

Additional File 2 : Comparison of PKU-QOL scores according to overall assessment of health status

Table 2_a : Comparison of the Child PKU-QOL scores according to overall assessment of health status

Modules	Domains		Poor (N=0)	Fair (N=2)	Good (N=19)	Very good (N=41)	Excellent (N=28)	p-value*
Symptoms	Headaches	Mean (SD)	na	12.5 (17.7)	16.7 (17.1)	13.1 (18.8)	19.2 (28.6)	0.829
		Median (Q1 – Q3)	na	12.5 (0.0 – 25.0)	25.0 (0.0 – 50.0)	0.0 (0.0 – 50.0)	0.0 (0.0 – 100.0)	
	Stomach aches	Mean (SD)	na	50.0 (70.7)	15.3 (21.2)	17.5 (22.1)	10.6 (17.6)	0.518
		Median (Q1 – Q3)	na	50.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 50.0)	
	Tiredness	Mean (SD)	na	62.5 (53.0)	38.9 (24.6)	30.6 (26.2)	36.5 (26.7)	0.382
		Median (Q1 – Q3)	na	62.5 (25.0 – 100.0)	50.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	
	Irritability	Mean (SD)	na	50.0 (70.7)	16.7 (24.3)	21.3 (29.7)	15.4 (26.5)	0.738
		Median (Q1 – Q3)	na	50.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	
	Aggressiveness	Mean (SD)	na	50.0 (70.7)	18.1 (30.7)	11.3 (20.4)	7.7 (17.0)	0.534
		Median (Q1 – Q3)	na	50.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	
	Moodiness	Mean (SD)	na	50.0 (70.7)	11.1 (19.6)	21.3 (24.4)	11.5 (19.0)	0.205
		Median (Q1 – Q3)	na	50.0 (0.0 – 100.0)	0.0 (0.0 – 50.0)	12.5 (0.0 – 75.0)	0.0 (0.0 – 50.0)	
	Sadness	Mean (SD)	na	62.5 (53.0)	9.7 (15.2)	18.8 (25.8)	14.8 (18.7)	0.180
		Median (Q1 – Q3)	na	62.5 (25.0 - 100.0)	0.0 (0.0 – 50.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 50.0)	
	Anxiety	Mean (SD)	na	25.0 (35.4)	5.6 (13.7)	11.9 (20.4)	6.7 (16.7)	0.347
		Median (Q1 – Q3)	na	25.0 (0.0 – 50.0)	0.0 (0.0 – 50.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 50.0)	
	Lack of concentration	Mean (SD)	na	50.0 (35.4)	20.6 (28.3)	21.3 (26.2)	23.0 (31.4)	0.539
		Median (Q1 – Q3)	na	50.0 (25.0 – 75.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
Slow thinking	Mean (SD)	na	37.5 (17.7)	26.5 (22.5)	23.5 (31.2)	19.0 (22.0)	0.453	
	Median (Q1 – Q3)	na	37.5 (25.0 – 50.0)	25.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	25.0 (0.0 – 75.0)		
PKU in general	Emotional impact of PKU	Mean (SD)	na	41.7 (23.6)	38.0 (24.5)	32.7 (23.3)	30.1 (18.0)	0.690
		Median (Q1 – Q3)	na	41.7 (25.0 – 58.3)	37.5 (0.0 – 75.0)	29.2 (0.0 – 83.3)	29.2 (0.0 – 66.7)	
	Practical impact of PKU	Mean (SD)	na	25.0 (35.4)	9.0 (12.7)	11.3 (18.5)	11.6 (16.2)	0.900
		Median (Q1 – Q3)	na	25.0 (0.0 – 50.0)	0.0 (0.0 – 37.5)	0.0 (0.0 – 75.0)	0.0 (0.0 – 50.0)	
	Social impact of PKU	Mean (SD)	na	37.5 (29.5)	23.6 (12.9)	21.5 (21.5)	13.5 (12.5)	0.086
		Median (Q1 – Q3)	na	37.5 (16.7 – 58.3)	25.0 (0.0 – 50.0)	16.7 (0.0 – 91.7)	8.3 (0.0 – 33.3)	
	Overall impact of PKU	Mean (SD)	na	35.9 (11.0)	25.3 (12.4)	23.1 (17.9)	19.2 (12.2)	0.161
		Median (Q1 – Q3)	na	35.9 (28.1 – 43.8)	28.1 (0.0 – 43.8)	18.8 (0.0 – 78.1)	17.2 (0.0 – 46.9)	
	Anxiety - Blood test	Mean (SD)	na	0.0 (0.0)	13.2 (21.4)	14.7 (17.2)	9.7 (13.6)	0.266
		Median (Q1 – Q3)	na	0.0 (0.0 – 0.0)	0.0 (0.0 – 50.0)	12.5 (0.0 – 62.5)	0.0 (0.0 – 50.0)	
	Anxiety - Phe levels	Mean (SD)	na	50.0 (35.4)	40.3 (31.1)	48.8 (38.4)	44.4 (39.4)	0.902
		Median (Q1 – Q3)	na	50.0 (25.0 – 75.0)	25.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	

