

Additional File 3 : Comparison of PKU-QOL scores according to BH4 intake

Table 3_a : Comparison of the Child PKU-QOL scores according to BH4 intake

Modules	Domains		BH4 (N=27)	No BH4 (N=65)	p-value*	
Symptoms	Headaches	Mean (SD)	15.7 (25.1)	15.2 (20.0)	0.778	
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)		
	Stomach aches	Mean (SD)	18.5 (23.6)	14.8 (22.1)	0.483	
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)		
	Tiredness	Mean (SD)	34.3 (29.5)	35.2 (25.2)	0.891	
		Median (Q1 – Q3)	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)		
	Irritability	Mean (SD)	22.2 (33.5)	17.2 (26.0)	0.800	
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)		
	Aggressiveness	Mean (SD)	13.0 (26.3)	11.9 (22.6)	0.936	
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)		
	Moodiness	Mean (SD)	23.1 (25.9)	13.5 (22.2)	0.091	
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)		
	Sadness	Mean (SD)	17.6 (22.8)	15.7 (23.6)	0.655	
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)		
	Anxiety	Mean (SD)	8.7 (18.6)	9.3 (18.2)	0.717	
		Median (Q1 – Q3)	0.0 (0.0 – 50.0)	0.0 (0.0 – 75.0)		
	Lack of concentration	Mean (SD)	17.7 (27.1)	23.7 (28.8)	0.350	
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)		
	Slow thinking	Mean (SD)	24.0 (27.1)	22.3 (25.8)	0.802	
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)		
PKU in general	Emotional impact of PKU	Mean (SD)	32.1 (21.2)	33.3 (22.4)	0.850	
		Median (Q1 – Q3)	33.3 (0.0 – 66.7)	33.3 (0.0 – 83.3)		
	Practical impact of PKU	Mean (SD)	12.0 (19.5)	10.5 (15.7)	0.908	
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)		
	Social impact of PKU	Mean (SD)	18.9 (21.9)	19.6 (16.4)	0.468	
		Median (Q1 – Q3)	16.7 (0.0 – 91.7)	16.7 (0.0 – 75.0)		
	Overall impact of PKU	Mean (SD)	22.1 (16.7)	22.5 (14.8)	0.640	
		Median (Q1 – Q3)	17.2 (0.0 – 71.9)	18.8 (0.0 – 78.1)		
	Anxiety - Blood test	Mean (SD)	13.4 (17.3)	11.9 (16.7)	0.531	
		Median (Q1 – Q3)	12.5 (0.0 – 62.5)	0.0 (0.0 – 50.0)		
	Anxiety - Phe levels	Mean (SD)	53.7 (43.1)	41.1 (33.3)	0.219	
		Median (Q1 – Q3)	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)		
	Supplement administration	Adherence to supplements	Mean (SD)	12.5 (15.9)	10.5 (17.7)	0.366
			Median (Q1 – Q3)	12.5 (0.0 – 50.0)	0.0 (0.0 – 100.0)	
Practical impact of supplements		Mean (SD)	15.2 (29.9)	19.4 (32.1)	0.581	
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)		
Guilt if poor adherence to supplements		Mean (SD)	31.0 (37.0)	50.4 (38.3)	0.040	
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)		
Relationships within family because of supplements		Mean (SD)	12.0 (26.0)	8.5 (19.7)	0.808	
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)		
Taste - Supplements		Mean (SD)	42.0 (27.2)	33.6 (33.5)	0.186	
		Median (Q1 – Q3)	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)		

Table 3_a : Comparison of the Child PKU-QOL scores according to BH4 intake (cont'd)

