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Pre-presentation management, metabolic state and outcome of children admitted for diarrhoea disease in Calabar, Nigeria

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Abstract: *Background:* Much of the activities for the prevention and management of diarrhoea should occur or be initiated at home. In addition, management is guided by clinical and metabolic state. This work was designed to examine some pre-hospital management practices and the metabolic state of children with diarrhoea in Calabar, Nigeria.

Methodology: Prospective study of the pre-presentation management, clinical and metabolic states of children with diarrhoea presenting at University of Calabar Teaching Hospital, Calabar from April to November 2016. History was used to determine pre-presentation management while blood glucose and electrolytes were determined before initiation of treatment.

Results: Forty-four children aged 4 – 13years, mean 18.65 ± 17.37 months were recruited into the study. Forty (90.91%) had acute watery diarrhoea and 4 (9.09%) had dysentery. Of these, 72.27% had oral electrolyte solu-

tions at home and nine (20.45%) zinc. Twelve (27.27%) had mild, 26 (45.4%) moderate and 3 (6.82%) severe dehydration.

Twenty (63.64%) had metabolic acidosis, 18(40.91%) hyponatraemia and 18(40.9%) azotaemia suggesting Acute Kidney Injury. Two (4.50%) had hypoglycaemia while 16(36.30%) had hypozincaemia.

Conclusion: A majority of children with diarrhoea have oral electrolytes solution before presentation in the hospital and all had continued feeding. There is a high proportion with hyponatremia and more than a third had azotaemia. Modality of treatment at home did not seem to affect these parameters, perhaps because all subjects had continued feeding. Further studies on the appropriateness of low osmolarity ORS in this environment and renal function of children with diarrhoea are recommended.

Key Word: *Diarrhoea, Pre-hospital Management, electrolytes derangement*

Introduction

While there has been remarkable reduction in diarrhoea morbidity and mortality in recent years, it still causes more than half a million (525,000) under five childhood deaths annually.¹⁻⁴ When diarrhoea occurs good management with the use of appropriate oral rehydration fluids to prevent or correct dehydration, vitamin A, zinc, and continued feeding are essential to lessen morbidity and prevent mortality.³⁻⁶ Much of these should be done or initiated at home.⁵

Home management of diarrhoea can significantly affect ultimate outcome. Several studies in Africa and other parts of the developed have shown widely different modalities of pre-facility presentation management of diarrhoeal disease in children. In India Ahmed et al⁷ re-

ported in 2009 that only 8.7% of children received ORS for current episode of diarrhoea and 77.9% received antibiotics while 4.0% of parents withheld foods. Otho et al⁸ in Kenya reported in 2005 that 27.8% of the children had no fluids at all during diarrhoeal episodes, 70% of mothers decreased fluid intake and only 10.0% increased fluid intake. Almost 90% of the mothers withheld milk, including breast milk.⁸ On the other hand, in Nigeria, Uchendu et al⁹ in a hospital based study in 2011 reported that 73.1% of the children with diarrhoea had some form of oral rehydration fluid before presentation at hospital, out of which 80.7% had (standard) ORS and 94.8% continued feeds. A recent study in Port Harcourt, Nigeria showed that more than 70 percent presenting with diarrhoea at a cottage hospital had some form of oral rehydration fluid before presentation, with 97.4% of

these having standard ORS solution, with a majority (57.4%) having antimicrobials while more than a third had anti-emetics.¹⁰ Thus home/pre-presentation management of childhood appears to vary between and within country, with implications for in-facility management and health education activities.

This work was designed to examine pre-presentation management, metabolic state and outcome of children presenting with diarrhoea at the University of Calabar Teaching Hospital, Calabar, Nigeria.

Materials and Methods

This was a prospective study of all children admitted into the DTTU of the University of Calabar Teaching Hospital from April to November, 2016. A self-developed questionnaire with four sections covering socio-demographic characteristics of child and mother, pre-presentation management, clinical and metabolic states, and outcome, was used to collect data. The instrument was assessed for face validity with a test-re-test used to ascertain its reliability.

