani, H. I., & Abdel-Dayem, M. (2002). Hyperprolactinemia and hypergonadotropins in infertile males with severe oligospermia and azoospermia. *Internet J Endocrinol*, *3*, 1540-2606.
- Al-Janabi, A. S., Al-Mehdawi, F. A., & Al-Lami, M. Q. (2012). Relationship of seminal biochemical parameters and serum reproductive hhormones with sperm function tests in asthenospermic patients. *Jordan Med J*, *46*, 97-107.
- Ambulkar, S. S., Darves-Bornoz, A. L., Fantus, R. J., Wren, J., Bennett, N. E., Halpern, J. A., & Brannigan, R. E. (2022). Prevalence of hyperprolactinemia and clinically apparent prolactinomas in men undergoing fertility evaluation. *Urology*, *159*, 114-119.
- Anawalt, B. D. (2013). Approach to male infertility and induction of spermatogenesis. *The Journal of Clinical Endocrinology & Metabolism*, 98(9), 3532-3542.
- Asare-Anane, H., Bannison, S. B., Ofori, E. K., Ateko, R. O., Bawah, A. T., Amanquah, S. D., ... & Ziem, J. B. (2016). Tobacco smoking is associated with decreased semen quality. *Reproductive Health*, *13*(1), 1-6.
- Babakhanzadeh, E., Nazari, M., Ghasemifar, S., & Khodadadian, A. (2020). Some of the factors involved in male infertility: a prospective review. *International journal of general medicine*, 29-41.
- Balen, A. H., Morley, L. C., Misso, M., Franks, S., Legro, R. S., Wijeyaratne, C. N., ... & Teede, H. (2016). The management of anovulatory infertility in women with polycystic ovary syndrome: an analysis of the evidence to support the development of global WHO guidance. *Human reproduction update*, 22(6), 687-708.
- Benjamin, U. O., Akhere, T. I., & Orhue, A. A. (2014). prevalence and The patterns of male endocrinopathies amongs azoospermic partners at a fertility clinic Benin in city. Endocrinol Metab Int J, 1(1), 8-13.
- Coutton, C., Fissore, R. A., Palermo, G. D., Stouffs, K., & Touré, A. (2016). Male infertility: genetics, mechanism, and therapies, *Hindawi*, 2016.
- Emokpae, M. A., Uadia, P. O., Omale-Itodo, A., & Orok, T. N. (2007). Male infertility and endocrinopathies in Kano, Northwestern Nigeria. *Annals of African Medicine*, *6*(2), 64.
- Espino, J., Mediero, M., Lozano, G. M., Bejarano, I., Ortiz, Á., García, J. F., ... & Rodríguez, A. B. (2009). Reduced levels of intracellular calcium releasing in spermatozoa from asthenozoospermic patients. *Reproductive Biology and Endocrinology*, 7(1), 1-11.

- Geidam, A. D., Yawe, K. D. T., Adebayo, A. E. A., & Idrisa, A. (2008). Hormonal profile of men investigated for infertility at the University of Maiduguri in northern Nigeria. *Singapore medical journal*, 49(7), 538.
- Green, K. I., & Amadi, C. (2020). Status of serum prolactin levels among male cohort in infertile couples. *International Journal of Applied and Basic Medical Research* 10(4), 245.
- Hasan, R., & Wijesinghe, J. A. A. S. (2016). Implications of prolactin abnormalities on the male reproductive tract and male factor infertility.
- Ibadin, M. O., & Akpede, G. O. (2021). A revised scoring scheme for the classification of socioeconomic status in Nigeria. *Nigerian Journal of Paediatrics*, 48(1), 26-33.
- Juneja, P., Phukan, P. K., & Changmai, D. (2019). A study of abnormal semen parameters in infertile couples in Assam, India. *International Journal of Reproduction, Contraception, Obstetrics and Gynecology*, 8(3), 997-1001.
- Kaiser, U. B. (2012). Hyperprolactinemia and infertility: new insights. *The Journal of clinical investigation* 122(10), 3467-3468.
- Kumar, N., & Singh, A. K. (2015). Trends of male factor infertility, an important cause of infertility: A review of literature. *Journal of human reproductive sciences*, 8(4), 191.
