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Article

Assessment of Determinants of Paediatric Diarrhoea Case Management Adherence in Pakistan

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Abstract: Worldwide, diarrhoea in children under-five years of age is the second leading cause of death. Despite having high morbidity and mortality, diarrhoeal diseases can be averted by simple and cost-effective interventions. The Integrated Management of Childhood Illness (IMCI) has proposed the use of Oral Rehydration Salt (ORS) and zinc together with adequate food and fluid intake for the management of acute non-dysenteric watery diarrhoea in children. In the past, few studies examined the determinants of adherence to diarrhoea case management. Therefore, this study measured the determinants of therapeutic and dietary adherence to diarrhoea case management using the third and fourth wave of Pakistan Demographics & Health Surveys (PDHS) datasets. Data from 4,068 children between 0 to 59.9 months with positive history of diarrhoea were included, while data on children with dysentery, severe dehydration, and co-morbid condition was excluded. This study reported therapeutic adherence in less than 10% of children in Pakistan, while dietary adherence was reported in 39.2% of children (37.7% in 2012-2013 ~ 40.7% in 2017-2018). A significant improvement in therapeutic (0.8% in 2012-2013 ~ 8.1% in 2017-2018) and dietary adherence (37.7% in 2012-2013 ~ 40.7% in 2017-2018) was reported in the 2017-2018 survey, compared to the 2012-2013 survey. In general, children over the age of one year (compared to children <1 year) and of the richer/richest socioeconomic class (compared to poorest/poorer socioeconomic class) have showed higher therapeutic and dietary adherence. Therapeutic and dietary adherence among diarrhoeal children can be improved by increasing the awareness and accessibility of ORS, zinc, and essential foods.

Keywords: adherence; case management; determinants; diarrhoea; paediatrics

Introduction:

Children under five years of age are highly vulnerable to various infections (1). Pneumonia, diarrhoea, malaria, tuberculosis, and Human Immunodeficiency Virus (HIV) are the five common but preventable infectious diseases known to kill millions of children before their fifth birthday (2, 3). Among various paediatric illnesses, diarrhoea is the second leading cause of death after pneumonia in children below five years (4, 5). The disease is characterized by the passage of three or more stools of a loose consistency on any given day, which leads to fluid and electrolyte loss. If not managed promptly, fluid and electrolyte loss during the diarrhoea results in hypovolemic shock and even death (5, 6).

Despite having high mortality associated with diarrhoea, the disease can be averted by simple and cost-effective interventions (7). To prevent health adversities associated with diarrhoea, it is necessary to break the transmission cycle of the common pathogens causing diarrhoea (5). This can be executed by providing health education, immunization with the rotavirus vaccine, exclusive breastfeeding for children under six months, access to safe drinking water, and improved sanitation, such as safe excreta disposal, washing hands with soap, and personal & food hygiene (5, 8, 9). However, medical cases of paediatric diarrhoea can be managed by using either oral and/or intravenous fluids (10). The Integrated Management of Childhood Illness (IMCI) guideline has recommended the use of Oral Rehydration Solution (ORS), zinc, and adequate use of both fluid and food for the management of non-dysenteric diarrhoea with none/some dehydration (7).

ORS was recommended among non-dysenteric diarrhoeal children with none/some dehydration over fifty years ago in 1968 (11). The ORS is a mixture of glucose and different salts, such as sodium chloride, potassium chloride, and sodium citrate. When mixed with water, this mixture of glucose with different salt can replenish the necessary electrolytes and water loss in faeces during diarrhoea (12). In addition to ORS, the World Health Organization (WHO) and UNICEF in 2004 advised using oral zinc because zinc reduces the severity and frequency of diarrhoeal episodes among children, reducing the complications and mortality associated with diarrhoea (11, 13). The global adherence to the guidelines set out by the WHO and UNICEF is still inadequately adhered to on a large scale (7). In 2012 less than fifty percent of children with diarrhoea received the recommended treatment proposed by the WHO and UNICEF in the IMCI guidelines (7, 11, 13, 14).

Paediatric diarrhoea case management approach in Pakistan is not different from other countries of the World. Most of the studies conducted previously have individually examined the use of ORS and zinc, but a few studies have only assessed the adherence to the co-administration of zinc with ORS. To the best of our knowledge, no studies in Pakistan have thoroughly examined the adherence to the co-administration of zinc with ORS at a large scale. Moreover, adherence to dietary practices has not been investigated in most countries. To investigate the treatment and dietary adherence to the recommended WHO and IMCI guidelines, this study was conducted among children below five years of age using the third and fourth wave of Pakistan Demographics & Health Surveys (PDHS) datasets.

2. Methods

Study design: This study was based on the secondary analysis of Pakistan Demographic & Health Surveys (PDHS). This study included data from the third and fourth waves of PDHS conducted in 2012-2013 and 2017-2018. In each PDHS, the data was collected by a team of data collectors, who received adequate training of around three to four weeks related to data collection and interviewing technique. The data collectors in each PDHS interviewed women of reproductive age by means of a survey questionnaire, and there is no predefined exposure and outcome measurement. Owing to this reason, each PDHS has a cross-sectional study design.

Study population: The study included data from all children aged 0 to 59.9 months with positive diarrhoea history. However, data on all children with dysentery (blood in stool), severe dehydration (received intravenous treatment), and enteric disease (diarrhoea with fever) were excluded. Additionally, data on children simultaneously suffering from diarrhoea with cold, cough, flu, shortness of breath, and malaria, were also excluded.

