# Determinants of Obstetricians' Pattern of Care for Sickle Cell Disease in Pregnancy

### Abstract

Background: Pregnancy in sickle cell disease (SCD) is high risk. With improved comprehensive obstetric care, pregnant females with SCD can achieve successful pregnancy outcomes, especially in resource-poor settings. Objectives: To determine the predictors of Obstetricians' pattern of care for SCD in pregnancy in Nigeria. Materials and Methods: Self-administered, pre-tested, pre-validated questionnaires containing 18 questions on demographic details of obstetricians, and their pattern of practice towards antenatal care for pregnant SCD patients were distributed to attendees of the 2018 conference of the Society of Obstetrics and Gynaecology of Nigeria (SOGON). Regression analysis was done to determine the possible predictors, and a significant level was <0.05. Result: Almost all the respondents (98.4%) considered pregnancy in SCD as high risk, and 96.2% proposed for preconception care in a tertiary hospital. The majority, (62%) agreed that antenatal visits in the first and second trimesters should be more frequent. The majority (96.2%) reported they would routinely order urine tests among other investigations. Majority of respondents, 74.9% and 98.4% knew that foetal medicine specialists and haematologists should be part of preconception care team, respectively. Respondents' practice centre and designation, significantly contributed to their "willingness to consult a haematologist" (P = 0.004)," and willingness to consult a foetal specialist" (P = 0.047), while practice centre and practice population significantly contributed to their response to "ideal centre for management of SCD pregnancy": (P = 0.049), (P = 0.024) respectively. Conclusion: Obstetricians' level of training, practice centre, and practice population of pregnant women with SCD are significant contributors to their pattern of care towards antenatal care for pregnancy in SCD.

Keywords: Antenatal services, blood transfusion, obstetricians, sickle cell disease

### Introduction

Sickle cell disease (SCD) refers to a group of genetic diseases that results from the replacement of normal adulttype haemoglobin (HbA) with abnormal mutant haemoglobins notably the sickle haemoglobin.<sup>[1-3]</sup> The presence of haemoglobin S in a homozygous state is sickle cell anaemia, while the coexistence of mutant haemoglobin S in a heterozygous state with any other abnormal haemoglobin results in the classical clinical findings is referred to as sickle cell disease4. SCD is a protean manifestation as it affects virtually all organ systems of the body. The most common recognizable clinical presentation is chronic haemolysis, bone pains, and fever, while the most striking diagnostic finding is the presence of sickled red blood cells on peripheral blood smears.<sup>[2,4,5]</sup> The most common inherited disorders of haemoglobin. Inheritance is mendelian and usually autosomal recessive in nature.<sup>[6]</sup>

It is of enormous public health importance in the sub-Saharan region countries, especially Nigeria, which carry the greatest burden. 25% of the population has this trait, while the prevalence of the disease prevalence is 1-3% of live births.<sup>[5,7,8]</sup> The clinical findings in SCD usually result from an intrinsic inability of Sickle haemoglobin to resist drops in oxygen tension. Low oxygen pathologically leads to polymerization of S-haemoglobin, consequently, sickling of the affected red cells occurs.<sup>[4,9]</sup>

Pregnancy in sickle cell disease (SCD) is a high risk, both for the mothers and foetus.<sup>[10]</sup> Also, the frequencies of complications of pregnancies are more in women with SCD than observed in the normal population.<sup>[11]</sup> This is because the clinicopathological findings of chronic haemolytic anaemia, repeated vaso-occlusion and resultant multi-

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organ dysfunction syndrome seen in sickle cell disease will be severely negatively impacted by the physiologic changes seen in pregnancy.<sup>[6,11,12]</sup>

Despite this seemingly bleak outlook, pregnancy is not contraindicated in sickle cell disease.<sup>[13]</sup> This is important to note because improvements in the health care delivery system over the years have significantly improved the survival of SCD patients to adulthood, with an associated desire of many to achieve pregnancy.<sup>[6,14]</sup> There is thus an increased need for improved obstetric care especially antenatal care within the context of a multi-disciplinary approach for this group especially in resource-poor settings to reduce poor outcomes resulting in increased maternal morbidity and mortality.<sup>[12-14]</sup> This work is critically directed at evaluating the practice pattern of Nigerian obstetricians towards the delivery of effective antenatal services and the management of pregnancy in sickle cell disease.

The findings of this study are believed to fill an important knowledge gap, as literature searches have shown a severe dearth of published information on the practice patterns and clinical skills of Nigerian obstetricians in the management of pregnancy in sickle cell disease.