Modules	Domains		BH4 (N=27)	No BH4 (N=65)	p-value*
Dietary protein restriction	Food temptations	Mean (SD)	37.5 (30.4)	26.4 (26.4)	0.115
		Median (Q1 – Q3)	50.0 (0.0 – 87.5)	25.0 (0.0 – 100.0)	
Dietary protein restriction	Adherence to dietary protein restriction	Mean (SD)	9.9 (18.1)	11.1 (16.3)	0.510
		Median (Q1 – Q3)	0.0 (0.0 – 62.5)	0.0 (0.0 – 62.5)	
	Social impact of dietary protein restriction	Mean (SD)	21.5 (23.3)	17.4 (16.6)	0.819
		Median (Q1 – Q3)	12.5 (0.0 – 80.0)	15.0 (0.0 – 70.0)	
	Taste – low protein food	Mean (SD)	16.2 (19.6)	18.5 (24.4)	0.875
		Median (Q1 – Q3)	0.0 (0.0 – 50.0)	25.0 (0.0 – 100.0)	
	Food enjoyment	Mean (SD)	15.7 (21.0)	17.7 (24.5)	0.808
		Median (Q1 – Q3)	0.0 (0.0 – 50.0)	0.0 (0.0 – 100.0)	
	Guilt if dietary protein restriction not followed	Mean (SD)	44.6 (44.6)	55.0 (39.8)	0.281
		Median (Q1 – Q3)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	
	Overall difficulty following dietary protein restriction	Mean (SD)	24.0 (36.5)	16.5 (23.5)	0.858
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	

*Non-parametric P-value for between-group comparisons: Mann-Whitney-Wilcoxon
 In bold p-value<0.05

Table 3_b : Comparison of the Adolescent PKU-QOL scores according to BH4 intake

Modules	Domains		BH4 (N=27)	No BH4 (N=83)	p-value*
Symptoms	Self-rated status health	Mean (SD)	33.3 (25.9)	35.3 (24.0)	0.680
		Median (Q1 – Q3)	25.0 (0.0 – 75.0)	50.0 (0.0 – 75.0)	
	Headaches	Mean (SD)	18.5 (23.6)	20.5 (26.0)	0.795
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Stomach aches	Mean (SD)	14.8 (24.3)	14.4 (20.0)	0.802
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	0.0 (0.0 – 75.0)	
	Tiredness	Mean (SD)	40.7 (28.7)	40.4 (25.2)	0.911
		Median (Q1 – Q3)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	
	Irritability	Mean (SD)	24.1 (26.4)	25.0 (25.8)	0.849
		Median (Q1 – Q3)	25.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)	
	Aggressiveness	Mean (SD)	12.0 (22.3)	11.9 (22.8)	0.985
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Moodiness	Mean (SD)	16.7 (19.6)	20.9 (23.7)	0.515
		Median (Q1 – Q3)	0.0 (0.0 – 50.0)	25.0 (0.0 – 75.0)	
	Sadness	Mean (SD)	4.6 (12.1)	14.7 (20.9)	0.022
		Median (Q1 – Q3)	0.0 (0.0 – 50.0)	0.0 (0.0 – 75.0)	
	Anxiety	Mean (SD)	13.0 (24.4)	14.4 (23.5)	0.694
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	
	Lack of concentration	Mean (SD)	17.6 (26.7)	20.3 (23.6)	0.424
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	12.5 (0.0 – 75.0)	
	Slow thinking	Mean (SD)	13.0 (18.8)	14.1 (20.2)	0.869
		Median (Q1 – Q3)	0.0 (0.0 – 50.0)	0.0 (0.0 – 75.0)	
PKU in general	Emotional impact of PKU	Mean (SD)	27.1 (15.3)	33.8 (20.9)	0.209
		Median (Q1 – Q3)	30.0 (0.0 – 70.0)	30.0 (0.0 – 90.0)	
	Practical impact of PKU	Mean (SD)	10.0 (12.9)	12.5 (14.2)	0.707
		Median (Q1 – Q3)	8.3 (0.0 – 33.3)	8.3 (0.0 – 50.0)	
	Social impact of PKU	Mean (SD)	11.2 (11.8)	17.1 (15.0)	0.052
		Median (Q1 – Q3)	8.3 (0.0 – 41.7)	16.7 (0.0 – 91.7)	
	Overall impact of PKU	Mean (SD)	17.8 (10.7)	24.4 (16.1)	0.066
		Median (Q1 – Q3)	17.5 (2.3 – 50.0)	22.7 (0.0 – 85.0)	
	Anxiety - Blood test	Mean (SD)	11.1 (21.9)	10.8 (20.4)	0.857
		Median (Q1 – Q3)	0.0 (0.0 - 87.5)	0.0 (0.0 – 100.0)	
	Anxiety - Phe levels	Mean (SD)	35.6 (33.3)	35.6 (31.3)	0.943
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	