After informed consent of parent/care-giver, history and physical examination with attention to state of hydration was obtained. Three milliliters (3ml) of venous blood was drawn from each patient; 1.5ml each was added into a plain bottle and fluoride oxalate bottle. The specimens were taken to the Chemical Pathology Laboratory of the same hospital where they were centrifuged at 3000rpm for 10 minutes. The supernatant serum (from the plain bottle) for electrolytes and zinc analyses were stored at -20°C for a maximum period of two weeks before batch analysis. The supernatant plasma (from the fluoride oxalate bottle) was separated and immediately used for glucose analysis.

The plasma glucose was analyzed using the glucose oxidase photometric method produced by Biolabo® (Biolabo SA 02160, maicy France). The serum electrolytes were analyzed using an ion selective electrode (ISE) machine (ISE model unit – 910C, URIT Medical Electronic Co, Ltd, China). The serum zinc analysis was carried out using an atomic absorption spectrophotometer (AAS Model 205, Bulk Scientific, United States of America).

Data Analysis

Data was entered into an Excel spreadsheet then transferred to stata 10 (stata Corp Texas) for analysis. Frequencies, simple proportions and percentages were used to analyze the data. Chi-square test was used to test associations.

Ethical Issues

Ethical clearance for the study was obtained from the Health Research Ethics Committee of the University of Calabar Teaching Hospital, Calabar, with reference number: UCTH/HREC/33/337. Informed consent was

obtained from all participating parents/caregivers, with assurance that refusal to participate would not prejudice the care of their children/wards.

Results

Forty-four children were recruited into the study. The age range was 4 – 13 months with a mean age of 18.7 ± 17.4 months. Twenty four of the children were males and 20 females give a male to female ratio of 1:2:1.

The mean duration of diarrhea before presentation was 3.0 ± 2.8 with a range of 2 - .4 days. Forty (90.9%) of the children had acute watery diarrhoea while 4(9.1%) had dysentery. None had persistent diarrhoea. Twenty eight (63.6%) had associated vomiting. Twenty six (45.5%) of the subjects presented with moderate dehydration. Twelve (27.3%) had mild dehydration while 3 (6.8%) had severe dehydration. Three (6.8%) had no signs of dehydration.

Pre-presentation management

Fifteen (34.1%) of the children had exclusive breast feeding in the first six months of life. All the children had continued feeding at home during the episode of diarrhoea.

Fluids

Thirty-two (72.3%) of the children had oral fluids before presentation. Of these 6(13.6%) had low osmolarity Oral Rehydration Salt Solution (ORS), 17(38.6%) standard ORS and 9(20.5%) salt-sugar-solution.

Nine (20.5%) had zinc, 13(29.6%) had anti-microbials as follows; 8(18.2%) metronidazole; 2 (4.6%) cotrimoxazole; 2(4.6%) cephalosporin and 1(2.3%) amoxicillin/clavitanate. One (2.3%) each had metoclopramide and “Lucozade”

Clinical state

Table 1 shows the clinical state of the children in relation to modalities of pre-presentation management. Degree of hydration was not influenced by the administration of zinc nor by the type of oral rehydration fluid used.

Metabolic State

Table 2 shows the metabolic state of the children on admission. The important abnormalities were metabolic acidosis in 28 or 63.6% of the children, hyponatraemia (16 or 36.3%), raised creatinine (14 or 31.8%) hypokalaemia (6 or 13.6%). Two (4.6%) of the children had hypoglycaemia. Eighteen (40.9%), 95CI 26.3% – 56.8% of the children had azotaemia, (increased urea and/or creatinine).

Table 1: Levels of some metabolites in 44 children with Diarrhoea

Metabolite(mmol/L)	Range	Mean	SD	Low	Normal	Elevated
Zinc(0.06 – 1.20)	0.0 – 70	1.14	1.35	16[36.36]	15[34.09]	13[29.38]
Sodium(135 – 145)	120.7 – 146.0	135.22	6.23	18[40.91]	25[36.82]	1[2.27]
Potassium(3.5 – 5.5)	3.0 – 5.4	4.01	0.55	6[13.64]	38[86.36]	0[0.00]
Chloride(98 – 106)	840 – 106.1	98.25	5.98	15[34.09]	27[61.36]	2[4.5]
Bicarbonate (2.00 – 26.0) –	18.1 – 26.0	21.46	2.35	28[63.64]	16[36.36]	0[0.00]
Urea (1.8 – 6.4)	1.8 – 7.8	4.46	1.69	3[6.82]	39[77.27]	7[15.91]
Creatinine (18.0 – 35.0)	18 – 46.2	23.39	8.95	0[0.00]	30[68.18]	14[31.82]
Glucose (3.6 – 5.8)	1.8 – 7.9	4.72	1.24	2[4.55]	42[95.45]	0[0.00]