Sample size and sampling methods: The primary sampling unit (PSU) in each Demographic & Health Survey (DHS) were women aged between 15 to 49 years, which were approached via a multistage stratified cluster sampling method for the interview. Initially, the target population of this study was bifurcated into groups based on their place of residence: urban and rural. Enumeration blocks (EBs) were selected randomly from each urban and rural group. In general, each EBs represents a cluster of 200 to 250 households. Finally, from each EBs, women were selected following a systematic random method. The National Institute of Population Sciences (NIPS) instructed the data collection team to select women from every 28th household for the survey. However, the sample size in each PDHS was calculated using the number of EBs. The list of the EBs in each PDHS was received from the Pakistan Bureau of Statistics and the last census records. During the

third wave of PDHS, there were 500 EBs, which increased to 580 during the fourth PDHS wave conducted in 2017-2018. The. The number of EB in 2012-2013 was 500, which increased to 580 in 2017-2018. The survey team multiplied the number of EB with the fixed number "28" and thus produced a sample size of 14,000 and 16,240 (15, 16).

In this study, children aged below five years with a history of diarrhoea in the last fourteen days were targeted. Data from all the children with dysentery, severe dehydration, enteric disease, and other comorbid condition were excluded. Similarly, data of all the children with normal health status, i.e., no episode of diarrhoea and/or any other common preventable illnesses, i.e., cold, cough, flu, fever, was also excluded. After excluding the data of ineligible children in this study, we received a total sample size of 5,105 (2,998 from 2012-2013 and 2,107 from 2017-2018).

Study outcome: The primary outcome of this study is to assess adherence to diarrhoea case management practices in children. The WHO has recommended using ORS, oral zinc sulphate, and adequate fluid & food to manage diarrhoea in children with no and/or some dehydration. The adherence to recommended diarrhoea case management was derived from the consumption of ORS, zinc, adequate fluid, and food intake during the diarrhoeal episodes. In this study, adherence to diarrhoea case management was examined in two steps: 1) Therapeutic adherence and 2) Dietary adherence. Further information about therapeutic adherence and dietary adherence pertaining to diarrhoea case management is presented in Table-1.

Table 1. Operational definition of therapeutic and dietary adherence pertaining to paediatric diarrhoea case-management.

	Adherence	Non-adherence			
Therapeutic	Coadministration of ORS and	The therapeutic non-adherence			
•	zinc for the management of	may encompass treatment with any			
	acute watery non-dysenteric	of the following medicine/remedies			
	diarrhoea with none/some	either without ORS or Zinc or both:			
	dehydration.	1) Home solution			
		2) Antibiotic pills			
		3) Antibiotic syrup			
		4) Antimotility drug			
		5) Antibiotic injection			
		6) Intravenous fluid			
		7) Any unknown pill			
		8) Non-antibiotic injection			
		9) Unknown injection			
		10) Rice starch			
		11) Mint extract			
		12) Herbal remedy			
		13) Other treatment			
		14) No treatment.			
Dietary adherence	Adequate intake ¹ of both food	Inadequate intake ² of either fluid or			
	and fluid during the diarrhoeal	food or both during the diarrheoal			
	illness.	illness.			

1. Adequate intake of food and fluid= Either equal or greater intake of food/fluid during the diarrhoeal illness than does the routine

food/fluid intake.

2. Inadequate intake of food and fluid= The intake of food/fluid during the diarrhoeal illness should be lower than does the routine

food/fluid intake. Food abstinence and fluid restrictions are examples of fluid/food inadequate intake.

Study covariates: Different covariates, such as child age (0 to 11 months, 12 to 23 months, 24 to 35 months, 36 to 47 months, and 48 to 59 months), gender (male and female), birth order (first and other), birth type (singlet and twin/triplet) to community factors such as place of residence (urban and rural) were selected. Additionally, maternal factors such as maternal age (less than 20, 20 to 34, and 35 plus), maternal education (none, primary, and secondary/higher), maternal occupation (working and non-working), number of parities (two or less and more than two), and birth in last five years (two or less and more than two) were also included. Similarly, household factors, such as time to get water (within the premises and obtained from outside), family size (four or less and more than four), number of children under five years (two or less than two and more than two), wealth index (poorest, poorer, middle, richer, and richest) were also included.

Data analysis: In this study, two different datasets were obtained from the DHS data repository. After variable selection, the datasets used in this study were merged into a single file. Before formal data analysis initially, preliminary analysis for each variable was performed before formal data analysis to assess the data quality and distribution. Following preliminary analysis, descriptive analysis was carried out. Later inferential analysis was carried out. The outcome variable of this study was adherence to diarrhoea case management practices, which has two categories: Yes and No. Thus, this study used binomial logistic regression to assess adherence to diarrhoea case management practices with different covariates. Variables with a p-value of ≤0.05 and a confidence interval of 95% were considered significant predictors.

Ethical principles of study: The data of this study was received from the DHS data repository. After receiving from the DHS data repository, the data were screened thoroughly. The data files were completely devoid of all the information, including any patient identifier, such as participant name, residential address, national identity card number, passport number, and phone number. The ethical principles of anonymity and confidentiality were maintained throughout this research.

3. Results

Description of the study population

This study analyzed data from 4,068 children aged 0 months to 59.9 months using datasets of Pakistan Demographic and Health Surveys conducted in 2012-2013 and 2017-2018, respectively. The detail regarding the sociodemographic profile of the study population is presented in Table 1.

More than 40% of children between 0 to 59.9 months received ORS during diarrhoeal illness, but less than a quarter of children received zinc. In 2012-2013, only 2% of the children received zinc, which increased to 12.5% in 2017-2018. Owing to this reason, the 2017-2018 survey reported an increase in treatment adherence to 8.1% from 0.8% in 2012-2013.

Across two survey period, an adequate intake of fluid was reported in around 60% of the children. However, less than half of the children reported an adequate intake of food. Overall, 39.2% of children showed dietary adherence during their diarrhoeal illness (Table 2)

Table 2. Sociodemographic features of the study population.

