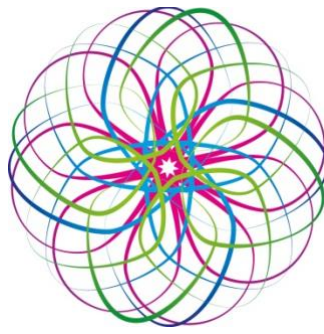




CHDR
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Statistical appendix: Analysis of pharmacodynamic parameters CHDR1939-B

Randomized, double-blind, placebo-controlled, two way crossover, single centre study evaluating the acute and chronic effect of clonazepam on cognitive tests and patientreported outcome measures in patients with ARID1B-related intellectual disability

PROTOCOL MANAGEMENT

CHDR protocol ID	CHDR1939-B
Short title	Clonazepam in ARID1B Evaluation (CARE study)
Sponsor protocol ID	
File name	CHDR1939-B_FPD_2023-01-18_CGI
Date	2023-01-18

Investigator site

Responsible investigator	R. (Rob) Zuiker, MD, PhD
Co-investigator(s)	G. (Gijs) Santen, MD, PhD; K. (Koshar) Safai Pour, MD; P. (Eline) van der Sluijs, MD; C. (Cecile) Berends. MSc
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Sponsor

Company name	Centre for Human Drug Research
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Description of the analysis

Changes in data

This report focuses on the **CGI** Questionnaire data.

The CGI Severity and Improvement score scales are as follows:

Severity scale	Description
0	Normal not at all ill
1	Borderline ill
2	Mildly ill
3	Moderately ill
4	Markedly ill
5	Severely ill
6	Among the most extremely ill patients
Improvement scale	Description
0	Very much improved
1	Much improved
2	Minimally improved
3	No change
4	Minimally worse
5	Much worse
6	Very much worse

Calculations

For every corresponding parent/teacher, Clinician Global Impression Score (CGI) is a combination of severity score (CGI-S) taken on day 1 and improvement score (CGI-I) assessed on day 22.

Analysis

All CGI variables are listed by subject, treatment, visit and time, and individual plots per treatment by time are generated. These variables are also summarized by frequency tables and graphs.

For every CGI variable, the improvement score is used as outcome and analysed using mixed model analysis of variance with fixed factors treatment and period (occasion), with random factor subject and CGI severity score as covariate.

The following contrast is calculated within the models:

- Clonazepam versus Placebo on day 22

Finally the full ANOVA SAS output is added, as is the used SAS program.

All calculations were performed using SAS for windows V9.4 (SAS Institute, Inc., Cary, NC, USA).

Individual pharmacodynamic response data

DRAFT

Listing 1 Parents: Clinical Global Impression of Severity**Listing 1 Parents: Clinical Global Impression of Severity**
Safety population

Subject	Visit	Treatment	Date	Time (hh:mm)	Time from dosing		Parents: Clinical Global Impression of Severity
					Protocol (hh:mm)	Actual (hh:mm)	
201	1	Placebo	11SEP2021	9:31	-0:08	-1:27	3
	2	Clonazepam	23OCT2021	14:49	-0:08	4:18	3
202	1	Clonazepam	11SEP2021	M	-0:08		3
	2	Placebo	23OCT2021	15:00	-0:08	4:01	3
203	1	Placebo	11SEP2021	M	-0:08		1
	2	Clonazepam	23OCT2021	15:42	-0:08	3:34	2
204	1	Clonazepam	25SEP2021	M	-0:08		1
	2	Placebo	06NOV2021	15:05	-0:08	4:35	2
205	1	Placebo	25SEP2021	M	-0:08		3
	2	Clonazepam	06NOV2021	15:19	-0:08	4:12	3
206	1	Clonazepam	25SEP2021	M	-0:08		2
	2	Placebo	21NOV2021	14:55	-0:08	4:50	3
207	1	Clonazepam	09OCT2021	13:52	-0:08	2:50	3
208	1	Placebo	10OCT2021	14:49	-0:08	3:18	3
	2	Clonazepam	21NOV2021	15:53	-0:08	5:05	3
209	1	Clonazepam	10OCT2021	14:48	-0:08	4:22	2
	2	Placebo	21NOV2021	16:33	-0:08	5:25	2
210	1	Placebo	10OCT2021	14:47	-0:08	3:35	3
	2	Clonazepam	21NOV2021	15:54	-0:08	5:17	3
211	1	Placebo	12MAR2022	16:32	-0:08	3:52	1
	2	Clonazepam	14MAY2022	11:33	-0:08	1:02	1
212	1	Clonazepam	29JAN2022	14:48	-0:08	4:13	2
	2	Placebo	12MAR2022	14:43	-0:08	4:48	2
213	1	Clonazepam	29JAN2022	15:14	-0:08	4:25	4
	2	Placebo	12MAR2022	14:45	-0:08	4:40	3
214	1	Placebo	29JAN2022	16:00	-0:08	4:39	3

