1 <u>Case Study</u>

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3 Case Report: Successful Management of Opioid Abuse and Addiction in a

Known SCD Patient at the University of Calabar Teaching Hospital,

Calabar, Nigeria

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7 ABSTRACT

- 8 **BACKGROUND:** Opioids are a group of potent analgesic with mixed receptor activities.
- 9 Pain related symptom accounts for the main reason for substance dependence among sickle
- 10 cell disease (SCD) patients.
- 11 AIMS: The report aims to elucidate the adverse effects of opioid and it's complication
- 12 (abuse, dependency and addiction) and provides a management strategy for health
- practitioners to curtail the dependency of SCD patients to opioid use.
- 14 **PRESENTATION OF CASE:** The patient was a 27 years old lady that was diagnosed with
- sickle cell disease at the age of two. She presented with a two years history of oral self-
- medication of DF118 and Tramadol. She became dependent on the opioid on the account of
- sickle cell bone pain crises affecting both her upper and lower limbs with a pain score of
- 18 9/10. Other anagelsic like Diclofenac and Pentazocin couldn't ameliorate her excruciating
- pain but the administration of 60mg of oral DF118 provided her with quick relieve. The
- 20 sedative effect of Tramadol and DF118 allows her to sleep comfortably and hence the
- beginning of her dependency. On review, patient's system was essentially intact and she was
- 22 further referred for psychiatrist evaluation and possible rehabilitation.
- 23 **DISCUSSION:** Recurrent bone pain crisis represent the most common reason patient with
- SCD seek acute medical care. Due to the quick analgesic relief and euphoric effect derive
- 25 from both medication, patients feign pain after the genuine pain had subsided in other to
- 26 continue getting the prescription. The immediate pain assessment and frequent reassessment
- at 15min, 30min, 1hour then 2hours with appropriate application of medication until pain
- relief is very important to prevent drug abuse.
- 29 **CONCLUSION:** Less addictive analgesic should be considered first after observing the
- and nature of the pain before moving to stronger analgesic that has a high potential for abuse and
- 31 when stronger analysesic is to be used it should be for a short duration.
- 32 Keywords: Opioid Abuse, Sickle Cell Disease, Dependency, Addiction, UCTH

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INTRODUCTION

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Sickle cell disease (SCD) is a heterogeneous group of autosomal recessive structural 36 haemoglobin disorder¹. The most prevalent form is sickle cell anaemia (HbSS), which is the 37 homozygous form. Other forms of SCD include the compound heterozygous forms for 38 example, HbC in HbSC, βthalassaemia in HbSβ thalassaemia among others. 1,2 39 40 SCD is the most common genetic disorder in Sub-Sahara Africa. Nigeria is bearing a high disease burden with an estimated 1-2% of its population affected by the disease. An 41 estimated 20 – 30% of her populace carry the sickle cell gene with a normal haemoglobin 42 gene (carrier state). The disease burden differs slightly from one geographical region to 43 another. Nwogoh et al³reported the prevalence rate of SCD to be 2.4% and a 23% carrier state 44 in Benin City. Invama et al⁴ reported a prevalence of 3.7% in a multi-centre study in Nigeria. 45 The pathophysiology of Sickle cell anaemia is the substitution of the glutamic acid which is 46 hydrophilic with a less polar hydrophobic, neutral amino acid valine. During hypoxic 47 condition, this abnormal valine amino acid causes intraerythrocytic hydrophobic interaction 48 of affected haemoglobin tetramers, thereafter causing their precipitation and finally polymer 49 formation, leading to the loss of potassium and water resulting in cellular dehydration which 50 also promotes further precipitation and red cell rigidity⁵. Other contributing factors include 51 52 Nitric oxide depletion, endothelia activation with increased expression of adhesion molecule, inflammation and activation of coagulation.⁵ Despite understanding the molecular basis for 53 this disease the mechanism of vaso-occlussive crisis remain so vast that it cannot be 54 55 completely avoided thereby predisposing many of this patient to unbearable bone pain crisis. Opioid are group of potent analgesic with mixed receptor activities. Opioid is said to be 56 57 absorbed from the gastrointestinal tract and metabolized in the liver, gastrointestinal tract and kidney. There are four types of opioid receptors (Mu, Kappa, Delta, Nociceptor-OR) with a 58

major analgesic effect and a subtype nociceptor-OR which is termed the MOP.⁶ Most opioid tend to cause a reduction in consciousness and euphoria predisposing them to abuse.⁶

Recurrent bone pain crisis represent the most common reason patient with SCD seek medical care. In a study among sickle cell anaemia patients that are substance dependent, pain related symptom accounted for about 88% of all symptoms. Opioid analgesic are the mainstay of therapy for bone pain crisis in SCD, thus before adulthood most SCD patients must have had intermittent exposure to opioids. Opioids are potent analgesic associated with decrease hospitalization. The management of bone pain crisis has been an issue of debate among physician. Some physician advocate minimal use of these drug for fear of addiction, while others believe that inadequate analgesia predisposes patients to pseudoaddiction. There have been several report in substance abuse by SCD patients with a prevalence of less than 10% worldwide, but varies from one region to another in Nigeria. Ahmed et al reported a prevalence of 17.8% of opiate dependence among patients with SCD in Maiduguri, North East Nigeria with a male preponderance. Similarly, in a study by Mabayoje et al an incident of less than 10% was reported in the South West. Furthermore, Iheanacho et al la laso reported a less than 18.2% with male preponderance. From the various studies, it could be said that the incidence varies with geographical location with a prevalence of male sex