Table 3_b : Comparison of the Adolescent PKU-QOL scores according to BH4 intake (cont'd)

Modules	Domains		BH4 (N=27)	No BH4 (N=83)	p-value*
Supplement administration	Adherence to supplements	Mean (SD)	7.6 (9.9)	16.7 (20.5)	0.103
		Median (Q1 – Q3)	3.1 (0.0 – 33.3)	7.3 (0.0 – 83.3)	
	Practical impact of supplements	Mean (SD)	6.6 (8.7)	22.6 (25.2)	0.006
		Median (Q1 – Q3)	3.1 (0.0 – 25.0)	12.5 (0.0 – 100.0)	
	Guilt if poor adherence to supplements	Mean (SD)	42.5 (33.5)	43.6 (36.6)	0.985
		Median (Q1 – Q3)	37.5 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Relationships within family because of supplements	Mean (SD)	10.0 (17.0)	15.5 (27.0)	0.727
		Median (Q1 – Q3)	0.0 (0.0 – 50.0)	0.0 (0.0 – 100.0)	
	Taste - Supplements	Mean (SD)	41.3 (20.3)	40.9 (26.0)	0.644
		Median (Q1 – Q3)	50.0 (0.0 – 75.0)	50.0 (0.0 – 100.0)	
Dietary protein restriction	Food temptations	Mean (SD)	14.6 (18.8)	29.3 (29.3)	0.049
		Median (Q1 – Q3)	0.0 (0.0 – 50.0)	25.0 (0.0 – 100.0)	
	Adherence to dietary protein restriction	Mean (SD)	7.2 (10.6)	15.0 (19.8)	0.124
		Median (Q1 – Q3)	0.0 (0.0 – 33.3)	8.3 (0.0 – 83.3)	
	Practical impact of dietary protein restriction	Mean (SD)	22.8 (13.7)	28.1 (19.0)	0.574
		Median (Q1 – Q3)	20.7 (0.0 – 53.6)	28.6 (0.0 – 85.7)	
	Social impact of dietary protein restriction	Mean (SD)	5.3 (10.9)	16.0 (21.8)	0.031
		Median (Q1 – Q3)	0.0 (0.0 – 45.0)	7.5 (0.0 – 90.0)	
	Overall impact of dietary protein restriction	Mean (SD)	14.7 (11.2)	21.8 (18.0)	0.155
		Median (Q1 – Q3)	13.6 (0.0 – 37.5)	18.2 (0.0 – 86.4)	
	Taste – low protein food	Mean (SD)	22.9 (16.7)	26.4 (26.8)	0.978
		Median (Q1 – Q3)	25.0 (0.0 – 50.0)	25.0 (0.0 – 100.0)	
	Food enjoyment	Mean (SD)	11.8 (21.9)	17.1 (27.6)	0.482
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Guilt if dietary protein restriction not followed	Mean (SD)	34.7 (36.5)	49.3 (34.9)	0.112
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	
	Overall difficulty following dietary protein restriction	Mean (SD)	3.9 (9.4)	20.3 (28.4)	0.010
		Median (Q1 – Q3)	0.0 (0.0 – 25.0)	0.0 (0.0 – 100.0)	

*Non-parametric P-value for between-group comparisons: Mann-Whitney-Wilcoxon
In bold p-value<0.05