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Listing 1 Parents: Clinical Global Impression of Severity
Safety population

Subject	Visit	Treatment	Date	Time (hh:mm)	Time from dosing		Parents: Clinical Global Impression of Severity
					Protocol (hh:mm)	Actual (hh:mm)	
215	2	Clonazepam	12MAR2022	15:03	-0:08	3:46	2
	1	Clonazepam	12MAR2022	16:33	-0:08	3:28	1
216	2	Placebo	23APR2022	12:32	-0:08	2:13	1
	1	Placebo	12MAR2022	16:50	-0:08	4:38	4
	2	Clonazepam	23APR2022	12:32	-0:08	2:49	4

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Listing 2 Parents: Clinical Global Impression of Improvement**Listing 2 Parents: Clinical Global Impression of Improvement
Safety population**

Subject	Visit	Treatment	Date	Time (hh:mm)	Time from dosing		Parents: Clinical Global Impression of Improvement
					Protocol (hh:mm)	Actual (hh:mm)	
201	1	Placebo	02OCT2021	14:15	503:52	507:17	3
	2	Clonazepam	13NOV2021	15:09	503:52	508:38	3
202	1	Clonazepam	02OCT2021	14:36	503:52	506:11	2
	2	Placebo	13NOV2021	15:10	503:52	508:11	3
203	1	Placebo	02OCT2021	14:50	503:52	506:14	3
	2	Clonazepam	13NOV2021	15:46	503:52	507:38	4
204	1	Clonazepam	16OCT2021	16:00	503:52	507:45	2
	2	Placebo	27NOV2021	14:53	503:52	508:23	1
205	1	Placebo	16OCT2021	17:00	503:52	509:07	3
	2	Clonazepam	27NOV2021	15:44	503:52	508:37	4
206	1	Clonazepam	16OCT2021	17:02	503:52	508:17	4
	2	Placebo	12DEC2021	15:02	503:52	508:57	3
207	1	Clonazepam	M	M	503:52	M	M
208	1	Placebo	31OCT2021	M	503:52		3
	2	Clonazepam	12DEC2021	15:02	503:52	508:14	0
209	1	Clonazepam	31OCT2021	M	503:52		2
	2	Placebo	12DEC2021	15:02	503:52	507:54	3
210	1	Placebo	31OCT2021	M	503:52		3
	2	Clonazepam	12DEC2021	15:27	503:52	508:50	5
211	1	Placebo	02APR2022	15:57	503:52	507:17	3
	2	Clonazepam	03JUN2022	15:05	503:52	484:34	4
212	1	Clonazepam	19FEB2022	14:51	503:52	508:16	1
	2	Placebo	02APR2022	13:05	503:52	507:10	3
213	1	Clonazepam	19FEB2022	15:20	503:52	508:31	3
	2	Placebo	02APR2022	12:53	503:52	506:48	0
214	1	Placebo	19FEB2022	14:06	503:52	506:45	3

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**Listing 2 Parents: Clinical Global Impression of Improvement
Safety population**

Subject	Visit	Treatment	Date	Time from dosing			Parents: Clinical Global Impression of Improvement
				Time (hh:mm)	Protocol (hh:mm)	Actual (hh:mm)	
215	2	Clonazepam	02APR2022	11:45	503:52	504:28	2
	1	Clonazepam	02APR2022	16:37	503:52	507:32	5
216	2	Placebo	14MAY2022	11:45	503:52	505:26	3
	1	Placebo	02APR2022	16:37	503:52	508:25	3
	2	Clonazepam	12MAY2022	12:28	503:52	458:45	2