Variable	2012-2013	2017-2018	Total	
	(n = 2002)	(n = 2066)	(n = 4068)	
	Child facto	rs		
Child age (categorical)				
0 to 11.9 months	506 (12.4%)	577 (14.2%)	1083 (26.6%)	
12 to 23.9 months	554 (13.6%)	567 (13.9%)	1121 (27.6%)	
24 to 35.9 months	415 (10.2%)	423 (10.4%)	838 (20.6%)	
36 to 47.9 months	312 (7.7%)	289 (7.1%)	601 (14.8%)	
48 to 59.9 months	215 (5.3%)	210 (5.2%)	425 (10.4%)	
Child sex				
Male	1065 (26.2%)	1093 (26.9%)	2158 (53%)	
Female	937 (23.0%)	973 (23.9%)	1910 (47%)	
Birth order	3.47 ± 2.73		3.36 ± 2.28	
Birth order (categorical)				
Primipara	471 (23.5%)	523 (25.3%)	994 (24.5%)	
Multipara	1531 (76.5%)	1543 (74.7%)	3074 (75.5%)	
Birth type				
Singlet	1958 (977.8%)	2016 (97.5%)	3974 (97.7%)	
Twin/triplet	44 (2.2%)	50 (2.5%)	94 (2.3%)	
	Maternal fac	tors		
Maternal age				
15 to 19 years	68 (3.4%)	86 (4.2%)	154 (3.8%)	
20 to 24 years	490 (24.5%)	479 (23.2%)	969 (23.8%)	
25 to 29 years	626 (31.3%)	660 (31.9%)	1286 (31.6%)	
30 to 34 years	454 (22.7%)	468 (22.7%)	922 (22.7%)	
35 to 39 years	256 (12.8%)	275 (13.3%)	531 (13.1%)	
40 to 44 years	86 (4.3%)	77 (3.7%)	163(4.0%)	
45 to 49 years	22 (1.1%)	21 (1%)	43 (1.1%)	
Maternal education				
None	1116 (55.7%)	1007 (48.7%)	2123 (52.2%)	
Primary	302 (15.1%)	318 (15.4%)	620 (15.2%)	
Secondary	380 (19.0%)	456 (22.1%)	836 (20.6%)	
Higher	204 (10.2%)	285 (13.8%)	489 (12.0%)	
Maternal work				
No	1602 (76.9%)	1857 (89.9%)	3459 (85%)	
Yes	400 (23.1%)	209 (10.1%)	609 (15%)	
Number of parities	3.73 ± 2.38	3.50 ± 2.20	3.61 ± 2.30	
Birth in last 5 years	1.80 ± 0.74	1.74 ± 0.71	1.77 ± 0.72	
ž.	Household fa	ctors		
Family size	9.43 ± 5.02	9.52 ± 5.04	9.47 ± 5.03	
Wealth index				
Poorest	446 (22.3%)	432 (21.7%)	894 (22%)	
Poorer	443 (22.1%)	482 (23.3%)	925 (22.7%)	
Middle	392 (19.6%)	448 (20.9%)	824 (20.3%)	
Richer	400 (20.0%)	366 (17.7%)	766 (18.8%)	
Richest	321 (16.0%)	338 (16.4%)	659 (16.2%)	
	Treatment fac		/-	
Use of ORS	,			
No	1148 (57.3%)	1213 (58.7%)	2361 (58%)	

Use of zinc			
No	1962 (98%)	1807 (87.4%)	4028 (92.7%)
Yes	40 (2%)	259 (12.5%)	299 (7.3%)
Treatment adherence			
No adherence	1984 (99.2%)	1882 (93.9%)	3866 (95%)
Adherence	18 (0.8%)	184 (8.1%)	202 (5%)
Use of fluid intake			
None or less than usual	821 (41%)	818 (39.5%)	1639 (40.3%)
More or same as usual	1181 (59%)	1248 (60.5%)	2429 (59.7%)
Food and diet intake			
None or less than usual	1175 (58.6%)	1110 (53.7%)	2285 (56.2%)
More or same as usual	956 (41.3%)	956 (42.3%)	1783 (43.8%)
Dietary adherence			
No adherence	1247 (62.3%)	1225 (59.3%)	2472 (60.8%)
Adherence	755 (37.7%)	841 (40.7%)	1596 (39.2%)
	Community fo	actors	
Place of residence			
Urban	829 (41.4%)	917 (44.3%)	1746 (42.9%)
Rural 1173 (58.6%)		1149 (55.7%) 2322 (

Determinants of therapeutic adherence during diarrhoea case management

Our study reported a significant improvement in the therapeutic adherence of paediatric diarrhoea case management across two survey periods. Similarly, zinc intake has improved significantly in the 2017-2018 survey, compared with the 2012-2013 survey. However, the intake of ORS didn't show any significant change across the two survey periods.

In general, the therapeutic adherence to paediatric diarrhoea case management significantly increased in children aged over 12 months compared to children aged between 0 to 11.9 months. Similarly, the intake of ORS and zinc was significantly higher in children over 12 months compared to children below 12 months.

Compared to children of the poorest socioeconomic class, children of the richest socioeconomic class showed 1.56 (1.01 to 243) higher odds of therapeutic adherence. However, socioeconomic class did not show any association with the use of either ORS or zinc during diarrhoea (Table 3).

Additionally, increased maternal education, increased family size, and less number of childbirth in the last five years significantly increased the intake of ORS during diarrhoea.

Determinants of dietary adherence during diarrhoea case management

Compared to the 2012-2013 survey, the 2017-2018 survey showed a significant improvement in dietary adherence during diarrhoea case management. Similarly, food intake during the diarrhoea case management increased to 1.26 (1.11 to 1.43) in the 2017-2018 survey, compared to the 2012-2013 survey. However, no association was observed between the intake of fluid and survey years.

In general, an improvement in the socioeconomic status from poorer to richest significantly increased the intake of fluid, intake of food, and both (dietary adherence), compared to the poorest children. Similarly, an increase in child age from 12 months to 59.9 significantly increases food intake and dietary adherence to around 1.5 to 2 folds. Additionally, dietary adherence during the diarrhoea case management decreased by 2% (95% CI: 1% to 3%) with each member's increased family size (Table 4).

Table 3. Assessing the determinants of Diarrhoea treatment adherence in children below five years of age.