Table 3_c : Comparison of the Adult PKU-QOL scores according to BH4 intake

Modules	Domains		BH4 (N=15)	No BH4 (N=89)	p-value*
Symptoms	Self-rated health status	Mean (SD)	28.6 (25.7)	34.3 (22.4)	0.430
		Median (Q1 – Q3)	25.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)	
	Headaches	Mean (SD)	17.9 (20.6)	21.9 (24.2)	0.654
		Median (Q1 – Q3)	12.5 (0.0 – 50.0)	25.0 (0.0 – 75.0)	
	Stomach aches	Mean (SD)	12.5 (23.5)	17.3 (26.1)	0.533
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Tiredness	Mean (SD)	37.5 (27.3)	45.5 (27.5)	0.455
		Median (Q1 – Q3)	50.0 (0.0 – 75.0)	50.0 (0.0 – 100.0)	
	Trembling hands	Mean (SD)	10.7 (18.9)	12.8 (23.7)	0.975
		Median (Q1 – Q3)	0.0 (0.0 – 50.0)	0.0 (0.0 – 100.0)	
	Irritability	Mean (SD)	25.0 (21.9)	34.8 (25.2)	0.180
		Median (Q1 – Q3)	25.0 (0.0 – 50.0)	50.0 (0.0 – 100.0)	
	Aggressiveness	Mean (SD)	5.4 (10.6)	13.6 (22.4)	0.294
		Median (Q1 – Q3)	0.0 (0.0 – 25.0)	0.0 (0.0 – 75.0)	
	Moodiness	Mean (SD)	19.6 (22.3)	23.6 (25.2)	0.650
		Median (Q1 – Q3)	12.5 (0.0 – 50.0)	25.0 (0.0 – 75.0)	
	Sadness	Mean (SD)	26.8 (30.2)	29.3 (28.2)	0.687
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Anxiety	Mean (SD)	23.2 (28.5)	21.9 (29.6)	0.751
		Median (Q1 – Q3)	12.5 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Lack of concentration	Mean (SD)	23.2 (24.9)	24.7 (26.4)	0.914
		Median (Q1 – Q3)	25.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)	
	Slow thinking	Mean (SD)	7.1 (15.3)	19.9 (27.6)	0.098
		Median (Q1 – Q3)	0.0 (0.0 – 50.0)	0.0 (0.0 – 100.0)	
PKU in general	Emotional impact of PKU	Mean (SD)	42.5 (22.3)	44.9 (22.2)	0.927
		Median (Q1 – Q3)	45.0 (0.0 – 70.0)	45.0 (0.0 – 100.0)	
	Practical impact of PKU	Mean (SD)	12.7 (17.0)	18.6 (16.9)	0.204
		Median (Q1 – Q3)	8.3 (0.0 – 58.3)	16.7 (0.0 – 75.0)	
	Social impact of PKU	Mean (SD)	14.1 (9.1)	18.4 (16.2)	0.601
		Median (Q1 – Q3)	16.7 (0.0 – 31.3)	16.7 (0.0 – 68.8)	
	Overall impact of PKU	Mean (SD)	22.4 (12.0)	30.9 (15.7)	0.373
		Median (Q1 – Q3)	24.0 (8.3 – 33.3)	27.1 (2.1 – 68.8)	
	Anxiety - Blood test	Mean (SD)	9.6 (11.6)	10.8 (23.4)	0.308
		Median (Q1 – Q3)	0.0 (0.0 – 25.0)	0.0 (0.0 – 100.0)	
	Anxiety - Phe levels	Mean (SD)	39.3 (21.3)	36.2 (28.2)	0.559
		Median (Q1 – Q3)	37.5 (0.0 – 75.0)	25.0 (0.0 – 100.0)	
	Anxiety - Phe levels during pregnancy	Mean (SD)	80.6 (24.3)	80.8 (24.8)	0.959
		Median (Q1 – Q3)	100.0 (50.0 – 100.0)	100.0 (0.0 – 100.0)	
	Financial impact of PKU	Mean (SD)	3.6 (9.1)	21.3 (28.3)	0.013
		Median (Q1 – Q3)	0.0 (0.0 – 25.0)	0.0 (0.0 – 100.0)	
	Information on PKU	Mean (SD)	30.4 (20.0)	34.8 (24.9)	0.570
		Median (Q1 – Q3)	25.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)	

Table 3_c : Comparison of the Adult PKU-QOL scores according to BH4 intake (cont'd)