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Listing 3 Teacher: Clinical Global Impression of Severity**Listing 3 Teacher: Clinical Global Impression of Severity**
Safety population

Subject	Visit	Treatment	Date Time (hh:mm)		Time from dosing		Teacher: Clinical Global Impression of Severity
					Protocol (hh:mm)	Actual (hh:mm)	
201	1	Placebo	M	M	24:06	M	M
	2	Clonazepam	02DEC2021	10:04	24:06	959:33	3
202	1	Clonazepam	M	M	24:06	M	M
	2	Placebo	25OCT2021	10:35	24:06	47:36	3
203	1	Placebo	M	M	24:06	M	M
	2	Clonazepam	26OCT2021	13:31	24:06	73:23	2
204	1	Clonazepam	M	M	24:06	M	M
	2	Placebo	02DEC2021	10:25	24:06	623:55	2
205	1	Placebo	M	M	24:06	M	M
	2	Clonazepam	01DEC2021	11:14	24:06	600:07	3
206	1	Clonazepam	M	M	24:06	M	M
	2	Placebo	M	M	24:06	M	M
207	1	Clonazepam	M	M	24:06	M	M
208	1	Placebo	09NOV2021	16:50	24:06	725:19	3
	2	Clonazepam	M	M	24:06	M	M
209	1	Clonazepam	11NOV2021	15:02	24:06	772:36	2
	2	Placebo	M	M	24:06	M	M
210	1	Placebo	10NOV2021	16:11	24:06	748:59	3
	2	Clonazepam	M	M	24:06	M	M
211	1	Placebo	14MAR2022	12:50	24:06	48:10	1
	2	Clonazepam	18MAY2022	11:38	24:06	97:07	1
212	1	Clonazepam	03FEB2022	16:59	24:06	126:24	2
	2	Placebo	14MAR2022	14:40	24:06	52:45	2
213	1	Clonazepam	08FEB2022	15:01	24:06	244:12	4
	2	Placebo	14MAR2022	14:41	24:06	52:36	4
214	1	Placebo	03FEB2022	16:35	24:06	125:14	3

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Listing 3 Teacher: Clinical Global Impression of Severity
Safety population

Subject	Visit	Treatment	Date	Time (hh:mm)	Time from dosing		Teacher: Clinical Global Impression of Severity
					Protocol (hh:mm)	Actual (hh:mm)	
215	2	Clonazepam	14MAR2022	14:00	24:06	50:43	3
	1	Clonazepam	17MAR2022	M	24:06		2
	2	Placebo	18MAY2022	11:39	24:06	601:20	2
216	1	Placebo	11MAY2022	12:33	24:06	1440	4
	2	Clonazepam	11MAY2022	12:33	24:06	434:50	4

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Listing 4 Teacher: Clinical Global Impression of Improvement**Listing 4 Teacher: Clinical Global Impression of Improvement
Safety population**

Subject	Visit	Treatment	Date	Time (hh:mm)	Time from dosing		Teacher: Clinical Global Impression of Improvement
					Protocol (hh:mm)	Actual (hh:mm)	
201	1	Placebo	21OCT2021	15:39	528:05	964:41	3
	2	Clonazepam	02DEC2021	9:45	528:05	959:14	5
202	1	Clonazepam	25OCT2021	10:35	528:05	1054	2
	2	Placebo	18NOV2021	16:14	528:05	629:15	3
203	1	Placebo	M	M	528:05	M	M
	2	Clonazepam	01DEC2021	10:47	528:05	934:39	3
204	1	Clonazepam	M	M	528:05	M	M
	2	Placebo	02DEC2021	10:23	528:05	623:53	2
205	1	Placebo	M	M	528:05	M	M
	2	Clonazepam	01DEC2021	11:15	528:05	600:08	6
206	1	Clonazepam	25OCT2021	10:00	528:05	717:15	4
	2	Placebo	18DEC2021	13:15	528:05	651:10	3
207	1	Clonazepam	M	M	528:05	M	M
208	1	Placebo	09NOV2021	16:50	528:05	725:19	3
	2	Clonazepam	15DEC2021	15:00	528:05	580:12	1
209	1	Clonazepam	11NOV2021	15:02	528:05	772:36	3
	2	Placebo	21DEC2021	12:00	528:05	720:52	3
210	1	Placebo	10NOV2021	16:11	528:05	748:59	3
	2	Clonazepam	16DEC2021	16:00	528:05	605:23	M
211	1	Placebo	07APR2022	14:00	528:05	625:20	3
	2	Clonazepam	22JUN2022	16:50	528:05	942:19	4
212	1	Clonazepam	M	M	528:05	M	1
	2	Placebo	08APR2022	14:00	528:05	652:05	3
213	1	Clonazepam	04MAR2022	16:15	528:05	821:26	3
	2	Placebo	07APR2022	12:00	528:05	625:55	0
214	1	Placebo	22FEB2022	12:30	528:05	577:09	3