Table 2_a : Comparison of the Child PKU-QOL scores according to overall assessment of health status (cont'd)

Modules	Domains		Poor (N=0)	Fair (N=2)	Good (N=19)	Very good (N=41)	Excellent (N=28)	p-value*
Supplement administration	Adherence to supplements	Mean (SD)	na	50.0 (70.7)	8.8 (13.1)	10.9 (15.2)	9.9 (14.3)	0.871
		Median (Q1 – Q3)	na	50.0 (0.0 – 100.0)	0.0 (0.0 – 37.5)	0.0 (0.0 – 50.0)	0.0 (0.0 – 50.0)	
	Practical impact of supplements	Mean (SD)	na	50.0 (70.7)	26.5 (40.0)	17.3 (29.3)	13.0 (25.1)	0.571
		Median (Q1 – Q3)	na	50.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)	
	Guilt if poor adherence to supplements	Mean (SD)	na	12.5 (17.7)	41.2 (35.3)	51.3 (39.0)	41.3 (40.3)	0.444
		Median (Q1 – Q3)	na	12.5 (0.0 – 25.0)	25.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Relationships within family because of supplements	Mean (SD)	na	25.0 (35.4)	7.4 (14.7)	13.5 (27.4)	4.0 (11.8)	0.364
		Median (Q1 – Q3)	na	25.0 (0.0 – 50.0)	0.0 (0.0 – 50.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 50.0)	
	Taste - Supplements	Mean (SD)	na	0.0 (0.0)	29.4 (29.6)	32.7 (29.9)	47.8 (32.8)	0.063
		Median (Q1 – Q3)	na	0.0 (0.0 – 0.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	
Dietary protein restriction	Food temptations	Mean (SD)	na	31.3 (8.8)	23.4 (29.2)	30.1 (27.8)	34.1 (28.2)	0.545
		Median (Q1 – Q3)	na	31.3 (25.0 – 37.5)	18.8 (0.0 – 100.0)	25.0 (0.0 – 100.0)	31.3 (0.0 – 87.5)	
	Adherence to dietary protein restriction	Mean (SD)	na	37.5 (17.7)	9.4 (13.3)	11.2 (16.4)	7.4 (16.7)	0.059
		Median (Q1 – Q3)	na	37.5 (25.0 – 50.0)	0.0 (0.0 – 37.5)	0.0 (0. – 62.5)	0.0 (0.0 – 62.5)	
	Social impact of dietary protein restriction	Mean (SD)	na	22.5 (24.7)	21.1 (19.3)	19.2 (19.6)	17.0 (17.8)	0.884
		Median (Q1 – Q3)	na	22.5 (5.0 – 40.0)	17.5 (0.0 – 62.5)	15.0 (0.0 – 80.0)	15.0 (0.0 – 60.0)	
	Taste – low protein food	Mean (SD)	na	0.0 (0.0)	17.2 (19.8)	20.3 (21.9)	18.2 (29.1)	0.390
		Median (Q1 – Q3)	na	0.0 (0.0 – 0.0)	12.5 (0.0 – 50.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	
	Food enjoyment	Mean (SD)	na	50.0 (70.7)	11.1 (15.4)	20.6 (22.6)	13.9 (24.4)	0.268
		Median (Q1 – Q3)	na	50.0 (0.0 – 100.0)	0.0 (0.0 – 50.0)	25.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Guilt if dietary protein restriction not followed	Mean (SD)	na	62.5 (53.0)	57.8 (35.0)	53.9 (41.3)	49.0 (44.8)	0.870
		Median (Q1 – Q3)	na	62.5 (25.0 – 100.0)	62.5 (0.0 – 100.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	
	Overall difficulty following dietary protein restriction	Mean (SD)	na	0.0 (0.0)	18.8 (23.3)	21.2 (27.2)	17.6 (32.4)	0.362
		Median (Q1 – Q3)	na	0.0 (0.0 – 0.0)	12.5 (0.0 – 75.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	

*Non-parametric P-value for between-group comparisons: Kruskal-Wallis
In bold p-value<0.05

Table 2_b : Comparison of the Adolescent PKU-QOL scores according to overall assessment of health status