Modules	Domains		BH4 (N=15)	No BH4 (N=89)	p-value*
Supplement administration	Adherence to supplements	Mean (SD)	11.1 (9.6)	20.8 (20.4)	0.567
		Median (Q1 – Q3)	16.7 (0.0 – 16.7)	16.7 (0.0 – 66.7)	
	Practical impact of supplements	Mean (SD)	9.4 (10.8)	22.6 (21.7)	0.257
		Median (Q1 – Q3)	6.3 (0.0 – 25.0)	18.8 (0.0 – 100.0)	
	Guilt if poor adherence to supplements	Mean (SD)	50.0 (28.9)	47.8 (33.3)	0.886
		Median (Q1 – Q3)	50.0 (25.0 – 75.0)	50.0 (0.0 – 100.0)	
	Relationships within family because of supplements	Mean (SD)	18.8 (37.5)	10.8 (21.4)	0.891
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Taste - Supplements	Mean (SD)	50.0 (20.4)	49.7 (25.8)	0.926
		Median (Q1 – Q3)	50.0 (25.0 – 75.0)	50.0 (0.0 – 100.0)	
Dietary protein restriction	Food temptations	Mean (SD)	12.5 (12.5)	36.7 (24.8)	0.027
		Median (Q1 – Q3)	12.5 (0.0 – 25.0)	37.5 (0.0 – 100.0)	
	Adherence to dietary protein restriction	Mean (SD)	4.7 (9.4)	20.6 (18.2)	0.064
		Median (Q1 – Q3)	0.0 (0.0 – 18.8)	20.0 (0.0 – 56.3)	
	Social impact of dietary protein-restriction	Mean (SD)	2.7 (3.8)	19.0 (18.6)	0.016
		Median (Q1 – Q3)	0.0 (0.0 – 8.3)	15.8 (0.0 – 79.2)	
	Practical impact of dietary protein restriction	Mean (SD)	15.5 (14.5)	36.5 (19.6)	0.019
		Median (Q1 – Q3)	12.5 (0.0 – 39.3)	35.7 (0.0 – 75.0)	
	Overall impact of dietary protein restriction	Mean (SD)	9.5 (9.4)	27.7 (17.1)	0.016
		Median (Q1 – Q3)	6.8 (0.0 – 25.0)	26.9 (0.0 – 75.0)	
	Taste – low protein food	Mean (SD)	50.0 (.)	33.1 (21.4)	0.292
		Median (Q1 – Q3)	50.0 (50.0 – 50.0)	25.0 (0.0 – 100.0)	
	Food enjoyment	Mean (SD)	10.0 (13.7)	27.2 (30.0)	0.223
		Median (Q1 – Q3)	0.0 (0.0 – 25.0)	25.0 (0.0 – 100.0)	
	Guilt if dietary protein restriction not followed	Mean (SD)	65.0 (28.5)	53.2 (31.9)	0.415
		Median (Q1 – Q3)	75.0 (25.0 – 100.0)	50.0 (0.0 – 100.0)	
	Overall difficulty following dietary protein restriction	Mean (SD)	16.7 (30.3)	29.1 (27.9)	0.232
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)	

*Non-parametric P-value for between-group comparisons: Mann-Whitney-Wilcoxon
 In bold p-value<0.05

Table 3_d : Comparison of the Parent PKU-QOL scores according to patient's BH4 intake

