

Original Research Article

Pregnancy Outcomes in Women with Sickle Cell Disease (SCD) - A Tertiary Centre Retrospective Study in Nigeria.

Comment [U1]:

ABSTRACT

Background

Sickle cell disease is the most common inherited condition. Due to recent advances in medical care for sicklers, more women with SCD are able to survive up to the reproductive age group. This has given rise to a higher number of pregnant women with SCD. It results in increased fetal and maternal risks due to the metabolic demands, hypercoagulable state and vascular stasis often associated with pregnancy. It has therefore become imperative to explore what progress has been made over the last decade as well as what can be done to achieve the best outcome in these women.

Method

This was a retrospective study of women with SCD who accessed antenatal care (ANC) at the NnamdiAzikiwe University Teaching Hospital (NAUTH) Nnewi over a ten year period. Folder numbers were obtained from the obstetrics clinic and thereafter the folders were assessed and data retrieved using a checklist. Data analysis was done using SPSS 24.

Results

The prevalence of sickle cell disease among pregnant women in this centre was 0.03%. All the women had the HbSS variants. The mean age of the subjects was 28.3 ± 5.6 years. Majority of the women were gainfully employed. The mean gestational age at booking was 26 9.5 weeks. Half of the neonates had fetal distress at birth. Commonest maternal complication was anemia(100%) followed by vaso-occlusive crisis(89%). Eighty nine percent of the women were delivered by emergency caesarean section prior to 37 weeks gestational age. Preterm birth and fetal distress were the commonest neonatal complications.

Conclusion

The study revealed that pregnancy associated with sickle cell disease still carries high maternal and fetal morbidity in this environment. Majority of the women with SCD in pregnancy did not receive preconception care and also did not commence their ANC early enough. There is need to educate women with SCD on the need for preconception care to ensure optimization of health prior to conception. They should also book early enough to facilitate close fetal monitoring throughout pregnancy.

Keywords: sickle cell disease, antenatal care, maternal outcome, neonatal outcome

Comment [U2]: This word is not acceptable, it is stigmatizing, use instead "patients with sickle cell anaemia"

INTRODUCTION

Nowadays, with recent advances and improved healthcare, the life expectancy of people with SCD has improved and many more women are reaching reproductive age. A pregnant woman with SCD is at risk of developing sickle cell crises. SCD is a group of haemoglobin disorders that result from the inheritance of the sickle cell β -globin gene. It can occur in the homozygous form as sickle cell anaemia (SS) or in double heterozygous form with other types of abnormal Hb such as the Sc and S β thal variants¹. It is a common disease of tropical origin affecting countries within the Mediterranean, Asia and Sub-Saharan Africa to which Nigeria belongs to². The inheritance pattern is autosomal recessive and as such both parents must have the sickle cell gene. The mainstay of treatment has often been prevention and appropriate management of any of the crises in individuals suffering from SCD with some advances in gene therapy.

Comment [U3]: This sentence should come after the explanation on the meaning of SCD

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During pregnancy, SCD can become more severe and pain episodes occur more frequently. This tends to pose a great challenge to their care because of the increased risk of pregnancy complications such as toxemia of pregnancy, postpartum haemorrhage, infections, prematurity, intrauterine growth, restriction (IUGR), low birth weight, abortions, stillbirth, urinary tract infections, stroke, thromboembolic episodes, heart failure, neonatal and maternal mortality³. Anaemia often occurs as a result of the deoxygenation of the sickle cells which become sickled and are then removed from the body by the reticuloendothelial system with a subsequent red blood cell life span reduction to 17 days and then a resultant chronic compensated anaemia⁶. There is also associated high perinatal mortality. Earlier studies on pregnancy outcomes in sickle cell patients revealed universal adverse outcome but with recent medical advances and care, especially the outcome seems to have greatly improved^{4,7}.

Maternal morbidity is increased in SCD due to infections (UTIs), cerebrovascular accident and sickle cell crisis. Maternal mortality is increased by as much as 25% due to pulmonary infarction, acute chest syndrome, congestive heart failure and embolism³. The frequency of crises increases significantly especially in 3rd trimester as a result of increased oxygen consumption, blood viscosity & red cell mass. In addition, SCD constitutes a financial stress in view of the repeated hospital admissions, repeated transfusion and frequent pain medications given to these patients. While some studies have revealed improved outcomes for pregnant women with SCD as well as the majority of them being able to achieve a successful live birth, pregnancy with SCD is still associated with an increased incidence of morbidity and mortality⁷.

In a study done in Enugu state, a state in the same geographical zone as the centre, the prevalence rate of SCD in pregnancy was 0.01% whereas that in Ebonyi state was 0.69%^{8,9}. The study done in Port Harcourt (SS zone) had a prevalence rate of 0.14% in a ten-year study and 0.2% in a five-year study⁹.

