Supplementary Material: Determining the Relationship between Seizure days, Seizure-Free Days and Other Predictors of Health-Related Quality of Life in Patients with Dravet Syndrome and Their Carers from FFA registration studies

Authors

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Tel: +44 (0)207 993 2930 Email: <u>Elisabeth.adams@aquariusph.com</u> Table S1. Testing interaction terms following the univariate analysis for patient PedsQL.

Covariate	Coefficient	p-value	Outcome variable
Age*28-day frequency of	-0.005	>0.1	PedsQL
seizure days at baseline			baseline
Age*28-day frequency of	0.005	>0.1	PedsQL
SFDs at baseline			baseline
Age * Study	-0.157	>0.5	PedsQL
			baseline
Age * Motor impairments	0.775	>0.1	PedsQL
	0.913		baseline
28-day frequency of	0.045	>0.1	PedsQL
seizure days at baseline			baseline
*Study			
28-day frequency of SFDs	-0.045	>0.1	PedsQL
at baseline * Study			baseline
28-day frequency of	0.003	>0.1	PedsQL
seizure days at baseline *	-0.128	>0.1	baseline
Motor impairments			
28-day frequency of SFDs	-0.003	>0.1	PedsQL
at baseline* Motor	0.128	>0.1	baseline
impairments			

 Table S2. Testing interaction terms following the univariate analysis for patient EQ-5D-Y data at baseline

Covariate	Coefficient	p-value	Outcome
			variable
Age* 28-day frequency of	0.027	>0.1	EQ-5D-Y
seizure days at baseline			baseline
Age* 28-day frequency of	-0.027	>0.1	EQ-5D-Y
SFDs at baseline			baseline
Age * Study	-0.338	>0.1	EQ-5D-Y
			baseline
Age * Motor impairments	1.238	>0.1	EQ-5D-Y
	1.736		baseline
28-day frequency of seizure	0.220	>0.1	EQ-5D-Y
days at baseline * Study			baseline
28-day frequency of SFDs at	-0.220	>0.1	EQ-5D-Y
baseline * Study			baseline
28-day frequency of seizure	0.037	>0.1	EQ-5D-Y
days at baseline * Motor	-0.031	>0.1	baseline
impairments			
28-day frequency of SFDs at	-0.037	>0.1	EQ-5D-Y
baseline * Motor	0.031	>0.1	baseline
impairments			

[†]For EQ-5D-Y coefficients refer to a 0-100 scale. All utility values predicted using these coefficients were divided by 100 before the predicted relationship was estimated (shown in Figure 1).

 Table S3. Stepwise testing of covariates with the random effects components of each model

 analysis for patient PedSQL and EQ-5D-Y data at the 3 follow-up time points

Covariate	Coefficient	p-value	Outcome
			variable
age + (1 variable) + (1 subjid)	-0.746	<0.001	PedsQL
seizure day 28 freq +	-0.344	<0.001	PedsQL
(1 variable) + (1 subjid)			
seizure free day 28 freq +	0.344	<0.001	PedsQL
(1 variable) + (1 subjid)			
Study + (1 variable) +	0.914	>0.5	PedsQL
(1 subjid)			
physicalissues + (1 variable) +	-4.098	<0.05	PedsQL
(1 subjid)	-10.476	<0.05	
age + (1 variable) + (1 subjid)	-0.767	<0.05	EQ-5D-Y
seizure day 28 freq +	-0.647	<0.001	EQ-5D-Y
(1 variable) + (1 subjid)			
seizure free day 28 freq +	0.647	<0.001	EQ-5D-Y
(1 variable) + (1 subjid)			
Study + (1 variable) +	1.409	>0.1	EQ-5D-Y
(1 subjid)			
physicalissues + (1 variable) +	-7.458	<0.01	EQ-5D-Y
(1 subjid)	-15.333	<0.05	

[†]For EQ-5D-Y coefficients refer to a 0-100 scale. All utility values predicted using these coefficients were divided by 100 before the predicted relationship was estimated (shown in Figure 1).