Modules	Domains		Poor (N=0)	Fair (N=0)	Good (N=38)	Very good (N=42)	Excellent (N=30)	p-value*
Symptoms	Self-rated status health	Mean (SD)	na	na	39.3 (21.3)	41.3 (23.0)	20.8 (24.6)	0.001
		Median (Q1 – Q3)	na	na	50.0 (0.0 – 75.0)	50.0 (0.0 – 75.0)	12.5 (0.0 – 75.0)	
	Headaches	Mean (SD)	na	na	18.6 (24.5)	20.6 (25.2)	20.8 (27.1)	0.908
		Median (Q1 – Q3)	na	na	0.0 (0.0 – 75.0)	12.5 (0.0 – 100.0)	12.5 (0.0 – 100.0)	
	Stomach aches	Mean (SD)	na	na	16.4 (21.8)	15.6 (20.2)	10.8 (21.5)	0.355
		Median (Q1 – Q3)	na	na	0.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Tiredness	Mean (SD)	na	na	43.6 (28.0)	41.9 (23.6)	35.0 (26.7)	0.264
		Median (Q1 – Q3)	na	na	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Irritability	Mean (SD)	na	na	32.9 (26.3)	20.2 (25.4)	21.7 (24.3)	0.062
		Median (Q1 – Q3)	na	na	50.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)	
	Aggressiveness	Mean (SD)	na	na	17.1 (26.3)	11.3 (24.2)	6.7 (13.0)	0.305
		Median (Q1 – Q3)	na	na	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 50.0)	
	Moodiness	Mean (SD)	na	na	24.3 (24.6)	16.7 (21.9)	19.2 (21.5)	0.337
		Median (Q1 – Q3)	na	na	25.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	12.5 (0.0 – 50.0)	
Sadness	Mean (SD)	na	na	14.3 (20.4)	11.9 (20.1)	10.0 (18.1)	0.570	
	Median (Q1 – Q3)	na	na	0.0 (0.0 – 75.0)	0.0 (0.0 – 50.0)	0.0 (0.0 – 50.0)		
Anxiety	Mean (SD)	na	na	20.0 (25.6)	10.7 (20.0)	11.7 (25.2)	0.077	
	Median (Q1 – Q3)	na	na	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)		
Lack of concentration	Mean (SD)	na	na	25.7 (22.3)	16.7 (23.9)	16.7 (26.5)	0.057	
	Median (Q1 – Q3)	na	na	25.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)		
Slow thinking	Mean (SD)	na	na	15.7 (19.3)	11.9 (20.1)	14.2 (20.4)	0.549	
	Median (Q1 – Q3)	na	na	0.0 (0.0 – 50.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 50.0)		
PKU in general	Emotional impact of PKU	Mean (SD)	na	na	38.9 (23.0)	28.2 (17.6)	29.7 (17.2)	0.165
		Median (Q1 – Q3)	na	na	30.0 (0.0 – 90.0)	30.0 (0.0 – 70.0)	30.0 (0.0 – 70.0)	
	Practical impact of PKU	Mean (SD)	na	na	19.3 (17.1)	9.0 (11.0)	8.3 (11.2)	0.095
		Median (Q1 – Q3)	na	na	16.7 (0.0 – 50.0)	8.3 (0.0 – 33.3)	8.3 (0.0 – 33.3)	
	Social impact of PKU	Mean (SD)	na	na	18.1 (18.1)	14.8 (12.7)	13.8 (11.6)	0.723
		Median (Q1 – Q3)	na	na	16.7 (0.0 – 91.7)	8.3 (0.0 – 50.0)	8.3 (0.0 – 33.3)	
	Overall impact of PKU	Mean (SD)	na	na	28.4 (18.8)	19.9 (12.1)	20.2 (12.7)	0.093
		Median (Q1 – Q3)	na	na	25.0 (0.0 – 85.0)	20.0 (2.3 – 50.0)	17.5 (0.0 – 55.0)	
	Anxiety - Blood test	Mean (SD)	na	na	13.6 (25.3)	9.8 (19.5)	9.2 (16.1)	0.990
		Median (Q1 – Q3)	na	na	0.0 (0.0 – 100.0)	0.0 (0.0 – 87.5)	0.0 (0.0 – 50.0)	
	Anxiety - Phe levels	Mean (SD)	na	na	40.7 (33.3)	33.5 (31.9)	32.5 (29.5)	0.530
		Median (Q1 – Q3)	na	na	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	

Table 2_b : Comparison of the Adolescent PKU-QOL scores according to overall assessment of health status (cont'd)