# **Subjects and Methods**

### Patients and design

This was a cross-sectional study of 183 obstetrician and specialist Registrar attendees of the 2018 annual conference of the society of obstetrics and Gynaecology of Nigeria (SOGON). Ethical approval with the number NHREC/08/2008B-FWA0002458-1RB00002323 was obtained from the UNTH research ethics committee. The study period was November 2018 to June 2019.

### **Study tool**

Investigators designed self-administered, structured, pretested, and prevalidated questionnaires were used. It consists of 18 items on demographic details of obstetricians, knowledge of antenatal services, and pattern of antenatal care or pregnant SCD patients, with a Cronbach alpha of 0.85. Practice pattern and perception of antenatal services were the main outcomes measured. Sociodemographic characteristics: designation, years of practice, practice centre, and practice populations were the primary dependent variables of interest

### **Ethical consideration**

This was sought for and obtained from the institutional review board (IRB) of University of Nigeria Teaching Hospital Enugu (NHREC/05/01/2008B-FWA00002458-1RB00002323). All participants gave informed consent to participate, and the research was carried out in accordance with the declaration of Helsinki.

### Statistical analysis

Descriptive statistics were calculated to assess the sociodemographic characteristics of the subject. Fischer's

exact test was used to examine the associations between sociodemographic and outcome variables. Regression analysis. was done to determine possible predictors. A significant level was set at <0.05. All analyses were performed using the Chicago Illinois Statistical Package for Social Sciences (SPSS, Version 22.0).

# Result

## Sociodemographic variables of the respondents

Of the 183 participating obstetricians, 53.6% were specialist senior registrars, while 46.4% were consultants. The malefemale ratio was 3:1 with 76% males and 24% females. Most of the participants were aged 37 years and older. The majority (92.9%) of the respondent's practice were in a tertiary health institution and almost half (47%) have a yearly practice population of 0–5 clients. [See Table 1] Respondents' Pattern of antenatal care for pregnant women with SCD

Of the study participants, the majority 62% and 54.6% reported that antenatal visits in the first trimester and second trimester should be weekly respectively [see Table 2]. Almost all (98.4%) considered pregnancy in SCD as high risk, and 96.2% proposed for preconception care in a tertiary hospital, respectively. Most 96.2% reported that they routinely order urine tests among other antenatal investigations. [See Figure 1]

Table 1: Sociodemographic variables		
Demographic	Frequency	Percent
Gender		
Male	139	76.0
Female	44	24.0
Age category		
22-26	2	1.1
27-31	12	6.6
32-36	37	20.2
37-41	55	30.1
41 and above	77	42.1
Training		
Senior Resident	98	53.6
Consultant	85	46.4
Practice Centre		
General Hospital	11	6.0
Tertiary	170	92.9
Private	-	-
Not sure	2	1.1
Yearly Practice population		
(SCD pregnancy)		
0-5	86	47.0
6-10	50	27.3
11-15	18	9.8
16-20	5	2.7
>20	21	11.5
Not sure	3	1.6

Table 2: Frequency of ante natal visits of pregnant women   with SCD			
Ante natal visits of pregnant women with SCD.	Frequency	Percent	
Frequency of antenatal visit (first trimester)			
Weekly	7	3.8	
Fortnightly	113	61.7	
Monthly	53	29.0	
Not sure	10	5.5	
Frequency of antenatal visit (2 <sup>nd</sup> trimester)			
Weekly	62	33.9	
Fortnightly	100	54.6	
Monthly	10	5.5	
Not sure	11	6.0	
Frequency of antenatal visit (3 <sup>rd</sup> Trimester)			
Weekly	157	85.8	
Fortnightly	12	6.6	
Monthly	3	1.6	
Not sure	11	6.0	



Figure 1: Routine investigation in antenatal delivery



Figure 2: frequency of lab work in the first trimester

On the issue of laboratory investigations during ante natal care, approximately 43.2% indicated that frequency of lab work during the first trimester should be monthly, while 25.1% indicated that it should be on each visit. [See Figure 2]. For the second trimester and third trimester, 24.6% 44.3% of the health workers indicated that laboratory work should be at each visit, respectively. See [Figures 3 and 4].

On using a multi-disciplinary management approach, the majority (74.9%) of the participants were willing to



Figure 3: Frequency of lab works 2nd trimester



Figure 4: Frequency of laboratory work in the third trimester

consult a foetal medicine specialist in the management of pregnant sickle cell patients and up to 19.7% indicated they were willing but never had a foetal medicine specialist in their facility. The proportion of respondents who indicated that foetal medicine specialists and haematologists should be part of the management team was 74.9% and 98.4%, respectively. Again, (98.4%) also perceive SCD pregnancy as a high-risk pregnancy. This is shown in [Table 3].