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Listing 4 Teacher: Clinical Global Impression of Improvement
Safety population

Safety population							
				Time from dosing		Teacher: Clinical Global Impression of Improvement	
Subject	Visit	Treatment	Date	Time (hh:mm)	Protocol (hh:mm)		Actual (hh:mm)
215	2	Clonazepam	06APR2022	12:00	528:05	600:43	4
	1	Clonazepam	08APR2022	16:00	528:05	650:55	4
216	2	Placebo	18MAY2022	11:39	528:05	601:20	3
	1	Placebo	11MAY2022	12:33	528:05	1440	3
	2	Clonazepam	11MAY2022	12:33	528:05	434:50	1

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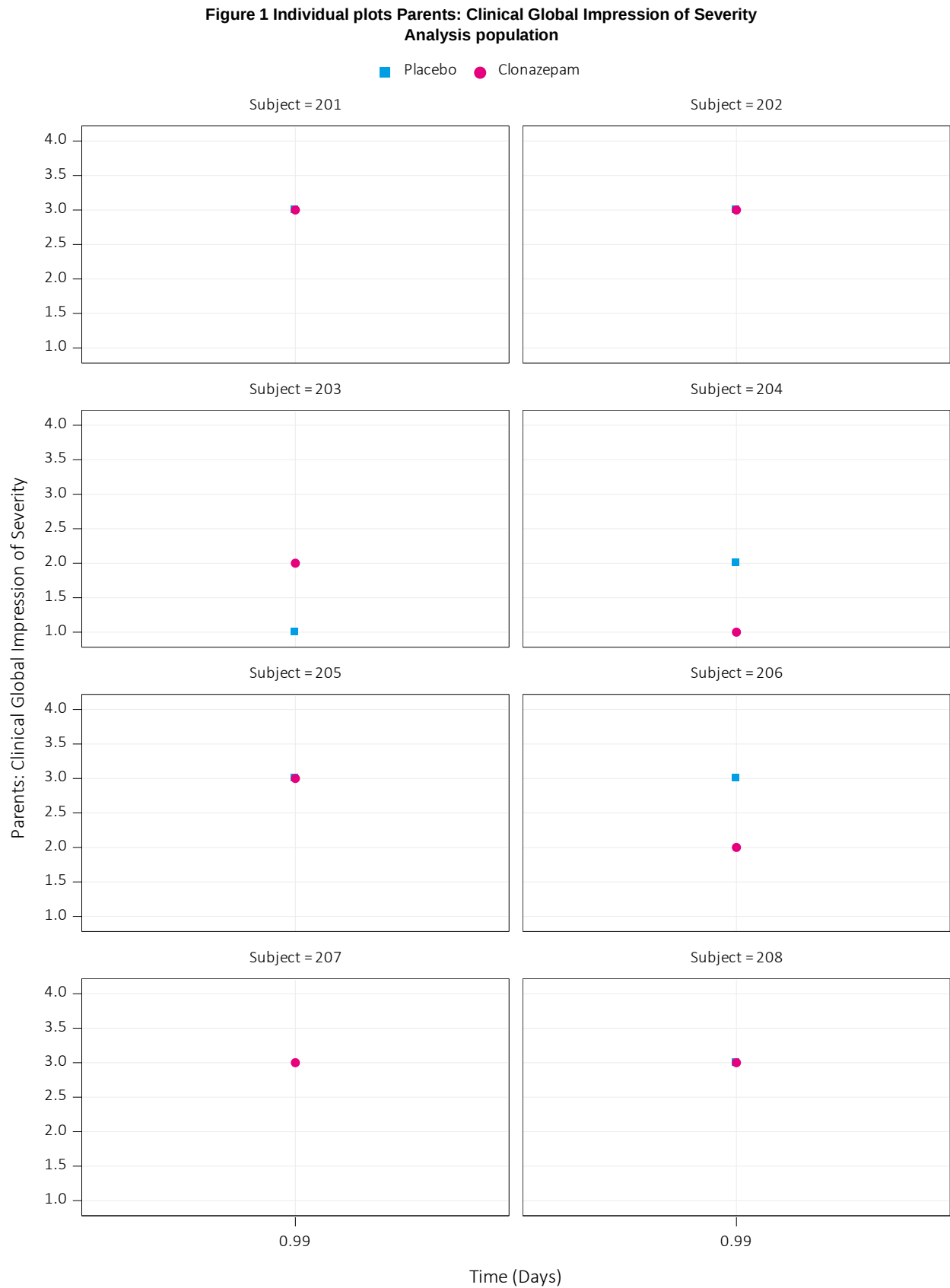
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Individual pharmacodynamic response plots