Modules	Domains		Poor (N=0)	Fair (N=0)	Good (N=38)	Very good (N=42)	Excellent (N=30)	p-value*
Supplement administration	Adherence to supplements	Mean (SD)	na	na	21.4 (25.1)	11.1 (12.9)	11.5 (16.4)	0.239
		Median (Q1 – Q3)	na	na	8.3 (0.0 – 83.3)	6.3 (0.0 – 50.0)	6.3 (0.0 -66.7)	
	Practical impact of supplements	Mean (SD)	na	na	24.0 (28.6)	14.2 (16.9)	20.3 (24.4)	0.447
		Median (Q1 – Q3)	na	na	12.5 (0.0 – 100.0)	12.5 (0.0 – 62.5)	6.3 (0.0 – 75.0)	
	Guilt if poor adherence to supplements	Mean (SD)	na	na	40.3 (34.0)	43.9 (37.5)	46.2 (36.5)	0.859
		Median (Q1 – Q3)	na	na	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	
	Relationships within family because of supplements	Mean (SD)	na	na	14.8 (28.3)	14.9 (26.0)	13.0 (20.6)	0.983
		Median (Q1 – Q3)	na	na	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 50.0)	
	Taste - Supplements	Mean (SD)	na	na	44.5 (26.0)	39.9 (23.2)	38.0 (26.1)	0.634
		Median (Q1 – Q3)	na	na	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	
Dietary protein restriction	Food temptations	Mean (SD)	na	na	30.8 (31.6)	27.0 (26.4)	19.6 (25.5)	0.360
		Median (Q1 – Q3)	na	na	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	12.5 (0.0 – 87.5)	
	Adherence to dietary protein restriction	Mean (SD)	na	na	21.3 (25.6)	10.7 (12.4)	7.6 (12.6)	0.083
		Median (Q1 – Q3)	na	na	12.5 (0.0 – 83.3)	8.3 (0.0 – 50.0)	0.0 (0.0 – 50.0)	
	Practical impact of dietary protein restriction	Mean (SD)	na	na	30.7 (21.8)	24.6 (16.4)	25.9 (14.7)	0.659
		Median (Q1 – Q3)	na	na	26.8 (0.0 – 85.7)	24.3 (0.0 – 64.3)	28.6 (0.0 – 67.9)	
	Social impact of dietary protein restriction	Mean (SD)	na	na	19.4 (25.2)	11.1 (18.3)	10.7 (15.2)	0.356
		Median (Q1 – Q3)	na	na	7.5 (0.0 – 90.0)	5.0 (0.0 – 80.0)	5.0 (0.0 – 65.0)	
	Overall impact of dietary protein restriction	Mean (SD)	na	na	26.0 (21.1)	17.2 (15.1)	18.3 (12.8)	0.177
		Median (Q1 – Q3)	na	na	19.4 (0.0 – 86.4)	13.1 (0.0 – 59.1)	18.2 (0.0 – 52.3)	
	Taste – low protein food	Mean (SD)	na	na	33.1 (31.2)	19.1 (17.5)	26.3 (25.6)	0.228
		Median (Q1 – Q3)	na	na	25.0 (0.0 – 100.0)	25.0 (0.0 – 50.0)	25.0 (0.0 – 100.0)	
	Food enjoyment	Mean (SD)	na	na	15.8 (26.7)	14.2 (26.7)	19.6 (27.1)	0.461
		Median (Q1 – Q3)	na	na	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	
Guilt if dietary protein restriction not followed	Mean (SD)	na	na	44.4 (35.8)	50.6 (38.3)	42.4 (30.6)	0.657	
	Median (Q1 – Q3)	na	na	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)		
Overall difficulty following dietary protein restriction	Mean (SD)	na	na	21.2 (30.1)	15.8 (23.6)	13.0 (26.0)	0.458	
	Median (Q1 – Q3)	na	na	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)		

*Non-parametric P-value for between-group comparisons: Kruskal-Wallis
In bold p-value<0.05

Table 2_c : Comparison of the Adult PKU-QOL scores according to overall assessment of health status