	Intake of ORS ¹		Intake of Zinc ²		Treatment adherence $^{3\alpha}$	
Variable	Unadjusted odd (95% CI)	Adjusted odds (95% CI)	Unadjusted odd (95% CI)	Adjusted odds (95% CI)	Unadjusted odd (95% CI)	Adjusted odds (95% CI)
Year	D (D (D. C	ъ. с	D. C
2012-2013	Ref		Ref	Ref	Ref	Ref
2017-2018	0.94 (0.83 to		7.03 (50.1 to	7.14 (5.08 to	10.77 (6.61 to	10.97 (6.73 to
	1.07)		9.86) *	10.03) *	17.55) *	17.89) *
	Ref	Ref	Ref	Ref		
Child age	1.94 (1.63 to	1.95 (1.64 to	1.69 (1.21 to	1.80 (1.28 to	Ref	Ref
(categorical)	2.31) *	2.32) *	2.36) *	2.54) *	1.96 (1.28 to 2.98)	
0 to 11.9 months	1.74 (1.44 to	1.78 (1.47 to	1.48 (1.03 to	1.57 (1.08 to	*	*
<i>12 to 23.9 months</i>	2.09) *	2.15) *	2.14) *	2.28) *	1.87 (1.19 to 2.93)	2.02 (1.34 to 3.17)
$24\ to\ 35.9\ months$	1.57 (1.27 to	1.58 (1.28 to	1.02 (0.70 to	1.19 (0.76 to	*	*
$36\ to\ 47.9\ months$	1.92) *	1.94) *	1.63)	1.13 (0.76 to	1.33 (0.79 to 2.26)	1.49 (0.87 to 2.56)
$48\ to\ 59.9\ months$			1.98 (1.31 to		2.17 (1.30 to 3.63)	2.43 (1.44 to 4.12)
	1.29 (1.02 to	1.31 (1.04 to		2.18 (1.43 to	*	*
	1.63) *	1.66) *	3.01) *	3.32) *		
Child sex	Ref		Ref		ъ (
Male	0.97 (0.85 to		1.10 (0.86 to		Ref	
Female	1.09)		1.39)		1.00 (0.75 to 1.33)	
	0.98 (0.96 to		0.97 (0.92 to			
Birth order	1.01)		1.02)		0.98 (0.92 to 1.05)	
Birth order						
(categorical)	Ref		Ref		Dof	
Primipara	0.95 (0.82 to		0.94 (0.72 to		Ref	
Multipara	1.10)		1.23)		0.92 (0.67 to 1.28)	
Birth type	D. C		D. C			
Singlet	Ref		Ref		Ref	
Twin/triplet	0.81 (0.53 to		0.41 (0.12 to		0.65 (0.19 to 1.99)	
	1.24)		1.30)			
	Ref		Ref			
	1.23 (0.86 to		0.91 (0.47 to			
Maternal age	1.76)		1.71)			
15 to 19 years	1.30 (0.91 to		0.84 (0.45 to		Ref	
20 to 24 years	1.84)		1.59)		1.09 (0.48 to 2.46)	
25 to 29 years	1.48 (1.03 to		0.95 (0.50 to		0.93 (0.41 to 2.09)	
30 to 34 years	2.11) *		1.81)		1.10 (0.48 to 2.48)	
35 to 39 years	1.53 (1.06 to		1.01) 1.14 (0.59 to		1.39 (0.60 to 3.21)	
40 to 44 years	2.23) *		2.22)		1.66 (0.63 to 4.35)	
v	2.23) * 1.19 (0.75 to		2.22) 1.19 (0.54 to			
45 to 49 years					0.50 (0.05 to 4.17)	
	1.89)		2.65)			
	1.46 (0.73 to 2.91)		0.28 (0.03 to 2.23)			
Maternal	2.01/		2.20/			
education	Ref	Ref	Ref		Ref	
None	0.98 (0.81 to	0.95 (0.79 to	1.28 (0.91 to		1.18 (0.78 to 1.81)	
Primary	1.18)	1.15)	1.79)		1.25 (0.86 to 1.81)	

Secondary	1.44 (1.23 to	1.36 (1.15 to	1.25 (0.92 to	1.84 (1.24 to 2.74)
Higher	1.70) *	1.61) *	1.69)	*
	1.41 (1.16 to	1.28 (1.03 to	1.55 (1.09 to	
	1.72) *	1.58) *	2.19) *	
Maternal work				
No	Ref		Ref	Ref
Yes	0.89 (0.75 to		0.73 (0.51 to	0.90 (0.61 to 1.36)
	1.07)	-	1.06)	
Number of	0.98 (0.95 to		0.97 (0.92 to	0.98 (0.92 to 1.05)
<u>parities</u>	1.01)		1.02)	0.50 (0.52 to 1.05)
Birth in last 5	0.87 (0.80 to	0.89 (0.81 to	0.91 (0.77 to	0.83 (0.68 to 1.02)
years	0.95) *	0.97) *	1.07)	0.03 (0.00 to 1.02)
Family size	1.01 (1.00 to	1.02 (1.01 to	1.00 (0.97 to	0.99 (0.96 to 1.02)
Tumity Size	1.02) *	1.03) *	1.02)	0.99 (0.90 to 1.02)
	D. C		D (
TAT 1:1 · 1	Ref		Ref	D. C
Wealth index	1.18 (0.97 to		1.05 (0.73 to	Ref Ref
Poorest	1.43)		1.51)	1.03 (0.64 to 1.59)
Poorer	1.35 (1.11 to		1.07 (0.73 to	1.01 (0.64 to 1.57)
Middle	1.64) *		1.55)	0.79 (0.49 to 1.29) 0.95 (0.60 to 1.51)
Richer	1.32 (1.08 to		1.01 (0.68 to	1.55 (1.01 to 2.39) 0.84 (0.51 to 1.38)
Richest	1.61) *		1.48)	* 1.56 (1.01 to 2.43)
	1.74 (1.41 to		1.46 (1.01 to	*
	2.13) *		2.12) *	
Place of residence		D (5 (
Urban	Ref	Ref	Ref	Ref
Rural	0.71 (0.63 to	0.79 (0.69 to	0.86 (0.68 to	0.89 (0.67 to 1.18)
Киги	0.81) *	0.91) *	1.10)	0.00 (0.0. 10 1.10)

Where, * = A significant association with various study covariates having p-value \leq 0.05, 1= Odds of ORS intake was adjusted with child age, maternal education, birth in last five years, family size, and place of residence, 2 = Odds of zinc intake was adjusted with year of survey, and child age, 3 = Odds of treatment adherence was adjusted with year of survey, child age, and wealth index, α = treatment adherence is the sum of adherence to the intake of both ORS and zinc during the diarrhoeal illness.