On the issue of blood transfusion support, the highest indication for blood transfusion as described by the respondents was acute anaemia (93.4%), followed by a painful crisis (33.3%) and acute chest syndrome (26.2%). Approximately 86.9% of the participants transfused pregnant women with SCD only when indicated. Also, about 6% and 2% of the respondents used regular top transfusion and exchange blood transfusion, respectively while 3% were not sure of the transfusion pattern they commonly used.

Only 51.4% of the participants administered routine aspirin for the prevention of preeclampsia in pregnant women with SCD and 2.2% of them did not know about it.

# Association between sociodemographic characteristics and some outcome measures

Fisher's exact analysis showed that the designation of the respondents significantly contributed to their 'willingness to consult a foetal specialist' (P = 0.047) See [Table 4]. Their practice centre significantly contributed to their 'willingness to consult a haematologist' (P = 0.004) See

Table 3: Attitudes to multi-disciplinary management of SCD pregnant women		
Multi-disciplinary approach antenatal care management of SCD pregnant women	Frequency	Percent
Are you willing to consult a foetal medicine specialist in the management of the pregnant sickle cell patient		
Yes	137	74.9
No	7	3.8
Yes but I don't have a foetal medicine specialist in my centre	36	19.7
Not sure	3	1.6
Are you willing to consult a Haematologist		
Yes	180	98.4
Yes but I don't have	3	1.6
Do you regard related pregnancy as high-risk pregnancy		
Yes	180	98.4
Not sure	3	1.6

# Table 4: Association of the sociodemographic characteristics of the respondents and their perception of the willingness to consult a specialist in foetal medicine

Factors	Willing to consult a f	oetal medicine specialist	Total	Fisher test (P-value)
	No	Yes		
Age category				
22-26	0(0)]	2(1.5)	2(1.1)	0.535
27-31	1(2.2)	11(8.0)	12(6.6)	
32-36	12(26.1)	25(18.2)	37(20.2)	
37-41	14(30.4)	41(29.9)	55(30.1)	
42 and above	19(41.3)	58(42.3)	77(42.1)	
Gender				
Male	34(73.9)	105(76.6)	139(76.0)	0.84
Female	12(26.1)	32(23.4)	44(24.0)	
Training				
Senior resident	24(52.2)	74(54.0)	98(53.6)	0.047*
Consultant	22(47.8)	63(46.0)	85(46.4)	
Practice centre				
General hospital	3(6.5)	8(5.8)	11(6.0)	0.09
Tertiary	41(89.1)	129(94.2)	170(92.9)	
Private	2(4.3)	0(0)	2(1.1)	
The practice population				
0-5	29(63.0)	59(43.1)	88(48.1)	0.11
6-10	9(19.6)	42(30.7)	51(27.9)	
11-15	2(4.3)	16(11.7)	18(9.8)	
16-20	0(0)	5(3.6)	21(11.5)	
>20	6(13.0)	15(10.9)	21(11.5)	

[Table 5] and their practice centre and the population of practice significantly contributed to their perception of the 'ideal centre for the management of pregnancy with SCD' (P = 0.049) and (P = 0.024), respectively. In [Table 6], we saw an association between the practice population and their perception on ideal centre for the management of pregnancy in SCD.

# Discussion

Our study was able to assess Obstetricians' pattern of service in antenatal care delivery to pregnant sickle cell disease patients as well as their practice determinants aimed at improving care rendered to these patients. Almost all our respondents affirmed that pregnancy in SCD is high risk. Pregnancy is considered high risk if it is associated with a condition which poses an actual or a potential risk to either or both the mother and fetus.<sup>[15]</sup> Normal pregnancy is associated with some physiological changes including hypercoagulability, hyperviscosity and increased metabolic demand. These changes are exaggerated in SCD patients predisposing them to higher frequency of sickle cell crises and complications.<sup>[6]</sup> The risks of both obstetrics and non-obstetrics complications of pregnancy like preeclampsia and pneumonia are increased in SCD as well.<sup>[6]</sup> These place pregnancy in SCD as high risk.<sup>[6,16]</sup> The WHO recommends a 4-visit focussed antenatal care of which the first contact is between 8 to 12 weeks of gestation, second visit between 24–26 weeks, third visit at 32 weeks while the forth visit is between 36 and 40 weeks.<sup>[17]</sup> The recommendation is