Modules	Domains	Poor (N=0)	Fair (N=3)	Good (N=28)	Very good (N=46)	Excellent (N=26)	p-value*	
Symptoms	Self-rated health status	Mean (SD)	na	66.7 (28.9)	46.3 (16.6)	30.4 (21.0)	22.1 (22.7)	<0.001
		Median (Q1 – Q3)	na	50.0 (50.0 – 100.0)	50.0 (25.0 – 75.0)	25.0 (0.0 – 75.0)	25.0 (0.0 – 50.0)	
Headaches	Mean (SD)	na	33.3 (28.9)	26.9 (25.4)	19.0 (24.3)	18.3 (20.7)	0.395	
	Median (Q1 – Q3)	na	50.0 (0.0 – 50.0)	25.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	12.5 (0.0 – 50.0)		
Stomach aches	Mean (SD)	na	50.0 (25.0)	26.0 (32.0)	13.6 (22.8)	9.6 (18.8)	0.013	
	Median (Q1 – Q3)	na	50.0 (25.0 – 75.0)	12.5 (0.0 – 100.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 50.0)		
Tiredness	Mean (SD)	na	41.7 (14.4)	51.0 (26.9)	44.0 (27.5)	38.5 (29.4)	0.479	
	Median (Q1 – Q3)	na	50.0 (25.0 – 50.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	37.5 (0.0 – 100.0)		
Trembling hands	Mean (SD)	na	16.7 (28.9)	15.4 (22.4)	10.3 (22.7)	13.5 (24.7)	0.517	
	Median (Q1 – Q3)	na	0.0 (0.0 – 50.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)		
Irritability	Mean (SD)	na	66.7 (14.4)	35.2 (25.2)	34.2 (22.6)	26.9 (27.3)	0.050	
	Median (Q1 – Q3)	na	75.0 (50.0 – 75.0)	50.0 (0.0 – 75.0)	50.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)		
Aggressiveness	Mean (SD)	na	41.7 (38.2)	14.4 (21.4)	9.2 (17.8)	13.5 (23.7)	0.243	
	Median (Q1 – Q3)	na	50.0 (0.0 – 75.0)	0.0 (0.0 – 50.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)		
Moodiness	Mean (SD)	na	33.3 (28.9)	27.9 (24.8)	23.4 (25.5)	16.3 (23.4)	0.297	
	Median (Q1 – Q3)	na	50.0 (0.0 – 50.0)	25.0 (0.0 – 75.0)	25.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)		
Sadness	Mean (SD)	na	50.0 (25.0)	40.4 (30.9)	26.1 (26.3)	21.2 (26.2)	0.044	
	Median (Q1 – Q3)	na	50.0 (25.0 – 75.0)	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)		
Anxiety	Mean (SD)	na	50.0 (43.3)	24.0 (28.7)	15.2 (23.9)	29.8 (34.7)	0.130	
	Median (Q1 – Q3)	na	75.0 (0.0 – 75.0)	12.5 (0.0 – 75.0)	0.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)		
Lack of concentration	Mean (SD)	na	33.3 (14.4)	31.7 (27.9)	23.4 (25.5)	17.3 (25.3)	0.160	
	Median (Q1 – Q3)	na	25.0 (25.0 – 50.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)		
Slow thinking	Mean (SD)	na	25.0 (25.0)	28.8 (30.6)	15.2 (25.0)	11.5 (23.7)	0.038	
	Median (Q1 – Q3)	na	25.0 (0.0 – 50.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)		

Table 2_c : Comparison of the Adult PKU-QOL scores according to overall assessment of health status (cont'd)

Modules	Domains		Poor (N=0)	Fair (N=3)	Good (N=28)	Very good (N=46)	Excellent (N=26)	p-value*
PKU in general	Emotional impact of PKU	Mean (SD)	na	55.0 (10.0)	58.5 (16.7)	43.0 (22.0)	32.5 (20.6)	<0.001
		Median (Q1 – Q3)	na	55.0 (45.0 – 65.0)	60.0 (20.0 – 85.0)	45.0 (0.0 – 100.0)	30.0 (0.0 – 95.0)	
	Practical impact of PKU	Mean (SD)	na	21.5 (14.8)	21.9 (17.8)	20.0 (17.6)	9.9 (13.0)	0.023
		Median (Q1 – Q3)	na	18.8 (8.3 – 37.5)	25.0 (0.0 – 58.3)	16.7 (0.0 – 75.0)	6.3 (0.0 – 43.8)	
	Social impact of PKU	Mean (SD)	na	27.8 (30.1)	23.0 (19.2)	17.1 (13.5)	12.3 (10.7)	0.173
		Median (Q1 – Q3)	na	12.5 (8.3 – 62.5)	18.8 (0.0 – 68.8)	14.6 (0.0 – 43.8)	10.4 (0.0 – 41.7)	
	Overall impact of PKU	Mean (SD)	na	41.7 (14.7)	39.1 (14.2)	28.8 (15.5)	21.3 (12.5)	0.004
		Median (Q1 – Q3)	na	41.7 (31.3 – 52.1)	39.6 (12.5 – 68.8)	27.1 (2.1 – 62.5)	17.7 (6.3 – 52.1)	
	Anxiety - Blood test	Mean (SD)	na	18.8 (8.8)	6.0 (13.5)	14.2 (27.8)	8.3 (17.2)	0.218
		Median (Q1 – Q3)	na	18.8 (12.5 – 25.0)	0.0 (0.0 – 50.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)	
	Anxiety - Phe levels	Mean (SD)	na	75.0 (25.0)	35.2 (28.8)	34.8 (25.0)	37.5 (28.5)	0.170
		Median (Q1 – Q3)	na	75.0 (50.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 75.0)	
	Anxiety - Phe levels during pregnancy	Mean (SD)	na	33.3 (38.2)	86.5 (16.5)	83.6 (22.5)	79.7 (24.5)	0.092
		Median (Q1 – Q3)	na	25.0 (0.0 – 75.0)	100.0 (50.0 – 100.0)	100.0 (25.0 – 100.0)	100.0 (50.0 – 100.0)	
	Financial impact of PKU	Mean (SD)	na	41.7 (52.0)	30.6 (31.3)	14.1 (23.4)	13.5 (22.6)	0.035
		Median (Q1 – Q3)	na	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)	
Information on PKU	Mean (SD)	na	50.0 (50.0)	38.9 (23.3)	34.2 (25.5)	27.9 (19.1)	0.265	
	Median (Q1 – Q3)	na	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 75.0)		
Supplement administration	Adherence to supplements	Mean (SD)	na	16.7 (11.8)	20.2 (20.1)	19.3 (18.1)	21.8 (25.1)	0.991
		Median (Q1 – Q3)	na	16.7 (8.3 – 25.0)	16.7 (0.0 – 66.7)	16.7 (0.0 – 66.7)	8.3 (0.0 – 66.47)	
	Practical impact of supplements	Mean (SD)	na	60.4 (20.1)	28.5 (26.8)	18.8 (17.7)	15.9 (14.7)	0.033
		Median (Q1 – Q3)	na	68.8 (37.5 – 75.0)	18.8 (0.0 – 100.0)	18.8 (0.0 – 56.3)	12.5 (0.0 – 37.5)	
	Guilt if poor adherence to supplements	Mean (SD)	na	41.7 (52.0)	46.7 (34.0)	49.3 (35.1)	47.2 (27.0)	0.969
		Median (Q1 – Q3)	na	25.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 75.0)	
	Relationships within family because of supplements	Mean (SD)	na	8.3 (14.4)	9.1 (26.2)	12.8 (22.5)	10.0 (18.8)	0.634
		Median (Q1 – Q3)	na	0.0 (0.0 – 25.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	
	Taste - Supplements	Mean (SD)	na	58.3 (14.4)	45.2 (33.2)	48.6 (19.5)	55.3 (28.4)	0.543
		Median (Q1 – Q3)	na	50.0 (50.0 – 75.0)	50.0 (0.0 – 100.0)	50.0 (25.0 – 100.0)	50.0 (0.0 – 100.0)	