Table 4. Dietary adherence during diarrhoea case management among children below five years of age.

	Adequate fluid intake		Adequate food intake		Dietary adherence	
Variable	Unadjusted odd (95% CI)	Adjusted odds (95% CI)	Unadjusted odd (95% CI)	Adjusted odds (95% CI)	Unadjusted odd (95% CI)	Adjusted odds (95% CI)
Year	D - C		D -C	D - C	D of	D a C
2012-2013	Ref		Ref	Ref	Ref	Ref
2017-2018	1.06 (0.93 to		1.22 (1.08 to	1.26 (1.11 to	1.13 (1.01 to 1.28)	1.16 (1.02 to
	1.20)		1.35) *	1.43) *		1.32) *
a	Ref		Ref	Ref	Ref	Ref
Child age	0.91 (0.76 to		1.83 (1.54 to	1.86 (1.56 to	1.71 (1.41 to 2.07)	1.78 (1.49 to
(categorical)	1.08)		2.18) *	2.21) *	*	2.13) *
0 to 11.9 months	0.89 (0.74 to		1.74 (1.44 to	1.78 (1.48 to	1.77 (1.41 to 2.07)	1.74 (1.44 to
12 to 23.9 months	1.07)		2.09) *	2.15) *	*	2.11) *
24 to 35.9 months	0.86 (0.71 to		2.15 (1.75 to	2.22 (1.81 to	2.00 (1.62 to 2.46)	2.06 (1.67 to
<i>36 to 47.9 months</i>	1.06)		2.63) *	2.73) *	*	2.54) *
48 to 59.9 months	0.79 (0.63 to		2.03) 2.011 (1.60 to	2.73) 2.06 (1.63 to	2.01 (1.59 to 2.53)	2.04 (1.61 to
	1.00)		2.011 (1.60 to 2.52) *	2.06 (1.65 to 2.58) *	2.01 (1.59 to 2.55) *	2.04 (1.61 10
CI 11 I				,		
Child sex	Ref		Ref		D.C	
Male	0.94 (0.83 to		1.05 (0.93 to		Ref	
Female	1.07)		1.19)		1.01 (0.89 to 1.14)	
	0.95 (0.93 to		0.97 (0.94		0.97 (0.94 to 0.99)	
Birth order	0.98) *		to1.001)		*	
Birth order						
(categorical)	Ref		Ref		D (
Primipara	0.92 (0.79 to		0.98 (0.85 to		Ref	
Multipara	1.07)		1.14)		0.98 (0.85 to 1.14)	
Birth type						
Singlet	Ref		Ref		Ref	
Twin/triplet	0.99 (0.65 to		0.82 (0.54 to		0.75 (0.49 to 1.17)	
1 with triplet	1.51)		1.29)		0.73 (0.13 to 1.17)	
	Daf					
	Ref		Ref			
36 . 1	1.08 (0.76 to		1.08 (0.76 to		D (
Maternal age	1.53)		1.53)		Ref	
15 to 19 years	1.19 (0.85 to		1.40 (0.99 to		1.27 (0.88 to 1.83)	
20 to 24 years	1.68)		1.97)		1.61 (1.12 to 2.31)	
25 to 29 years	1.08 (0.76 to		1.38 (097 to 1.96)		*	
30 to 34 years	1.53)		1.18 (0.82 to		1.54 (1.07 to 2.22)	
35 to 39 years	0.94 (0.65 to		1.71)		*	
40 to 44 years	1.35)		1.71) 1.15 (0.73 to		1.27 (0.86 to 1.86)	
45 to 49 years	1.04 (0.66 to		1.13 (0.73 to		1.42 (0.89 to 2.27)	
	1.64)		1.05 (0.52 to		1.06 (0.52 to 2.19)	
	0.52 (2.65 to		2.10)			
	1.04)		2.10)			
Maternal						
education	Ref		Ref		Ref	
None	1.44 (1.19 to		1.31 (1.10 to		1.38 (1.15 to 1.65)	
Primary	1.73) *		1.57) *		*	