Key: () = Reference values

() = Percentage

Table 2: Home Treatment and Clinical State on Presentation

Home Treatment	No dehydration	Mild dehydration	Moderate dehydration	Severe dehydration	X ²	P
No Zinc	2	4	15	2	0.39	0.94
Zinc	1	4	11	1		
SSS	1	1	6	1		
Low osmololomity ORS	0	2	2	4	2.93	0.18
Standard ORS	1	5	9	2		

Table 3 shows the metabolic state in relation to type of pre-presentation oral rehydration solution used. Type of home fluid did not affect the distribution of electrolyte changes.

Outcome

Four (9.1%) children had intravenous fluids initially and later low osmolarity ORS. Forty (90.9%) had low osmolarity ORS till discharge home. Median duration of stay in the hospital was 24hours (IQR 1- 48hours). There was zero case fatality.

Table 3: Home Management and Metabolic State

	SSS	LOORS	SORS	X ²	P
Hyponatraemia	4	2	8	0.03	0.98
Hypokalaemia	3	0	2	2.89	0.24
Hypochloraemia	3	0	6	1.99	0.37
Azotaemia	4	1	5	0.74	0.69

Key

SSS: Salt sugar solution

LOORS: Low osmolarity ORS

SORS: Standard ORS

Discussion

The age distribution of the children was expected and is in keeping with the known epidemiology of the diarrhoea disease. The slight male preponderance is also typical, so also the preponderance of acute watery diarrhoea.^{1-3,5} Appropriate home management of diarrhoea, including appropriate oral dehydration fluid, continuing feeding, zinc therapy, are important modalities of reducing morbidity and mortality from diarrhoea disease.^{4,5}

The low exclusive breastfeeding rate among these children is in keeping with the situation in Nigeria, reported to be the poorest in the world.⁷⁻⁹ Gratifyingly, all the subjects continued feeding during the episodes of diarrhoea. This is different from the situation in the past⁴ and reflects some gains in the health education on the management of diarrhoea disease at home. More than seventy percent of the children had some form of oral fluid, though the correctness of the constitution was not assessed. This is not satisfactory as all children with diarrhoea should have extra fluids as soon as the diarrhoea starts.⁴ All the children had continued feeding during the episodes of diarrhea. This is most salutary and is much different from the report in India in 2009 where up to four percent had no feeds during diarrhoeal episodes⁷ are in Kenya where about 90% of the children had milk, or including breast milk withheld in a 2005 report.⁸ This may reflect the high level of maternal education in Cross River state of Nigeria which has adult English language literacy rate for both sexes of more than seventy six percent,¹¹ and similar to report in the nearby Enugu state in Nigeria with adult English literacy rate of more than sixty four percent,¹¹ and more than seventy three percent of children with diarrhea were offered oral rehydration fluid at home.⁹ Maternal education is known to affect both the incidence and home management of childhood diarrhoea.^{12,13}

Only a fifth of the children had zinc. This reflects weak knowledge of this relatively recent recommendation by the pre-presentation care-givers. Zinc supplementation has been demonstrated to reduce the duration of acute diarrhoea; and for persistent diarrhoea duration and the probability of treatment failure or death.¹⁴ Zinc supplements for 10-14 days during diarrhoeal episode has also been shown to reduce further occurrences in the subsequent 2-3 months¹⁴ and is currently recommended as a routine.¹⁵ Mechanism of action of zinc is not fully understood but includes improvement of water and electrolytes absorption, regeneration of intestinal epithelium, increase in the level of brush boarder enzymes and enhancement of immune response against the diarrhoea pathogens.¹⁴ The rate of antimicrobial use in the present study was 18.2% though only 4(9.1%) had dysentery. This is a lot better than the situation in Kashmir, India where the rate of antibiotic use was close to 78%⁷ and in Port Harcourt, Nigeria where nearly sixty percent had antimicrobials.¹⁰ Only one child had anti-emetic in the current study. This again may reflect the high maternal literacy rate in the area and is markedly different from the report from Port Harcourt where more than a third of the children had antiemetic is¹⁰ with dire consequences

for some.^{10,16}

While only nine percent had dysentery, thirteen percent had antimicrobials. Abuse of anti-microbials and anti-emetics in children with diarrhoea is a well-known phenomenon in Nigeria.¹⁰ Fortunately only one child had an anti-emetic.