consult a haematologist				
Factors	Willing to consult a Haematologist		Total	Fisher test
	No	Yes		(P-value)
Age category				
22-26	0(0)	2(1.1)	2(1.1)	0.84
27-31	0(0)	12(6.7)	12(6.6)	
32-36	1(33.3)	36(20.0)	37(20.2)	
37-41	1(33.3)	54(30.0)	55(30.1)	
42 and above	1(33.3)	76(42.2)	77(42.1)	
Gender				
Male	2(66.7)	137(76.1)	139(76.0)	0.56
Female	1(33.3)	43(23.9)	44(24.0)	
Training				
Senior resident	2(66.7)	96(53.3)	98(53.6)	1.00
Consultant	1(33.3)	84(46.7)	85(46.4)	
Practice centre				
General hospital	1(33.3)	10(5.6)	11(6.0)	0.004*
Tertiary	1(33.3)	169(93.9)	170(92.9)	
Private	1(33.3)	1(0.6)	2(1.1)	
The practice population				
0-5	2(66.7)	86(47.8)	88(48.1)	1.00
6-10	1(33.3)	50(27.8)	51(27.9)	
11-15	0(0)	18(10)	18(9.8)	
16-20	0(0)	5(2.8)	5(2.7)	
>20	0(0)	21(11.7)	21(11.5)	

Table 5: Association of the sociodemographic characteristics of the respondents and their perception of willingness to

# Table 6: Association of the sociodemographic characteristics of the respondents and their perception on ideal centre for the management of pregnancy in SCD

Factors	Ideal centre for	Ideal centre for management of pregnancy in SCD			Fischer
	General	Tertiary	Not sure		(P-value)
Age category					
22-26	0(0)	2(1.2)	0(0)	2(1.1)	0.34
27-31	0(0)	12(7.1)	0(0)	12(6.6)	
32-36	4(50.0)	32(18.8)	1(20.0)	37(20.2)	
37-41	1(12.5)	54(31.8)	0(0)	55(30.1)	
42 and above	3(37.5)	70(41.2)	4(80.0)	77(42.1)	
Gender					
Male	7(87.5)	129(75.9)	3(60.0)	139(76.0)	0.52
Female	1(12.5)	41(24.1)	2(40.0)	44(24.0)	
Training					
Senior resident	6(75.0)	91(53.5)	1(20.0)	98(53.6)	0.17
Consultant	2(25.0)	79(46.5)	4(80.0)	85(46.4)	
Practice centre					
General hospital	1(12.5)	8(4.7)	2(40.0)	11(6.0)	0.049*
Tertiary	7(87.5)	160(94.1)	3(60.0)	170(92.9)	
Private	0(0)	2(1.2)	0(0)	2(1.1)	
The practice population					
0-5	3(37.5)	83(48.8)	2(40.0)	88(48.1)	0.024*
6-10	1(12.5)	50(29.4)	0(0)	51(27.9)	
11-15	1(12.5)	16(9.4)	1(20.0)	18(9.8)	
16-20	2(25.0)	3(1.8)	0(0)	21(11.5)	
>20	1(12.5)	18(10.6)	2(40.0)	21(11.5)	

also that antenatal visit for a normal pregnancy should be 4 weekly from the time of booking until about the 36th week of gestation and then 2 weekly till 40 weeks.<sup>[17]</sup> Our respondents opined that antenatal visits should be twice weekly in the first and second trimester, while it should be weekly in the third trimester. The frequency of visits is higher for high-risk pregnancy and this frequency is determined by the nature of the risk.

Antenatal care (ANC) is aimed at preventing or minimizing the effects of health conditions during pregnancy from negatively affecting the mother and foetus.<sup>[18]</sup> The minimum ANC package by WHO called focused ANC comes with it some basic investigations to be done and they include packed cell volume/haemoglobin concentration, syphilis, HIV infection, urinalysis and urine microscopy for proteinuria and bacteriuria, and blood group/Rhesus factors determination.<sup>[17,18]</sup> It is recommended that all investigations be done at first contact while packed cell volume (PCV) or haemoglobin (Hb) estimation and urinalysis with microscopy be done in all four visits or where indicated.<sup>[17]</sup> Our study shows that over 60% of the participants would request for all the WHO recommended basic investigations during antenatal the frequency of which varied between individuals and trimester. Majority opined that these tests should be carried out monthly during the first trimester, at each scheduled visit during the second and third trimesters. These differences are understandable as it is recommended that the frequency of ANC visits for high-risk pregnancies should depend on the nature of the risk. It is only natural one would investigate his/her patients on contact. In addition to the basic recommended tests, our responded listed they would also test their SCD clients for end organ damage by doing both renal and liver function tests. This practice is in accordance with the recommendations from the Royal College of Obstetrics and Gynaecology guideline on the management of SCD in pregnancy.<sup>[19]</sup>