Secondary	1.27 (1.08 to		1.31 (1.11 to		1.31 (1.12 to 1.55)	
Higher	1.50) *		1.54) *			
	1.33 (1.08 to		1.29 (1.06 to		1.37 (1.12 to 1.67)	
	1.63) *		1.58) *		*	
Maternal work	D (D (
No	Ref		Ref		Ref	
Yes	0.87 (0.73 to		0.86 (0.72 to		0.91 (0.75 to 1.08)	
	1.04)		1.02)		0.01 (0.70 to 1.00)	
Number of parities	0.95 (0.93 to		0.98 (0.95 to		0.98 (0.95 to 1.01)	
	0.98) *		1.01)		0.30 (0.33 to 1.01)	
Birth in last 5	0.99 (0.91 to		0.98 (0.91 to		1.00 (0.91 to 1.09)	
years	1.08)		1.07)		1.00 (0.91 10 1.09)	
Family size	0.99 (0.97to		0.99 (0.98 to		0.98 (0.97 to 1.00)	0.98 (0.97 to
	1.00)		1.01)		0.98 (0.97 to 1.00)	0.99) *
	Ref	Ref	Ref	Ref		Ref
Wealth index	1.29 (1.07 to	1.29 (1.07 to	1.29 (1.01 to	1.23 (1.02 to	Ref	1.23 (1.01 to
Poorest	1.55) *	1.55) *	1.47) *	1.49) *	1.21 (0.99 to 1.46)	1.49) *
Poorer	1.48 (1.22 to	1.48 (1.22 to	1.23 (1.02 to	1.43) 1.24 (1.02 to	1.24 (1.02 to 1.51)	1.43) 1.26 (1.03 to
Middle	1.40 (1.22 10	1.40 (1.22 10	1.49) *	1.51) *	*	1.54) *
Richer	1.85 (1.51 to	1.85 (1.51 to	1.54 (1.27 to	1.61 (1.31 to	1.66 (1.36 to 2.03)	1.73 (1.41 to
Richest	1.82) *	2.26) *	1.88) *	1.01 (1.31 10	*	2.11) *
Richesi					1.47 (1.20 to 1.82)	
	1.48 (1.21 to	1.48 (1.21 to	1.44 (1.17 to	1.47 (1.20 to	*	1.53 (1.24 to
	1.82) *	1.82) *	1.76) *	1.81) *		1.90) *
Place of residence	Ref		Ref		Ref	
Urban	0.91 (0.81 to		0.91 (0.80 to		0.87 (0.77 to 0.99)	
Rural	0.51 (0.01 10		0.51 (0.00 10		0.07 (0.77 10 0.33)	

Where, * = A significant association with various study covariates having p-value \leq 0.05, 1= Odds of fluid intake was adjusted with wealth index, 2 = Odds of food intake was adjusted with year of survey, child age, and wealth index, 3 = Odds of dietary adherence was adjusted with year of survey, child age, family size, and wealth index, α = dietary adherence is the sum of adequate intake of both fluid and food during the diarrhoeal illness.

4. Discussion

Diarrhoea is a common preventable paediatric illness that can be managed by simple and cost-effective interventions (17). Our study examined the adherence to diarrhoea case management practices among paediatric population using datasets of the last two waves of PDHS. To the best of our knowledge, our study is amongst the fewest studies that measured both therapeutic adherence (coadministration of zinc with ORS) and dietary adherence (adequate fluid intake and food) among diarrhoeal children of Pakistan. Additionally, our study also examined the individual as well as the cumulative adherence of each therapeutic and feeding indicator of diarrhoeal children using nationwide datasets of Pakistan (PDHS datasets). However, other studies from Pakistan had not assessed the therapeutic and dietary adherence exclusively among diarrhoeal children (18-21).

Our study reported a significant improvement in diarrhoea case management across two survey periods. The 2017-2018 survey showed over ten-fold improvement in therapeutic adherence to diarrhoea case management compared to the former survey of 2012-2013. This increase in therapeutic adherence in 2017-2018 was largely attributed to in-

creased zinc utilization (Table 3). Despite significant improvement in therapeutic adherence to diarrhoea case management practices, less than 10% of diarrhoeal cases showed therapeutic adherence (Table 2). Researchers from Asian and African regions reported that less than a third of children receive zinc together with ORS (22-26). However, around 40% of diarrhoeal children received ORS for fluid and electrolyte replenishment (23-26). The ORS has ability to reduce diarrhoea associated complications and mortality by replenishing the lost fluids and electrolytes (13), but it fails to alleviate the symptoms of diarrhoea, such as nausea, abdominal pain, and abdominal cramps (27). Owing to this reason, both the caregiver and care providers opt other methods for the therapeutic management of paediatric diarrhoea, including antimicrobial therapy (27, 28).

Across two survey periods, our study demonstrated a remarkable improvement in dietary adherence during diarrhoea case management (Table 3). Our study reported significantly higher dietary adherence among children of richer/richest socioeconomic classes than children of poor/poorer socioeconomic classes (Table 3). The low feeding adherence among children of poor/poorer socioeconomic classes is mainly attributed to food insecurity due to poverty. A report by the Pakistan Institute of Development Economics (PIDE) indicated that around one-third of the Pakistani population had been food insecure for over a decade (29). Poverty is the major cause of food insecurity. Inequalities in food distribution between various members of the household also contribute to low feeding adherence among children of poor/poorer socioeconomic status (30, 31). In general, in most societies, women, and children received food at the end, and our study also supported that each member increase in a family also reduces the odds of feeding adherence by 2% (1% to 3%), but on the other hands, increase the therapeutic adherence to ORS intake by 2% (1% to 3%). This reflects how the family size and dynamics contribute to diarrhoea case management.

Our study reported significantly higher odds of therapeutic and dietary adherence among children aged 12 to 59.9 months than those aged below one year. The reasons for the low therapeutic and dietary adherence among children aged below one year are unknown. We can postulate that exclusive breastfeeding (EBF) recommendation for children aged below six months can be one of the preponderant reasons for the low adherence among young children. During the EBF period, it is advisable to use ORS, medicine, multivitamins, and vaccines in conjunction with breast milk (32, 33). However, any solid, semi-solid, liquid, and soft food during the first six months of life reflect a deviation from the recommended feeding standards (32).

Moreover, delayed weaning after eight months of age also lowers the odds of adherence among children of this age group (21, 33). Similarly, the cultural practice of food abstinence during diarrhoeal illness and lack of maternal knowledge about diarrhoea case management also contributes to the low adherence (34). Different studies conducted in Pakistan reflected that most mothers are well-aware of zinc and ORS use for managing paediatric diarrhoea (18, 35, 36). However, a study conducted in the urban slums of Karachi reported inappropriate diarrhoea case management practices despite having adequate knowledge regarding diarrhoea case management (18). The knowledge-practice gap for diarrhoea case management can be averted by performing ethnographic studies, and these ethnographic studies would explore the underlying barriers to diarrhoea case management among Pakistani children.