- Kutlešić, R., Popović, J., Stefanović, M., Janošević, D. R., Živadinović, R., Petrić, A., & Trenkić, M. (2016). Association between hyperprolactinaemia and other causes of female infertility. *Facta Universitatis, Series: Medicine and Biology*, *17*(1), 36-41.
- Lingappa, H. A., Govindashetty, A. M., Puttaveerachary, A. K., Manchaiah, S., Krishnamurthy, A., Bashir, S., & Doddaiah, N. (2015). Evaluation of effect of cigarette smoking on vital seminal parameters which influence fertility. *Journal of clinical and diagnostic research: JCDR*, 9(7), EC13.
- Madziyire, M. G., Magwali, T. L., Chikwasha, V., & Mhlanga, T. (2021). The causes of infertility in women presenting to gynaecology clinics in Harare, Zimbabwe; a cross sectional study. *Fertility Research and Practice*, 7, 1-8.
- Muthusami, K. R., & Chinnaswamy, P. (2005). Effect of chronic alcoholism on male fertility hormones and semen quality. *Fertility and sterility*, 84(4), 919-924.
- Okada, H., Iwamoto, T., Fujioka, H., Shirakawa, T., Tatsumi, N., Kanzaki, M., ... & Ishigami, J. (1996). Hyperprolactinaemia among infertile patients and its effect on sperm functions. *Andrologia*, 28(4), 197-202.

- Oladosu, W. O., Biliaminu, S. A., Abdulazeez, I. M., Aliyu, G. G., Adelekan, A., & Okesina, A. B. (2017). Reproductive hormonal profile patterns among male partners of infertile couples at the University of Ilorin teaching hospital. *Afr J Infert Assist Concept*, 2, 6-10.
- Omokanye, L. O., Olatinwo, A. O., Durowade, K. A., Abdul, I. F., & Biliaminu, S. A. (2016). Determinants of infertility in male partners of infertile couples at a public health facility in Ilorin, Nigeria.
- Onyebuchi, A. K., Ifeoma, E. C., Mamah, J., Iwe, B., Afogu, E., & Obi, V. O. (2018). Seminal fluid analysis of male partners of infertile couples in Abakaliki, Ebonyi State, Nigeria. *Tropical Journal of Obstetrics and Gynaecology*, *35*(3), 261-265.
- Organization, W. H. (2023). Infertility prevalence estimates: 1990–2021.
- Osaikhuwuomwan James, A., & Osemwenkha Abieyuwa, P. (2015). Etiological pattern of infertility; an appraisal of contemporary trend in the region of Niger-Delta. *International Journal of Medical and Health Research Sep*, 1(2), 75-77.
- Ozoemena, O. F. N., Ezugworie, J. O., Mbah, A. U., Esom, E. A., Ayogu, B. O., & Ejezie, F. E. (2011). Abnormality of pituitary gonadal axis among nigerian males with infertility: study of patterns and possible etiologic interrelationships. *Open Access Journal of Urology*, *3*, 133.
- O'Leary, K. (2020). Hyperprolactinemia: Effect on Reproduction, Diagnosis, and Management. *Textbook of Assisted Reproduction*, 141-148.
- Samperi, I., Lithgow, K., & Karavitaki, N. (2019). Hyperprolactinaemia. *Journal of clinical medicine*, 8(12), 2203.
- Shahroona, M., Fatima, M., & Muhammad, B. (2007). Severe hyperprolactinemia directly depresses the gonadal activity causing infertility.
- Sharma, A. (2017). Male infertility; evidence, risk factors, causes, diagnosis and management in human. *Ann Clin Lab Res*, *5*(3), 188.
- Silvestroni, L., &. Menditto, A. (1989). Calcium uptake in human spermatozoa: characterization and mechanisms. *Archives of andrology*, 23(2), 87-96.
- Tritos, N. A., & Klibanski, A. (2019). Prolactin and its role in human reproduction. *Yen and Jaffe's Reproductive Endocrinology*, Elsevier: 58-74. e58.
- Uadia, P. O., & Emokpae, A. M. (2015). Male infertility in Nigeria: A neglected reproductive health issue requiring attention. *Journal of Basic and Clinical Reproductive Sciences*, 4(2),45-53.
- Wong, A., Eloy, J. A., Caldwell, W. T., & Liu, J. K. (2015). Update on prolactinomas. Part 1: Clinical manifestations and diagnostic challenges. *Journal of clinical neuroscience*, 22(10), 1562-1567.