Table 2_c : Comparison of the Adult PKU-QOL scores according to overall assessment of health status (cont'd)

Modules	Domains		Poor (N=0)	Fair (N=3)	Good (N=28)	Very good (N=46)	Excellent (N=26)	p-value*
Dietary protein restriction	Food temptations	Mean (SD)	na	50.0 (12.5)	45.3 (29.2)	34.5 (22.4)	23.7 (19.0)	0.038
		Median (Q1 – Q3)	na	50.0 (37.5 – 62.5)	50.0 (0.0 – 100.0)	43.8 (0.0 – 75.0)	25.0 (0.0 – 50.0)	
	Adherence to dietary protein restriction	Mean (SD)	na	12.5 (17.7)	21.2 (17.4)	21.0 (18.4)	16.4 (19.7)	0.671
		Median (Q1 – Q3)	na	12.5 (0.0 – 25.0)	18.8 (0.0 – 56.3)	22.5 (0.0 – 55.0)	10.0 (0.0 56.3)	
	Social impact of dietary protein restriction	Mean (SD)	na	51.4 (31.8)	21.7 (19.8)	15.4 (15.1)	13.9 (16.7)	0.064
		Median (Q1 – Q3)	na	58.3 (16.7 – 79.2)	16.7 (0.0 – 75.0)	11.3 (0.0 – 54.2)	8.3 (0.0 – 54.2)	
	Practical impact of dietary protein restriction	Mean (SD)	na	64.3 (15.6)	38.5 (16.5)	31.7 (18.6)	33.7 (23.0)	0.079
		Median (Q1 – Q3)	na	71.4 (46.4 – 75.0)	39.6 (15.0 – 71.4)	30.4 (0.0 – 64.3)	32.1 (7.1 – 67.9)	
	Overall impact of dietary protein restriction	Mean (SD)	na	58.3 (22.5)	29.5 (14.9)	23.9 (15.0)	23.7 (18.8)	0.063
		Median (Q1 – Q3)	na	67.3 (32.7 – 75.0)	27.1 (9.1 – 58.3)	22.8 (0.0 – 53.8)	20.8 (2.5 – 59.6)	
	Taste – low protein food	Mean (SD)	na	58.3 (14.4)	36.8 (25.5)	30.5 (19.8)	30.4 (17.5)	0.125
		Median (Q1 – Q3)	na	50.0 (50.0 – 75.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 75.0)	25.0 (0.0 – 50.0)	
	Food enjoyment	Mean (SD)	na	58.3 (52.0)	29.3 (28.9)	25.7 (29.3)	18.4 (26.1)	0.335
		Median (Q1 – Q3)	na	75.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	
Guilt if dietary protein restriction not followed	Mean (SD)	na	50.0 (43.3)	50.0 (29.9)	57.2 (35.3)	52.5 (26.8)	0.854	
	Median (Q1 – Q3)	na	25.0 (25.0 – 100.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)		
Overall difficulty following dietary protein restriction	Mean (SD)	na	41.7 (38.2)	34.8 (29.9)	23.1 (25.9)	28.9 (29.2)	0.383	
	Median (Q1 – Q3)	na	50.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 75.0)		