Policy implications and recommendations:

Despite the significant increase in therapeutic and dietary adherence across two survey periods, this study reported low therapeutic and dietary guidelines of less than 10% and 40%, respectively. Therapeutic and dietary adherence among diarrhoeal cases can be improved by following various prevention and treatment strategies for managing acute paediatric diarrhoea. UNICEF, WHO, and various other national & international bodies have proposed numerous prevention and treatment packages to alleviate the morbidity

and mortality associated with diarrhoea. The diarrhoea prevention package includes immunization against rotavirus and measles, water and sanitation hygiene (WASH), hand hygiene, EBF, and vitamin-A supplementation. However, the treatment package consists of coadministration of zinc & ORS with adequate intake of food & fluid (37). Our study demonstrated that the therapeutic and dietary indicators for managing acute watery non-dysenteric diarrhoea are not practiced appropriately. In this regard, there is a need to define therapeutic and dietary indicators for the management and control of paediatric diarrhoea. These indicators should be part of national, regional, and institutional surveil-lance programs (19). Based on the findings of our study, we proposed a need for standardized dietary practices to manage paediatric diarrhoea aged between 6 to 59.9 months (38). Different literatures published in the past proposed using yogurt, banana, and rice with lentils because of their pre-biotic, hyperkalemic, and antidiarrhoeal properties (18, 20, 21). In this regard, promoting the coadministration of zinc & ORS with adequate food and fluid intake during diarrhoea case management is essential.

Strengths and limitations of the study:

Using third and fourth waves of PDHS datasets, this study is among the few studies that measure the therapeutic and dietary adherence of paediatric diarrhoea case management. This study presented paediatric diarrhoea case management at the national level because each PDHS survey's sample size was calculated from the estimates of the last census (39). Another more significant strength of this study was the multistage stratified sampling method for selecting participants. Similarly, excluding diarrhoea cases with blood and intravenous therapy from this study clearly defined the study population.

Besides various study strengths, the cross-sectional design of this study weakens the internal validity of this study. The data collectors in each PDHS survey relied on the mother's response regarding the diarrhoea prevalence during the last fourteen days. Similarly, the information related to diarrhoea case management was also subjective. There might be high chances of reporting bias, recall bias, or both. Moreover, the food and fluid intake question were very general, which asked about the amount of food and/or fluid given to a child during diarrhoeal episodes. Hence, this study did not assess dietary adherence properly.

5. Conclusion

Adherence to diarrhoea case management is a pivotal tool for reducing the morbidity and mortality associated with paediatric diarrhoea. The WHO and UNICEF have proposed using ORS, zinc, and adequate food & fluid intake for the proper case management of paediatric diarrhoea. Worldwide, millions of children die from lack of adherence, although Pakistan showed significant improvement in therapeutic and dietary adherence between two survey periods (from 2012-2013 to 2017-2018). Still, less than half of the children under five years are adherent to the recommended diarrhoea case management guidelines. A set of therapeutic strategies, such as coadministration of zinc & ORS with adequate intake of food & fluid, have the potential to avert the therapeutic and dietary non-adherence among diarrhoeal children.

Author Contributions: The idea of this project was conceptualized by A.K. The data management, data analysis, and original draft writing were performed by A.K.; R.H-S assisted A.K. with introduction writing, while D.A assisted A.K with discussion writing. S.K. and Z.l. provided technical support based on their expertise. The paper has been reviewed and edited by S.K and Z.L. All authors have read and agreed to the published version of the manuscript.

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Informed Consent Statement: In this study, the research team received data from DHS repository which was de-identified, i.e., it does not contain information in which a participant can either be

identified or traced by any means. Due to this reason, this study does not involve a statement for informed consent.

Data Availability Statement: The data from this study can be retrieved from the DHS program (www.dhsprogram.com).

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References

- 1. Bhutta ZA, Saeed M. Childhood infectious diseases: overview. International Encyclopedia of Public Health. 2008:620.
- 2. Jones G, Steketee RW, Black RE, Bhutta ZA, Morris SS, Group BCSS. How many child deaths can we prevent this year? The lancet. 2003;362(9377):65-71.
- 3. Roy RB, Whittaker E, Seddon JA, Kampmann B. Children and Mycobacterium tuberculosis: a review of susceptibility and protection. The Lancet Infectious diseases. 2019;19(3):e96.
- 4. DuPont HL. Diarrheal diseases in the developing world. Infectious Disease Clinics of North America. 1995;9(2):313-24.
- 5. World Health Organization. Diarrhoeal disease. 2017 [Available from: Available from: https://www.who.int/news-room/fact-sheets/detail/diarrhoeal-disease.
- 6. Taghavi S, Askari R. Hypovolemic shock. StatPearls [Internet]: StatPearls Publishing; 2021.
- 7. Desta T. Integrated management of childhood illness (IMCI): an overview. Ethiopian Journal of Health Sciences. 1999;9(1):59-60.
- 8. Troeger C, Blacker BF, Khalil IA, Rao PC, Cao S, Zimsen SR, et al. Estimates of the global, regional, and national morbidity, mortality, and aetiologies of diarrhoea in 195 countries: a systematic analysis for the Global Burden of Disease Study 2016. The Lancet Infectious Diseases. 2018;18(11):1211-28.
- 9. Turin CG, Ochoa TJ. The role of maternal breast milk in preventing infantile diarrhea in the developing world. Current tropical medicine reports. 2014;1(2):97-105.
- 10. Lakshminarayanan S, Jayalakshmy R. Diarrheal diseases among children in India: Current scenario and future perspectives. Journal of natural science, biology, and medicine. 2015;6(1):24.
- 11. Black RE. Progress in the use of ORS and zinc for the treatment of childhood diarrhea. J Glob Health. 2019;9(1):010101.
- 12. Richards L, Claeson M, Pierce NF. Management of acute diarrhea in children: lessons learned. Pediatr Infect Dis J. 1993;12(1):5-9.
- 13. Guarino A, Vecchio AL, Dias JA, Berkley JA, Boey C, Bruzzese D, et al. Universal recommendations for the management of acute diarrhea in nonmalnourished children. Journal of pediatric gastroenterology and nutrition. 2018;67(5):586.
- 14. Sreeramareddy CT, Low Y-P, Forsberg BC. Slow progress in diarrhea case management in low and middle income countries: evidence from cross-sectional national surveys, 1985–2012. BMC pediatrics. 2017;17(1):1-8.
- 15. National Institute of Population Studies (NIPS) P, ICF,. Pakistan demographic and health survey 2012–13. NIPS and ICF International Islamabad, Pakistan, and Calverton, Maryland, USA; 2013.
- 16. National Institute of Population Studies (NIPS) P, ICF,. Pakistan Demographic and Health Survey 2017-18. Islamabad, Pakistan: NIPS/Pakistan and ICF; 2019.
- 17. Bernadeta Dadonaite, Hannah Ritchie, Max Roser. "Diarrheal diseases"-Published online at OurWorldInData.org 2018 [Available from: Retrieved from: 'https://ourworldindata.org/diarrheal-diseases' [Online Resource].
- 18. Khaliq A. Knowledge and Practices on the Prevention and Management of Diarrhea in Children Under-2 Years Among Women Dwelling in Urban Slums of Karachi, Pakistan. Maternal and child health journal. 2022;26(7):1442-52.
- 19. Baig LA, Thaver I. Supervision for diarrhoea case management at the oral rehydration therapy corners and diarrhoea training units in Sindh. J Pak Med Assoc. 1997;47(8):208-10.