The low incidence (4.5%) of hypoglycaemia in these children is almost identical to the 4.0% recorded in the same unit a few years earlier¹⁷ and is much lower than the 11.0 percent reported by Huq et al¹⁸ in Bangladesh and 7.7 percent by Onyiriuka et al¹⁹ in Benin city, Nigeria and the 5.3% in Lagos, Nigeria by Oyenusi et al.²⁰ This may reflect the continued feeding of all the children in the current study. The single child with hypoglycaemia had an anti-emetic. Anti-emetics have been associated with hypoglycaemia in childhood diarrhoea.¹⁶ Perhaps children with diarrhoea and history of initiation of anti-emetics, which interferes with feeding, should be screened for hypoglycaemia.

Forty percent of the children were hyponatraemic. However, their electrolyte levels were not re-assessed after rehydration. It would be needful to assess the electrolyte and osmolarity changes of children in response to low osmolarity ORS in this environment in view of the high proportion of them with hyponatraemia. While low osmolarity ORS has been demonstrated to be associated with less vomiting and lower stool volume in children admitted with diarrhoea and is currently the recommended fluid by WHO,²¹ it has also been associated with increased incidence of transient asymptomatic hyponatraemia.²² A little above a tenth of the children had metabolic acidosis which is eminently correctable by standard or low osmolarity ORS.^{4,5} The modality of pre-presentation management did not appear to affect the distribution of electrolyte changes in these children. This may reflect the effect of continued feeding during the diarrhoeal episodes and should be encouraged. The large proportion (more than a third) of the children with hypozinaemia in this study indicates that zinc deficiency and diarrhoea may be components of a vicious cycle. Diarrhoea management is therefore a useful entry point for the management of the widespread zinc deficiency in children reported in this environment.^{23,24} Medicine vendors/patient medicine dealers play important roles in the management of childhood diarrhoeal disease in Nigeria.^{25,26} These should receive targeted education on the use of zinc and other aspects of diarrhoeal management.

The relatively high incidence of azotaemia in this study, probably occurring as a result of pre-renal AKI, is noteworthy. Reversible acute kidney injury occurring in hospital has been shown to be associated with a significant risk for *de novo* chronic kidney disease, with implications for long term follow up of such patients.²⁷ Acute Kidney Injury and Chronic Kidney Disease (CKD), have been considered a continuous spectrum with vascular insufficiency, cell-cycle disruption and maladaptive repair mechanism as some of the modulators of progression from AKI to CKD.²⁸ This warrants long-term follow-up of patients with first episodes of AKI, even if they presented with normal renal function.²⁸ Since AKI

and CKD are currently considered inter-connected syndromes,^{29,30} it would be desirable to investigate the incidence of AKI in children with diarrhoea and establish the need and modalities to follow up these children.

Diarrhoea may be playing important role in the pathogenesis of CKD in this environment.

Conclusion

Acute watery diarrhoea is the main type of diarrhoea presenting in this centre, accounting for more than ninety percent of the cases. Most of the children had moderate dehydration. EBF rate is low among these children and all had continued feeding during diarrhoea while almost three quarters had some form of oral rehydration fluids before presentation. There was significant rate of abuse of anti-microbials. A large proportion were hyponatraemic on admission, indicating a need to investigate the appropriateness of low osmolarity ORS in children with diarrhoea in this environment. More than a third were zinc deficient. There is a high incidence of azotaemia suggesting AKI in these children. This has implications for the future development of CKD and the need for long term follow-up of these children. Larger studies to evaluate the suitability of low osmolarity ORS in this environment AKI, in diarrhoea are warranted. Health education on home management of diarrhoea in Calabar should emphasize zinc supplementation. This study has some obvious limitations. The sample size is small and details of the mixing of ORS solutions were not obtained. All the same, it has made important observations and raised important questions.

Conflict of Interest: None

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