CASE REPORT

Miss EO is a 27 years old Nigerian Female graduate with sickle cell anaemia. It was diagnosed when she was 2 years old using Haemoglobin electrophoresis. She presented on the 4th of January 2017 with a 2 years history of self-medication of DF118 and Tramadol. She said she got addicted to these drugs about 2 years ago while she was admitted at government hospital in Calabar on account of sickle cell bone pain crisis affecting her upper and lower limb which was so severe with a pain score of 9/10(based on numerical pain rating scale) and

lasted for about 48hours despite administration of several analgesic such as Diclofenac and Pentazocin. Pain began to subside on administration of oral DF118 at 60mg to alternate with Tramadol 100mg which was given for a week. Patient said while she was on admission she enjoyed the feelings of the quick relief of the pain and sedative effect that allows her to sleep comfortably following the administration of DF118 and Tramadol. Patient on account of this improvement sought to know the name of the medication that could give such a wonderful relief and also because of fear of reoccurrence of the pain. She also noticed that both medication become drug of choice each time she has severe bone pain and present to the same health centre. She said on account of the psychological burden of the disease on her parents, who were worried of the repeated bone pain crisis with frequent hospital visits and was also discovered that both medication give their daughter relief and reduce their hospital visit, therefore decided to purchase a card of each medication weekly for her. Administering 30mg of DF118 twice daily initially but after 5 months increased the dosage to 60mg twice daily for a year because the initial dosage could not control the pain and she was very uncomfortable. She started with the new dose in the absence of pain because she was enjoying the euphoric effect. Patient revealed she was purchasing the drug on her own and even exaggerates her pain to get the drug prescription from her physician and at most time she gets it without prescription from a private pharmacy whose identity she does not want to disclose. Patient said she spends about \$\frac{1}{300}\$ (approximately \$1) to purchase a card, which she finances with her pocket money, selling her belonging, borrowing and buying on credit. Patient said after a year of self-medication of oral DF118 at 60mg twice daily she discovered she was not getting the relief she used to get. Patient said she got depressed and decided to change to another potent oral opioid (Tramadol) not the injectable because she reacts to the injectable, with nausea and continuous vomiting. She said she started with 50mg of Tramadol twice daily, got relief and also enjoyed the euphoric effect and later increased the dose to

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100mg then 200mg twice daily which she took every day for 1year even in the absence of pain. She gets the drug from a pharmacy and each card costs between №1700 - №2000 (approximately \$7). She also gets prescription from a doctor whose she refuse to mention the name or address. She also claims that anytime she tries to stop the medication she develops withdrawal symptoms which include lack of sleep, restlessness, sweating, dizziness, blurred vision, headache, joint pain and abdominal cramping, depression, agitation and craving for the drugs. Thus, these made her to seek help.

On examination, a young lady, pale, anicteric, conscious, alert and coherent, well oriented in person, time and place, well groomed with good motor function with intact long and short term memory, sense of judgement was mildly impaired. A review of her systems were essentially intact, patient was referred to the psychiatric team for further evaluation and possible detoxification and rehabilitation.

The following were the full blood count; PCV was 27%, Hb 9g/dl, WBC 11.2 x 10^9 /L, Neutrophiles 68%, Lymphocyte 32%, and Platelet 452 x 10^9 /L

DISCUSSION

This is a case study of a SCA patient who is dependent and addicted to tramadol and DF118.

BPC is the most common presentation among SCD,¹⁴ which our index patients suffers about 10-12 episodes annually necessitating her to seek treatment from a health practitioner who prescribed Tramadol and DF118. It was also noticed that the patient was on these prescribed drug (DF118 & Tramadol) for too long with prescription note not properly controlled, which

made her to have access to this prescription note. Due to the quick analgesic relief and euphoric effect derived from both medications, the patient has to feign pain after the genuine pain had subsided, in order to continue getting the prescription¹⁴. Based on this it is pertinent to say the patient is addicted to both drug and the primary aim of both drugs is now being abused. Lack of proper orientation and counselling of the parents of the patient also contributed to the abuse of the above medications. At this juncture, clinical expertise and judgement of the physician is highly needed to distinguish genuine pain from feigned pain in a patient with SCD with DF118 & Tramadol abuse. There is a paucity of information on DF118 and Tramadol abuse among sickle cell disease patient. Alao et al¹³ reported the case of a 38-year-old female sickle cell anaemia patient, though the drug of choice in this instance was cocaine.