- 20. Imran Ali S, Naqvi B, Tasleem S. Mother's perception and attitude toward traditional dietary approaches concerning the management of childhood diarrhea. BAOJ Pharm Sci. 2019;4:52.
- 21. Syeda Sana Z, Nosheen Z, Shakir U, Nasir K, Muhammad Hamza J, Muneeba K. Mother's awareness and practices regarding home management of childhood diarrhea in a squatter settlement of Karachi. 2014.
- 22. Lazzerini M, Wanzira H. Oral zinc for treating diarrhoea in children. Cochrane database of systematic reviews. 2016(12).
- 23. Chopra M, Mason E, Borrazzo J, Campbell H, Rudan I, Liu L, et al. Ending of preventable deaths from pneumonia and diarrhoea: an achievable goal. The Lancet. 2013;381(9876):1499-506.
- 24. Lamberti LM, Taneja S, Mazumder S, LeFevre A, Black RE, Walker CLF. An external evaluation of the Diarrhea Alleviation through Zinc and ORS Treatment (DAZT) program in Gujarat and Uttar Pradesh, India. Journal of global health. 2015;5(2).
- 25. Schroder K, Battu A, Wentworth L, Houdek J, Fashanu C, Wiwa O, et al. Increasing coverage of pediatric diarrhea treatment in high-burden countries. Journal of global health. 2019;9(1).
- 26. Lam F, Pro G, Agrawal S, Shastri VD, Wentworth L, Stanley M, et al. Effect of enhanced detailing and mass media on community use of oral rehydration salts and zinc during a scale-up program in Gujarat and Uttar Pradesh. Journal of global health. 2019;9(1).
- 27. Harrell JE, Cheng SX. Inability to reduce morbidity of diarrhea by ORS: can we design a better therapy? Pediatric Research. 2018;83(3):559-63.
- 28. Chen J, Wan C-M, Gong S-T, Fang F, Sun M, Qian Y, et al. Chinese clinical practice guidelines for acute infectious diarrhea in children. World Journal of Pediatrics. 2018;14(5):429-36.
- 29. Ishaq A, Khalid M, Ahmad E. Food insecurity in Pakistan: A region-wise analysis of trends. Pakistan Insfitute of Development Economics (PIDE) Working Papers. 2018;157.
- 30. Harris-Fry H, Shrestha N, Costello A, Saville NM. Determinants of intra-household food allocation between adults in South Asia a systematic review. International Journal for Equity in Health. 2017;16(1):107.
- 31. Earnshaw VA, Karpyn A. Understanding stigma and food inequity: a conceptual framework to inform research, intervention, and policy. Translational Behavioral Medicine. 2020;10(6):1350-7.
- 32. Khaliq A. Association of Infant Feeding Indicators and Infant Feeding Practices with Coexisting Forms of Malnutrition in Children under Six Months of Age. Nutrients. 2022;14(20).
- 33. Khaliq A, Qamar M, Hussaini SA, Azam K, Zehra N, Hussain M, et al. Assessment of knowledge and practices about breastfeeding and weaning among working and non-working mothers. J Pak Med Assoc. 2017;67(3):332-8.
- 34. Khatun M, Sharmin A, Bithi IJ. Mothers' Knowledge, Attitude and Practice regarding Prevention of Diarrhea among Children: An Empirical Investigation. Randwick International of Social Science Journal. 2021;2(4):469-75.
- 35. Riaz N, Muntaha ST, Qibtia M, Sohail S. Use of Zinc and ORS in Home Management of Diarrhea: Knowledge of Mothers attending a Tertiary Care Hospital. Journal of Islamabad Medical & Dental College. 2019;8(3):135-40.
- 36. Duggal MN, Bari A, Mansoor A, Saeed S, Rizwan W, Nayar J. Awareness of quantity of ORS and Zinc among mothers of under five children with diarrhoea at a tertiary care Centre Lahore. J Pak Med Assoc. 2021;71(3):921-4.
- 37. Wardlaw T, Salama P, Brocklehurst C, Chopra M, Mason E. Diarrhoea: why children are still dying and what can be done. The lancet. 2010;375(9718):870-2.
- 38. Bajait C, Thawani V. Role of zinc in pediatric diarrhea. Indian J Pharmacol. 2011;43(3):232-5.
- 39. Khaliq A, Wraith D, Miller Y, Nambiar-Mann S. Prevalence, Trends, and Socioeconomic Determinants of Coexisting Forms of Malnutrition Amongst Children under Five Years of Age in Pakistan. Nutrients. 2021;13(12).