Most of the respondents would employ a multi-disciplinary approach in the management of SCD pregnancy-almost all (98.4%) would involve a haematologist while about three-quarters would involve a foetal medicine specialist. A multi-disciplinary approach in the ANC of SCD pregnancy is a good practice as there are evidence of better maternal and foetal outcome.[20,21] It is also in keeping in established management guidelines.<sup>[19]</sup> This notwithstanding, about 20% of our respondents who desired having a multi-disciplinary approach do not have access to foetal medicine specialist and may hamper on the expected pregnancy outcome. SCD is a chronic haemolytic anaemia and blood transfusion is a common practice in the management of SCD because patients may require transfusion at one time or the other for different reasons ranging from treatment of acute anaemia, acute chest syndrome, acute stroke, to prophylaxis of stroke or other severe disease complications.<sup>[2,5,22]</sup> During pregnancy, indications for transfusion may increase due to increased red cell demand as seen in multiple gestation and in women with severe medical, obstetrics and foetal complications.<sup>[19,23]</sup> Blood transfusion in SCD may be top-up transfusion or an exchange blood transfusion program.<sup>[2,5]</sup> A majority would transfuse a SCD patient when indicated of which over 90% of our respondents would transfuse for acute anaemia and just a few would transfuse for acute chest syndrome. Up to a third would wrongly transfuse a patient for painful crisis. Just a minority of the respondents understood the different transfusion practices, be it top-up or exchange blood transfusion. This shows that the transfusion practice among out respondents is not optimal. Again, despite positive reports from different group of researchers on the use of aspirin in the prevention of preeclampsia in SCD pregnant patients<sup>[24,25]</sup> and its recommendation in guidelines,<sup>[19]</sup> only about half of the respondents routinely administered aspirin in their patients. This may be due to lack of randomized controlled trials of the use of aspirin in a good number of pregnant SCD patients.<sup>[19]</sup>

We investigated the determinants of the different practices and observed that biological age, gender, centre of practice and practice population showed no association with their willingness to consult a foetal medicine specialist. It was interesting to observe that level of training (Consultant Vs Resident) showed a significant association. Our observation was that the more trained they are, the more unlikely they would seek the help of a foetal medicine specialist. Whereas we did not find any significant association between sociodemographic and willingness to seek expert advice from a Haematologist other than centre/place of practice. This may be explained by the fact that is more likely for Haematologists to be available or accessed in a tertiary institution than in secondary or private centres where they are unlikely to be employed.

Majority of the respondents believe that tertiary centres were the best for the management of SCD pregnancies. This is understandable because as recommended, management of SCD pregnancy is best under the care of a multi-disciplinary team<sup>[19]</sup> which is more likely to be available in a tertiary or bigger centres. Likewise, only practice centre and population of practice showed significant association with the ideal centre for SCD pregnancy management.

### **Conclusion and Recommendation**

A good number of the Obstetricians understand that SCD pregnancy is a high-risk pregnancy and should be managed by a multi-disciplinary team for a good outcome, however, factors like unavailability of specialist team members and poor understanding of transfusion practices in SCD are factors identified as drawbacks and may hamper the actualization of the goal of good maternal and foetal outcome. Obstetricians' level of training, practice centre, and practice population of pregnant women with SCD are significant contributors to their pattern of practice towards antenatal care for pregnancy in SCD. We therefore recommend establishment of national and local guidelines for management of pregnancy in SCD to guide Obstetricians in their practice.

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### **Conflicts of interest**

TUN, HCO, AOU, EOVU, AND, IUE, IEM, AOU and SCE declared no competing interest.

### Authors' contributions

TUN conceived and designed the study, interpreted the data and wrote the manuscript, HCO contributed to the study design, and interpretation of data, AOU, EOVU, AUD, IUE, IEM, AOU and SCE contributed in data interpretation. All authors contributed to manuscript writing and read and approved the final manuscript.

### List of Abbreviations

ANC	Antenatal care
Hb	Haemoglobin
HbA	Haemoglobin A
PCV	Packed cell volume
SCD	Sickle cell disease
SOGON	Society of Obstetrics and Gynaecology of Nigeria
WHO	World Health Organization

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