Table S4. Testing interaction terms following the univariate analysis for carer EQ-5D-3L data atbaseline

Covariate	Coefficient	p-value	Outcome variable
Age* 28-day frequency of seizure	-0.130	<0.05	EQ-5D-3L
days at baseline			baseline
Age* 28-day frequency of SFDs at	-0.130	<0.05	EQ-5D-3L
baseline			baseline
Age * Study	-1.451	>0.1	EQ-5D-3L
			baseline
Age * Motor impairments	0.465	>0.1	EQ-5D-3L
	0.919		baseline
28-day frequency of seizure days	-0.268	>0.1	EQ-5D-3L
at baseline * Study			baseline
28-day frequency of SFDs at	0.268	>0.1	EQ-5D-3L
baseline * study			baseline
28-day frequency of seizure days	0.366	>0.1	EQ-5D-3L
at baseline * Motor impairments	0.005	>0.1	baseline
28-day frequency of SFDs at	-0.366	>0.1	EQ-5D-3L
baseline * Motor impairments	-0.005	>0.1	baseline

⁺Coefficients refer to a 0-100 scale. All utility values predicted using these coefficients were divided by 100 before the predicted relationship was estimated

 Table S5. Stepwise testing of covariates with the random effects components of each model

 analysis for carer EQ-5D-3L data at the 3 follow-up time points

Covariate	Coefficient	p-value	Outcome
			variable
age + (1 variable) + (1 subjid)	-0.728	>0.05	EQ-5D-3L
seizure day 28 freq +	-0.473	<0.05	EQ-5D-3L
(1 variable) + (1 subjid)			
seizure free day 28 freq +	0.473	<0.05	EQ-5D-3L
(1 variable) + (1 subjid)			
Study + (1 variable) +	1.770	>0.1	EQ-5D-3L
(1 subjid)			
physicalissues + (1 variable) +	-26.821	<0.001	EQ-5D-3L
(1 subjid)	-14.998	>0.1	

⁺Coefficients refer to a 0-100 scale. All utility values predicted using these coefficients were divided by 100 before the predicted relationship was estimated

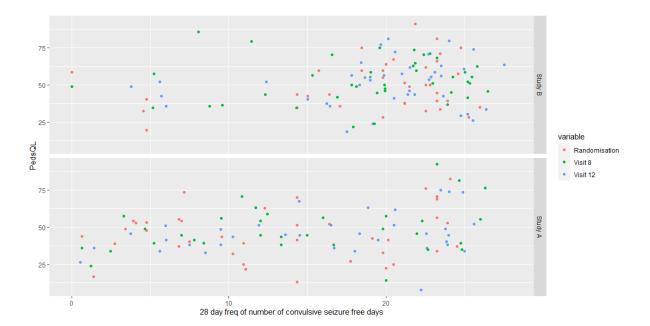
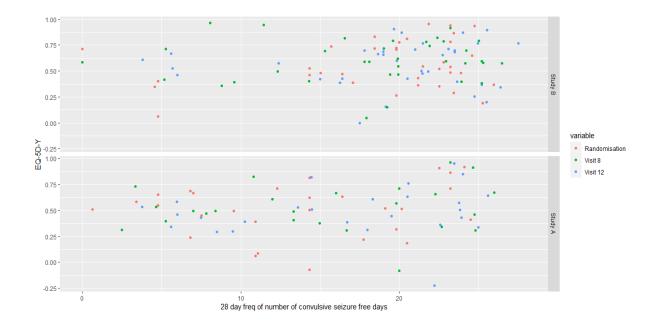


Figure S1. PedsQL data for 128 patients in both study groups of the trial for each follow-up point.

Figure S2. EQ-5D-Y data for 128 patients in both study groups of the trial for each follow-up point.



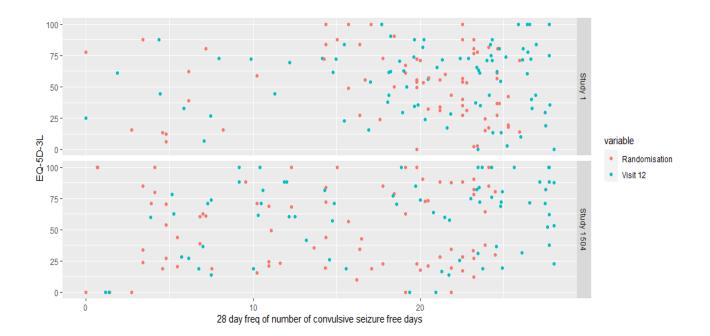


Figure S3. EQ-5D-3L data for 176 carers in both study groups of the trial for each follow-up point.