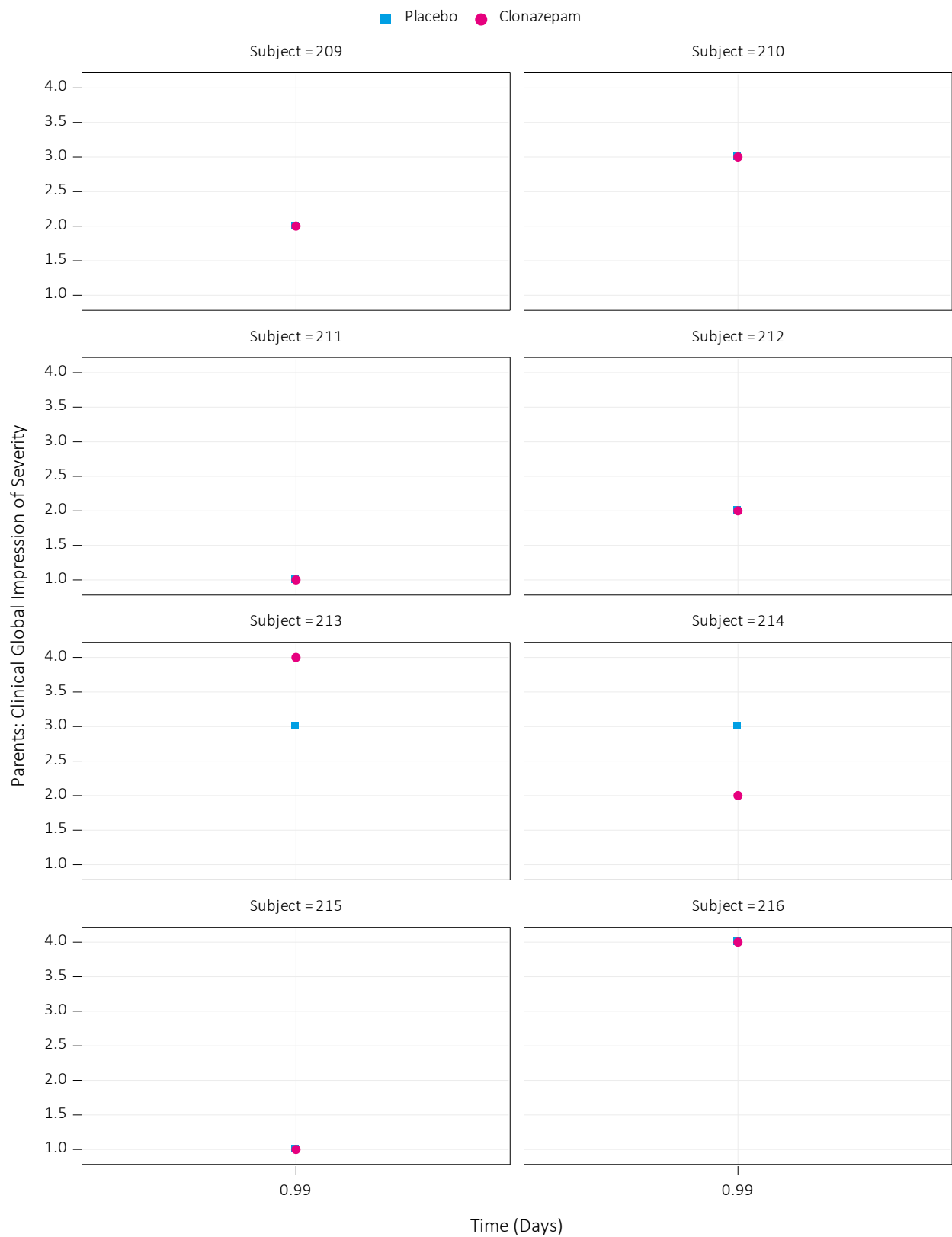
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Figure 1 Individual plots Parents: Clinical Global Impression of Severity