In West Africa, increasing rates of pregnant women with SCD are being seen. A study done in Ghana had a prevalence rate of 1.42% over a two year period¹⁰. Similar values were also seen in India where a 4 year study period revealed a prevalence of 1.2%¹¹. Considering the increase in various parts of the world, it has become obvious that

more pregnant women with SCD are being encountered now in medical practice. In the first half 20th century, women with SCD barely survived to reproductive age. Early medical experience with SCD and pregnancy was a cause for concern. The first successful pregnancy was in 1931 and with the major review that occurred a decade later, a 50% fetal loss was obtained. 11 Maternal mortality rates as high as 11.5% were reported from West Africa and from black American groups¹². There has been a number of varying reports on maternal and fetal outcome in Nigeria as well as Africa. This study was undertaken to assess the current pregnancy outcomes among women with SCD in our centre as there has been no such previous study in this environment.

Objectives

The study sought to determine the maternal and fetal outcomes in pregnant women with SCD attending antenatal clinic at NnamdiAzikiwe University Teaching Hospital, Nnewi, Anambra State

Materials and methods

Study Site: The study site was NnamdiAzikiwe University Teaching Hospital (NAUTH) Nnewi, Anambra state Nigeria. NAUTH is the only federal government owned tertiary hospital in the state. It offers primary, secondary and tertiary services in the state. Nnewi is the second most populated town after Onitsha in Anambra state and is also located very close to Onitsha and Awka which are two densely populated areas in the State. The hospital in addition also serves as a referral centre for neighboring states.

Study Population: All pregnant women with SCD who had accessed care from NAUTH, Nnewi from January 1st to December 31st 2020.

Inclusion Criteria:

Pregnant women who had been previously diagnosed with SCD and accessed antenatal care (ANC) as well as had their deliveries at NAUTH from January 1st 2011 to December 31st 2020.

Exclusion Criteria: Pregnant women who accessed ANC at the hospital but did not deliver in the hospital as well as those who had their delivery in the hospital but had accessed ANC care elsewhere

Sample size determination:

The sample size was determined using the Cochran formula¹³

$N = Z^2pq/d^2$ with a p-value of 0.01 (proportion of SCD patients accessing antenatal care from a previous study)⁹

$N = (2.96)^2 \times 0.01 \times 0.99$
 $(0.05)^2$

= 15

Sampling Technique: A non-random sampling technique was employed as all case folders that met the inclusion criteria within the study period were reviewed

Data Collection

With permission from the records department, obstetrics records of women with sickle cell disease were identified, studied and required data retrieved with a checklist that had been designed to record patient's profile.

This profile included age, highest educational qualification, occupation, marital status, parity, gestational age at booking, haemoglobin at booking, transfusion history, obstetric and non-obstetric complications in all trimesters, mode and outcome of delivery, neonatal complications. The diagnosis of SCD was made using Haemoglobin electrophoresis machine.

Anaemia was defined as a haemoglobin of less than 10g/dl

Vasocclusive crisis was defined as a painful episode with subjective complaint of pain and exclusion of other causes of pain based on related symptoms.

Malaria was defined as presence of malaria parasites in thick or thin peripheral blood films in the laboratory.

Data Analysis:

All available data were analyzed using SPSS 25 with results presented in form of frequency tables. Descriptive statistics in the form of the frequency and percentages were calculated.

Ethical Considerations:

Approval for the study was obtained from the NnamdiAzikiwe University Teaching Hospital Research and Ethics Committee.(date of approval 26th June 2021).

Results:

Out of 71, 737 women who accessed antenatal care during the period of study, 18 were diagnosed of SCD, giving a prevalence rate of 0.03%. The mean age of the respondents was 28.3 ± 5.6 years while the mean gestational age at booking was 26 ± 9.5 wks. Majority of them (77.8%) were married while 38.5% of them were nulliparous. More than half of them (55.6%) registered in their 3rd trimester. About 44% of the women had tertiary education. The booking haemoglobin was 6.5 ± 3.5 g/dl. All the women had the HbSS variant. The commonest complication was anaemia (100%) with 72.2% of them being transfused. Majority of them (77.8%) had vaso-occlusive crisis while 88.9% of them delivered by caesarean section.

Half of the women (50%) had pre-term delivery while 50% of the newborn had fetal distress. The recorded neonatal death was 11% while stillbirth recorded was 5.6%. There were 33% newborns with low birth weight.