*Non-parametric P-value for between-group comparisons: Kruskal-Wallis
 In bold p-value<0.05

Table 2_d : Comparison of the Parent PKU-QOL scores according to patient's overall assessment of health status

Modules	Domains	Poor (N=0)	Fair (N=2)	Good (N=64)	Very good (N=101)	Excellent (N=84)	p-value*	
Symptoms	Child health status	Mean (SD)	na	12.5 (17.7)	38.5 (19.7)	34.3 (25.4)	23.2 (22.9)	<0.001
		Median (Q1 – Q3)		12.5 (0.0 – 25.0)	50.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Headaches	Mean (SD)	na	25.0 (35.4)	20.1 (23.2)	15.4 (20.7)	9.2 (17.7)	0.012
		Median (Q1 – Q3)		25.0 (0.0 – 50.0)	25.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	
	Stomach aches	Mean (SD)	na	37.5 (53.0)	17.6 (22.5)	21.0 (24.1)	16.7 (25.2)	0.391
		Median (Q1 – Q3)		37.5 (0.0 – 75.0)	0.0 (0.0 – 75.0)	25.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Tiredness	Mean (SD)	na	12.5 (17.7)	35.7 (25.2)	31.3 (23.8)	30.1 (26.3)	0.313
		Median (Q1 – Q3)		12.5 (0.0 – 25.0)	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Irritability	Mean (SD)	na	62.5 (17.7)	43.9 (27.3)	34.8 (26.7)	31.3 (27.7)	0.015
		Median (Q1 – Q3)		62.5 (50.0 – 75.0)	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Aggressiveness	Mean (SD)	na	37.5 (17.7)	16.9 (24.3)	11.5 (21.0)	11.6 (20.9)	0.085
		Median (Q1 – Q3)		37.5 (25.0 – 50.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Moodiness	Mean (SD)	na	12.5 (17.7)	32.0 (24.2)	25.8 (26.2)	27.7 (26.2)	0.291
		Median (Q1 – Q3)		12.5 (0.0 – 25.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
Sadness	Mean (SD)	na	25.0 (0.0)	25.4 (21.6)	18.9 (22.6)	12.2 (17.7)	0.001	
	Median (Q1 – Q3)		25.0 (25.0 – 25.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)		
Anxiety	Mean (SD)	na	0.0 (0.0)	20.8 (26.7)	13.4 (20.7)	10.8 (18.4)	0.066	
	Median (Q1 – Q3)		0.0 (0.0 – 0.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)		
Lack of concentration	Mean (SD)	na	12.5 (17.7)	36.7 (30.7)	28.4 (27.2)	28.4 (27.1)	0.278	
	Median (Q1 – Q3)		12.5 (0.0 – 25.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)		
Slow thinking	Mean (SD)	na	12.5 (17.7)	27.6 (28.4)	18.0 (26.3)	16.3 (23.9)	0.053	
	Median (Q1 – Q3)		12.5 (0.0 – 25.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)		

Table 2_d : Comparison of the Parent PKU-QOL scores according to patient's overall assessment of health status (cont'd)