Modules	Domains		BH4 (N=67)	No BH4 (N=186)	p-value*
Symptoms	Child health status	Mean (SD)	28.0 (22.4)	32.5 (24.4)	0.221
		Median (Q1 – Q3)	25.0 (0.0 – 75.0)	25.0 (0.0 – 100.0)	
	Headaches	Mean (SD)	13.8 (21.0)	14.8 (20.7)	0.624
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 75.0)	
	Stomach aches	Mean (SD)	23.5 (26.8)	17.3 (23.2)	0.107
		Median (Q1 – Q3)	25.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Tiredness	Mean (SD)	32.1 (25.3)	31.9 (24.9)	0.926
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Irritability	Mean (SD)	38.1 (28.7)	35.2 (27.1)	0.523
		Median (Q1 – Q3)	50.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Aggressiveness	Mean (SD)	14.4 (22.0)	12.4 (21.8)	0.443
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Moodiness	Mean (SD)	34.1 (27.6)	25.3 (24.6)	0.023
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Sadness	Mean (SD)	18.2 (22.6)	18.2 (20.8)	0.819
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Anxiety	Mean (SD)	14.8 (21.7)	14.0 (21.9)	0.708
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)	
	Lack of concentration	Mean (SD)	34.1 (28.6)	29.0 (28.1)	0.192
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
Slow thinking	Mean (SD)	20.8 (25.8)	19.6 (26.8)	0.611	
	Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)		
PKU in general	Emotional impact of PKU	Mean (SD)	40.9 (25.3)	43.1 (22.1)	0.381
		Median (Q1 – Q3)	43.8 (0.0 – 93.8)	37.5 (0.0 – 100.0)	
	Practical impact of PKU	Mean (SD)	11.4 (12.9)	15.3 (14.8)	0.061
		Median (Q1 – Q3)	4.2 (0.0 – 41.7)	12.5 (0.0 – 70.0)	
	Social impact of PKU	Mean (SD)	15.8 (15.9)	17.5 (17.0)	0.346
		Median (Q1 – Q3)	10.0 (0.0 – 55.0)	12.5 (0.0 – 90.0)	
	Overall impact of PKU	Mean (SD)	21.4 (15.3)	24.7 (15.5)	0.083
		Median (Q1 – Q3)	18.8 (0.0 – 56.7)	21.7 (0.0 – 76.9)	
	Child anxiety - Blood test	Mean (SD)	23.3 (27.4)	23.7 (28.0)	0.981
		Median (Q1 – Q3)	12.5 (0.0 – 100.0)	12.5 (0.0 – 100.0)	
	Impact of child anxiety - Blood test	Mean (SD)	21.9 (27.1)	25.6 (29.1)	0.386
		Median (Q1 – Q3)	12.5 (0.0 – 100.0)	12.5 (0.0 – 100.0)	
	Anxiety - Phe levels	Mean (SD)	60.2 (33.7)	52.9 (31.0)	0.113
		Median (Q1 – Q3)	75.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)	
	Financial impact of PKU	Mean (SD)	24.2 (28.6)	27.2 (29.1)	0.428
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	
	Information on PKU	Mean (SD)	27.3 (20.0)	32.6 (25.7)	0.178
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)	

Table 3_d : Comparison of the Parent PKU-QOL scores according to patient's BH4 intake (cont'd)

Modules	Domains		BH4 Mean (SD) (N=67)	No BH4 Mean (SD) (N=186)	p-value*	
Supplement administration	Adherence to supplements	Mean (SD)	9.6 (13.3)	12.4 (19.3)	0.681	
		Median (Q1 – Q3)	0.0 (0.0 – 50.0)	0.0 (0.0 – 100.0)		
	Management of supplements	Mean (SD)	12.8 (26.0)	18.8 (27.7)	0.077	
		Median (Q1 – Q3)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)		
	Practical impact of supplements	Mean (SD)	16.8 (24.3)	21.6 (23.2)	0.113	
		Median (Q1 – Q3)	8.3 (0.0 – 100.0)	16.7 (0.0 – 100.0)		
	Guilt if poor adherence to supplements	Mean (SD)	51.6 (33.9)	52.7 (34.2)	0.849	
		Median (Q1 – Q3)	50.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)		
	Relationships within family because of supplements	Mean (SD)	15.1 (24.0)	20.6 (26.8)	0.189	
		Median (Q1 – Q3)	0.0 (0.0 – 75.0)	0.0 (0.0 – 100.0)		
	Dietary protein restriction	Adherence to dietary protein restriction	Mean (SD)	8.7 (20.1)	15.1 (26.2)	0.102
			Median (Q1 – Q3)	0.0 (0.0 – 100.0)	0.0 (0.0 – 100.0)	
Management of dietary protein restriction		Mean (SD)	24.0 (21.6)	24.8 (22.2)	0.906	
		Median (Q1 – Q3)	20.8 (0.0 – 83.3)	20.8 (0.0 – 100.0)		
Practical impact of dietary protein restriction		Mean (SD)	29.7 (18.1)	32.5 (21.5)	0.537	
		Median (Q1 – Q3)	28.6 (0.0 – 64.3)	28.6 (0.0 – 82.1)		
Child food enjoyment		Mean (SD)	19.7 (21.5)	20.5 (25.0)	0.812	
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	25.0 (0.0 – 100.0)		
Guilt if dietary protein restriction not followed		Mean (SD)	38.0 (34.7)	47.3 (35.1)	0.091	
		Median (Q1 – Q3)	25.0 (0.0 – 100.0)	50.0 (0.0 – 100.0)		

*Non-parametric P-value for between-group comparisons: Mann-Whitney-Wilcoxon
In bold p-value<0.05