Tables:

Table 1: SOCIO DEMOGRAPHIC CHARACTERISTICS OF THE RESPONDENTS

VARIABLE	FREQUENCY	PERCENTAGE
AGE(years)		
21-25	7	38.9
26-30	6	33.3
31-35	2	11.1
36-40	3	16.7
TOTAL	18	100.0
MEAN	28.3±5.6 years	
HIGHEST LEVEL OF EDUCATION		
JSCE	2	11.1
SSCE	5	27.8
B.Ed	2	11.1
B.Sc	6	33.3
MISSING	3	16.7
TOTAL	18	100.0
OCCUPATION		
STUDENT	1	5.6
TEACHER	2	11.0
TRADER	7	38.9
CIVIL SERVANT	5	27.8
MISSING	3	16.7
TOTAL	18	100.0
MARITAL STATUS		
SINGLE	4	22.2
MARRIED	14	77.8
TOTAL	18	100.0

PARITY		
P0	7	38.9
P1	6	33.3
P3	2	11.1
MISSING	3	16.7
TOTAL	18	100.0
GESTATIONAL AGE AT BOOKING		
(0 – 13WKS)	2	11.1
(14 – 26WKS)	6	33.3
(27 – 40wks)	10	55.6
TOTAL	18	100.0
MEAN	26 ± 9.5wks	
HAEMOGLOBIN AT BOOKING(g/dl)		
5.0	4	22.2
5.2	1	5.6
6.0	4	22.2
7.0	3	16.7
8.0	3	16.7
9.0	1	5.6
MISSING	2	11.1
TOTAL	16	88.9
MEAN HAEMOGLOBIN	6.5 ± 3.5g/dl	

TABLE 2: PREVALENCE OF SICKLE CELL DISEASE

VARIABLES	FREQUENCY	PERCENTAGE (%)
THOSE WITH SCD	18	0.03
THOSE WITHOUT SCD	71719	99.97
TOTAL	71737	

RECEIVED PRECONCEPTION CARE	
CARE	0
YES	16
NO	2
Missing	

TABLE 3: TABLE SHOWING THE VARIOUS TYPES OF SCD SEEN AMONG THE RESPONDENTS

VARIABLE	FREQUENCY	PERCENTAGE (%)
HbSS VARIANT	18	100
HbSC VARIANT	0	0
HβTHAL VARIANT	0	0
TOTAL	18	

TABLE 4: COMPLICATIONS IN WOMEN WITH SICKLE CELL DISEASE ACCESSING ANC AT NAUTH WITHIN THE TEN (10) YEARS OF STUDY

VARIABLE	FREQUENCY	PERCENTAGE (%)
PRE ECLAMPSIA		
YES	3	16.7
NO	13	72.2
MISSING	2	11.1
TOTAL	16	88.9
MALARIA		
YES	4	22.2
NO	14	77.8
TOTAL	18	100
TRANSFUSION HISTORY		
YES	13	72.2
NO	3	16.7
MISSING	2	11.1
TOTAL	16	88.9
ANAEMIA		
YES	18	100.0
NO	-	0.0
TOTAL	18	100.0
VASO-OCCLUSIVE CRISIS		
YES	14	77.8
NO	2	11.1

MISSING	2	11.1
TOTAL	16	88.9
INTRAUTERINE FETAL DEATH		
YES	2	11.1
NO	14	77.8
MISSING	2	11.1
TOTAL	16	88.9
RESPIRATORY TRACT INFECTION		
YES	3	16.7
NO	12	66.7
MISSING	3	16.7
TOTAL	15	83.3
EMERGENCY CESAREAN SECTION		
YES	16	88.9
NO	2	11.1
TOTAL	18	100.0
WOUND INFECTION		
YES	2	11.1
NO	16	88.9
TOTAL	18	100.0
MATERNAL DEATH		
YES	1	5.6
NO	17	94.4
TOTAL	18	100.0

Discussion

The prevalence of sickle cell disease in pregnancy in this study is 0.03%. This appears to be similar to the rate obtained at Enugu (0.01%) in Enugu state, a state in the same geographical zone as our centre. A higher value of 0.2% was obtained at Port Harcourt in South-South region of Nigeria¹⁷. Considering other West African countries, Ghana had a prevalence rate of 1.4% whereas 0.09% was obtained in Tanzania of East Africa. This lower value may have been as a result of the rule in this state where intending couples must present their genotype results before marriage institutions that end up discouraging those with the trait from getting married to each other. Studies done in other continents revealed relatively higher prevalence rates as seen in Saudi Arabia, India and USA.^{14,15,16}

All the SCD cases seen in this study were of the HbSS variant. This is contrary to the findings in other centres where there were also other variants.^{18, 8,19}. This could be

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attributed to the fact that this centre uses haemoglobin electrophoresis machine for diagnosis instead of High Performance Liquid Chromatography.