The immediate pain assessment and frequent reassessment at 15min, 30min, 1hour then

2hours with appropriate application of medication until pain relief, are very important to prevent drug abuse. ^{15,16} Therefore the less addictive analgesic should be administered first after considering the nature of the pain before moving to stronger analgesic that have high potential for abuse and when stronger analgesic is to be used it should be for a short duration. ^{15,16}

The psychiatrist made an impression of opoid abuse and addiction in a known SCD patient.

Patient was initially managed on outpatient basis because patient had full insight of her problem and also has the desire to stop but does not wish to be admitted.

On mental state examination, patient was calm with good hygiene, cooperative and appears motivated and emotionally stable. Her perception was normal, thoughts well collected with normal cognition.

On physical examination, the patient was a young slim tall lady, afebrile, anicteric, acyanose with long limbs. The patient has gradually tapered off tramadol with a 50mg weekly reduction for about 6weeks until she suddenly developed an episode of bone pain crisis. She was then admitted for five weeks where she was treated with NSAID (Arthrotec) 75mg which was alternated with paracetamol 1000mg. Patient was also given diazepam 10mg and was carefully observed all through the period of admission with total avoidance of opoid, she was then discharged and placed on a routine medication of folic acid, paludrine and was given 2 weekly clinic appointment to ensure proper follow up. The patient was also counselled to adopt pain tolerance.

CONCLUSION

It is suggested that regular orientation of health worker on the use of opioid particular DF118 and Tramadol among opoid naive SCD, a careful objective assessment of sickle cell patient presenting with painful episodes should be carried out by an experienced health caregiver with each case taken on its own merit. A non-opioid analgesic should be commenced first and if an opioid should be used, it should be used for a short duration. Prescription note of opioid analgesic should be properly controlled; there should be a drug unit established and also legislation against sales of this controlled drug. Opioid addictive patient should be taught how to tolerate pains and referred to a psychiatrist for detoxification and rehabilitation.

Consent Disclaimer:

As per international standard or university standard written patient consent has been collected and preserved by the author(s).

Ethical Disclaimer: NA

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REFERENCES

- 179 1 Ashley-koch A, yang Q, onley RS. Sickle haemoglobin alleles and sickle cell disease Am
- 180 J Epidemol 2000; 151(9) 839-45.
- Davies SC, Oni L. Management of sickle disease. Br. Med. J. 1997; 315: 655-660.
- 182 3 Nwogoh B, Adewoyin AS, Iheanacho OE, and Bazuaye GN. Prevalence of haemoglobin
- variant in Benin City, Nigeria. Annals of Biomedical Science. 2012; 11(2):60-64.
- 184 4 Inyama M et al stroke prevalence among sickle cell disease patients in Nigeria a muti-
- centre study. Africa Health. 2014; 14(2): 446-452.
- 5 Fronticelli C, Gold R.(1976) conformational relevance of the beta6glu replaced val
- mutation in the beta subunit and in the Beta(1-55) and Beta(1-30)peptides of haemoglobin
- 188 S.J Biol Chem251:4968.
- 189 6 Daya P, Desmeules J, Collart L. Pharmacology of tramadol Drug 1997; 53 suppl 2:18-24
- 190 7 Elander J, Lusher J, Bevan D, Telfer P (2003). Pain management and symptoms of
- substance dependence among patients with sickle cell disease. SocSci Med. 57(9), 1683-
- 192 1696
- 193 8 Makanjuola A B, Olatunji P O(2009). Pentazocine abuse in sickle cell anaemia patients: A
- report of two case vignettes. African J Drug & Alcohol Studies. 8(2):59-64
- 195 9 Kotila T, Management of acute painful crisis in sickle cell disease clinical and laboratory
- haematology 2005 27(4)221
- 197 10 -223
- 198 11 Mabayoje V O, Adeyemo M A, Akinola N O. Case Review; Drug Addiction in sickle cell
- disease, A possible ongoing challenge in management of pain? Journal of global
- 200 bioscience 2015,4(4):2021-2025

- 201 12 Ahmed S G, Ibrahim U A, The prevalence of therapeutic opiate dependence among patient
- with sickle cell disease in Maiduguri, North East Nigeria. Nigerian journal of pharmacy,
- 203 2001,32:56-59
- 204 13 Iheanacho O E, Ezenwenyi I P, Enosolease M. G. Pentazocine abuse in sickle cell disease
- patient seen at a tertiary hospital in Nigeria: A chronic menace International journal of
- 206 tropical disease & health 2015,9(1):1-8
- 207 14 Alao AO1, Westmoreland N, Jindal S (2003) Drug addiction in sickle cell disease: case
- 208 report. Int J Psychiatry Med. ;33(1):97-101.
- 209 15 Okpala I, Add T. management of pain in sickle cell disorder. J Rsoc med.2002
- 210 sep;95(9):456-458
- 211 16 D. C. Rees, A. D. Olujohungbe, N. E. Parker, A. D. Stephens, P. Telfer, and J. Wright,
- "Guidelines for the management of the acute painful crisis in sickle cell disease," British
- Journal of Haematology, vol. 120, no. 5, pp. 744–752, 2003.
- 214 17 Ibidapo M O, Akinyanju O O. Acute illness in Nigeria children with SCA annals of
- 215 Tropical paediatrics 1987, 7:181-186