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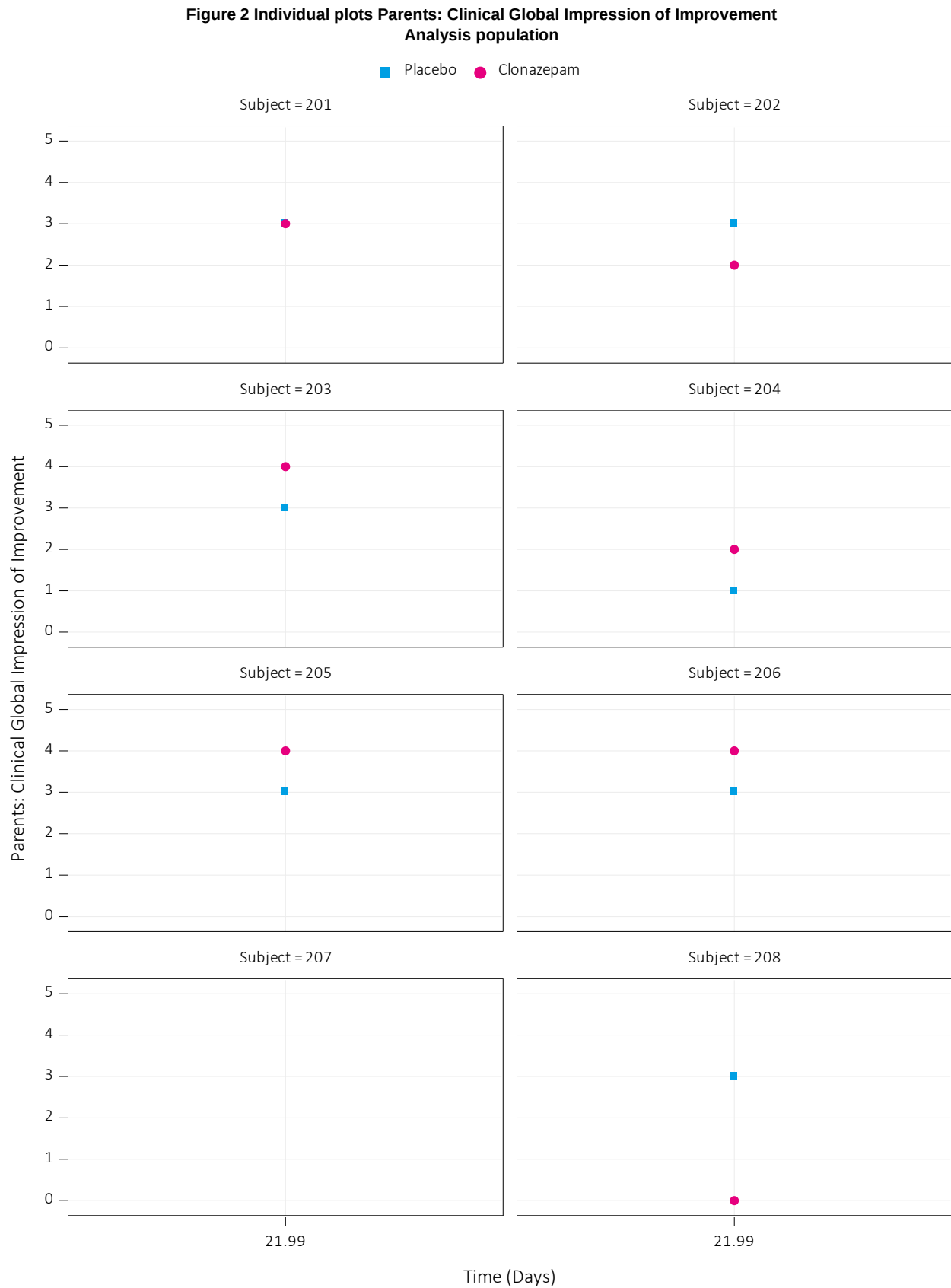
**Figure 1 Individual plots Parents: Clinical Global Impression of Severity
Analysis population**