Modules	Domains		Poor (N=0)	Fair (N=2)	Good (N=64)	Very good (N=101)	Excellent (N=84)	p-value*
PKU in general	Emotional impact of PKU	Mean (SD)	na	28.1 (13.3)	52.4 (22.4)	41.6 (21.8)	37.3 (22.9)	<0.001
		Median (Q1 – Q3)		28.1 (18.8 – 37.5)	56.3 (6.3 – 87.5)	37.5 (6.3 – 100.0)	34.4 (0.0 – 100.0)	
	Practical impact of PKU	Mean (SD)	na	25.4 (28.9)	18.9 (15.8)	14.9 (13.8)	10.4 (13.0)	0.007
		Median (Q1 – Q3)		25.4 (5.0 – 45.8)	16.7 (0.0 – 70.0)	12.5 (0.0 – 50.0)	5.0 (0.0 – 54.2)	
	Social impact of PKU	Mean (SD)	na	25.0 (21.2)	21.7 (17.5)	16.6 (17.0)	14.4 (15.3)	0.019
		Median (Q1 – Q3)		25.0 (10.0 – 40.0)	15.0 (0.0 – 90.0)	10.0 (0.0 – 80.0)	10.0 (0.0 – 70.0)	
	Overall impact of PKU	Mean (SD)	na	26.2 (21.9)	30.6 (15.9)	23.5 (14.9)	19.6 (14.2)	<0.001
		Median (Q1 – Q3)		26.2 (10.7 – 41.7)	28.3 (6.7 – 76.9)	21.2 (1.7 – 67.3)	15.8 (0.0 – 73.1)	
	Child anxiety - Blood test	Mean (SD)	na	25.0 (0.0)	26.0 (27.9)	22.3 (28.5)	23.8 (27.5)	0.670
		Median (Q1 – Q3)		25.0 (25.0 – 25.0)	12.5 (0.0 – 100.0)	12.5 (0.0 – 100.0)	12.5 (0.0 – 100.0)	
	Impact of child anxiety - Blood test	Mean (SD)	na	25.0 (17.7)	28.4 (31.2)	23.3 (27.2)	23.8 (28.7)	0.784
		Median (Q1 – Q3)		25.0 (12.5 – 37.5)	25.0 (0.0 – 100.0)	12.5 (0.0 – 100.0)	12.5 (0.0 – 100.0)	
Anxiety - Phe levels	Mean (SD)	na	50.0 (35.4)	61.9 (28.7)	54.6 (30.7)	50.6 (34.9)	0.206	
	Median (Q1 – Q3)		50.0 (25.0 – 75.0)	75.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)		
Financial impact of PKU	Mean (SD)	na	75.0 (35.4)	30.0 (30.5)	30.2 (30.0)	18.2 (24.3)	0.004	
	Median (Q1 – Q3)		75.0 (50.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)		
Information on PKU	Mean (SD)	na	37.5 (53.0)	38.1 (25.7)	31.1 (23.3)	26.5 (23.4)	0.032	
	Median (Q1 – Q3)		37.5 (0.0 – 75.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)		
Supplement administration	Adherence to supplements	Mean (SD)	na	0.0 (0.0)	16.8 (24.1)	9.9 (14.9)	10.0 (16.2)	0.224
		Median (Q1 – Q3)		0.0 (0.0 – 0.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 50.0)	0.0 (0.0 – 75.0)	
	Management of supplements	Mean (SD)	na	37.5 (17.7)	15.5 (27.8)	20.8 (29.5)	14.8 (24.5)	0.180
		Median (Q1 – Q3)		37.5 (25.0 – 50.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	
	Practical impact of supplements	Mean (SD)	na	25.0 (11.8)	21.5 (26.3)	22.0 (22.3)	18.1 (23.2)	0.353
Median (Q1 – Q3)			25.0 (16.7 – 33.3)	8.3 (0.0 – 100.0)	16.7 (0.0 – 100.0)	8.3 (0.0 – 83.3)		
Guilt if poor adherence to supplements	Mean (SD)	na	62.5 (53.0)	55.8 (36.3)	53.4 (33.8)	49.3 (32.9)	0.697	
	Median (Q1 – Q3)		62.5 (25.0 – 100.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)		
Relationships within family because of supplements	Mean (SD)	na	12.5 (17.7)	19.2 (27.4)	23.3 (28.4)	14.3 (21.7)	0.246	
	Median (Q1 – Q3)		12.5 (0.0 – 25.0)	0.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)		

Table 2_d : Comparison of the Parent PKU-QOL scores according to patient's overall assessment of health status (cont'd)

Modules	Domains		Poor (N=0)	Fair (N=2)	Good (N=64)	Very good (N=101)	Excellent (N=84)	p-value*
Dietary protein restriction	Adherence to dietary protein restriction	Mean (SD)	na	12.5 (17.7)	20.5 (26.8)	15.9 (28.5)	5.7 (16.0)	0.001
		Median (Q1 – Q3)	na	12.5 (0.0 – 25.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	
	Management of dietary protein restriction	Mean (SD)	na	37.5 (17.7)	31.8 (25.3)	23.4 (22.4)	20.6 (17.4)	0.048
		Median (Q1 – Q3)	na	37.5 (25.0 – 50.0)	25.0 (0.0 – 100.0)	16.7 (0.0 – 95.8)	16.7 (0.0 – 70.8)	
	Practical impact of dietary protein restriction	Mean (SD)	na	34.5 (1.7)	33.2 (20.9)	32.4 (20.0)	30.7 (22.5)	0.860
		Median (Q1 – Q3)	na	34.5 (33.3 – 35.7)	32.1 (0.0 – 82.1)	30.4 (0.0 – 78.6)	28.6 (0.0 – 82.1)	
	Child food enjoyment	Mean (SD)	na	12.5 (17.7)	20.5 (20.8)	21.4 (28.3)	18.7 (21.5)	0.900
		Median (Q1 – Q3)	na	12.5 (0.0 – 25.0)	25.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	25.0 (0.0 – 75.0)	
	Guilt if dietary protein restriction not followed	Mean (SD)	na	50.0 (35.4)	47.6 (36.1)	46.7 (34.8)	41.5 (35.6)	0.716
		Median (Q1 – Q3)	na	50.0 (25.0 – 75.0)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	

*Non-parametric P-value for between-group comparisons: Kruskal-Wallis
 In bold p-value<0.05