Majority of the women (88.9%) in this study booked in their 2nd or 3rd trimester. Only 11% of the women booked in first trimester. The mean gestational age at booking was 26 ± 9.5 wks with only 2 women booking in 1st trimester. This is even later than the mean gestational age of booking recorded by Ugboma & George (16.6 ± 3.3) in Port Harcourt and Kahansim ML et al (19.3 ± 7.7) in Jos. Booking early for antenatal care enables the health provider to monitor as well as identify risk factors related to prior maternal and fetal outcomes. Managing a pregnant woman with SCD requires close monitoring as anaemia can worsen very quickly⁶. Initiating appropriate medical interventions will then decrease the risks for both maternal and fetal morbidity and mortality.^{20,21} Other earlier studies had also recorded late booking in these women and most had almost all the women booking late in third trimester.^{21,17} The commonest maternal complication in this study was anaemia. Most of these women presenting late came in with severe anaemia as shown by the mean booking Hb of 6.5 ± 3.5 g/dl. This is similar to the finding in other centres where anaemia was present in many of the women. The value for anaemia obtained in this study is higher than that obtained in Jos (62.9%) but similar to the study finding of 92.2% in Abraka, Delta state.²³

Anaemia is one of the major complications of SCD and could be caused by hemolysis or red blood cell trapping in the spleen²². Anaemia in pregnancy has been found to be associated with increased risk of preterm premature rupture of membranes, spontaneous preterm labour, preterm delivery, intrauterine growth retardation as well as low birth weight babies. 8, 22

The anaemia resulted in all the women in this study being transfused prior to delivery. This high rate of antenatal transfusion (72.2%) has serious implications in view of the risk of infection transmission and alloimmunization. Anaemia was closely followed by vaso-occlusive crisis in form of bone pain with a frequency of 77.8%. This is much higher than that obtained in Port Harcourt (50%) and Jos (51.4%), Abraka (23.8%) and Ibadan (7.3%)²⁴. Studies outside the country also recorded lower values – USA (50%) and Benin Republic (57%). This higher incidence of vaso-occlusive crisis could be attributed to the fact that this centre serves mainly as a referral centre in this environment as more patients usually prefer to access care primarily from private hospitals²⁵.

There was a low incidence of malaria in pregnancy (22.2%) observed in this study. This is similar to the finding by Kahansim ML et al in Jos (25.7%) as well as that by Odum et al in Lagos (22.4%)²⁶ but contrary to that by Ugboma in Port Harcourt (80%). One would have expected a higher incidence of malaria in this study considering the fact that the study area is an area of high malaria transmission.

The incidence of pre-eclampsia in this study was 16.7%. This is quite lower than the 28.9% obtained in Jos. While Ugboma in Port Harcourt recorded maternal deaths from pre-eclampsia, some other studies had no significant increase in pre-eclampsia amongst pregnant women with SCD^{27,28}. More studies may be needed in this area.

The incidence of intrauterine fetal death (IUFD) in this study was 11%. This is less than the 20% obtained in both North Central and South-South region of Nigeria as well as in

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Jamaica²⁹. This highlights the need to monitor the pregnancy with serial sonograms for proper assessment of fetal growth. In this study, respiratory tract infection (16.7%) and wound infection (11.1%) were the least maternal complications. Identification and treatment of underlying infections is key in reducing the respiratory tract infections. Pneumococcal vaccines could be given where available.

The commonest mode of delivery was by caesarean section (88.9%) followed by spontaneous vaginal delivery (11%) and all of them were emergency caesarean section(C/S). This centre had a higher CS rate than that obtained in Port Harcourt (60%) and Jos (26.3%). No record of instrumental delivery or epidural analgesia was performed among the women studied. In view of economic cost of C/S with its morbidities and possible mortality, there is need for vaginal delivery especially assisted vaginal delivery with possible epidural analgesia to minimize pain. A similar study in Abakaliki also had a lower caesarean section rate of 21.6%.

Fetal distress and pre-term delivery were the commonest fetal complications seen in this study with each having an incidence of 50%. This brings to the fore the need for implementation of fetal surveillance in the third trimester.

This is much higher than the rate obtained in Jos where a 6.5% rate of fetal distress was recorded, 2 neonatal deaths (11.1%) were recorded with one stillbirth (5.6%). The study in Port Harcourt recorded 20% stillbirth whereas no still birth was recorded in

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Conclusion

This current study suggests that there are still high maternal and fetal complications in pregnant mothers with SCD in this environment. While a lot of advances have occurred in the management of SCD, there is need to ensure that these recent advances are utilized through a multidisciplinary management of the pregnant women with SCDS to ensure better outcome.

All the women did not receive preconception care and most did not commence antenatal care early enough. Women with SCD need to be educated on the need for preconception care to ensure optimization of health prior to conception. It is also important that pregnancy and conception discussions are held with women of child-bearing age with SCD during their periodic medical review.

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