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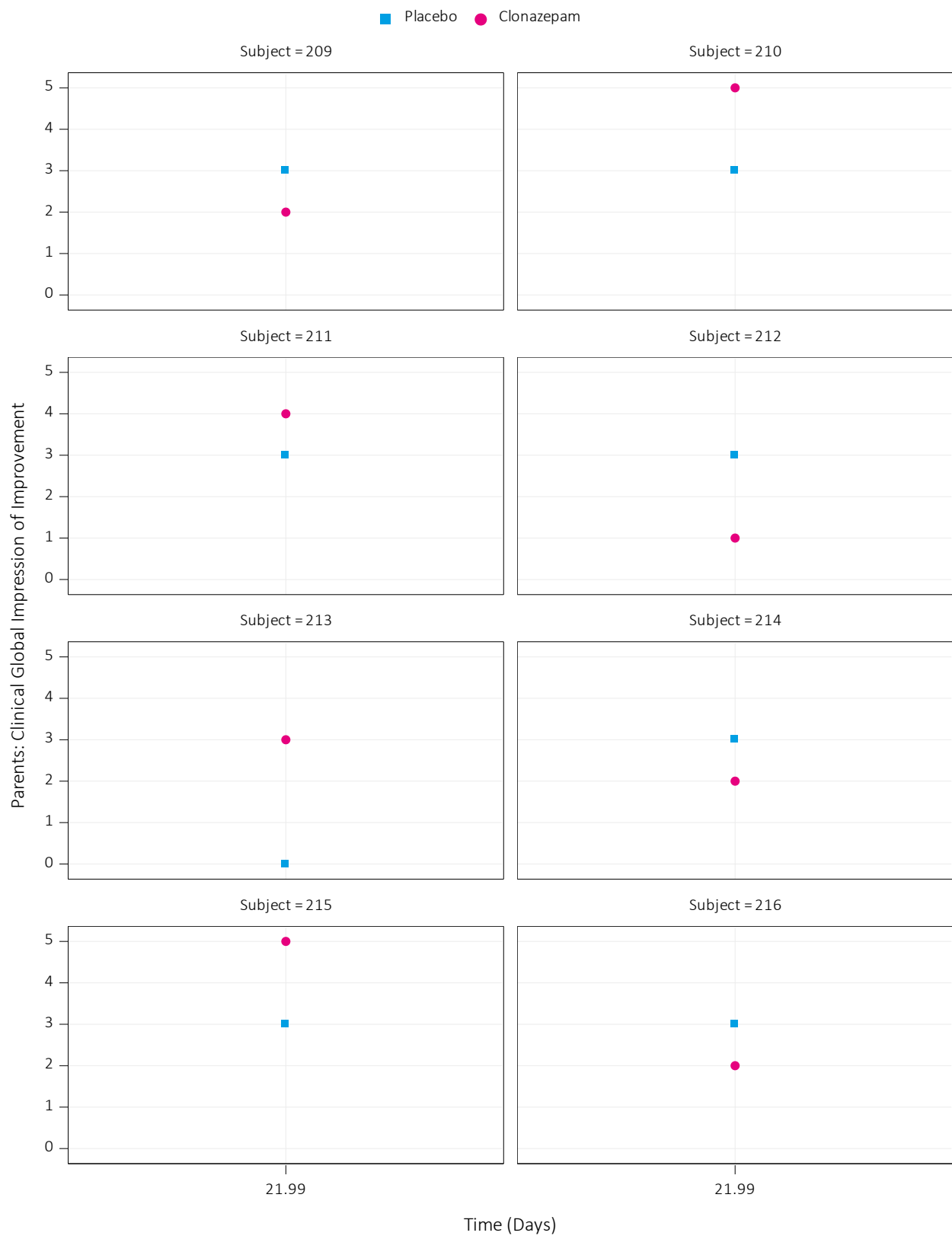
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Figure 2 Individual plots Parents: Clinical Global Impression of Improvement



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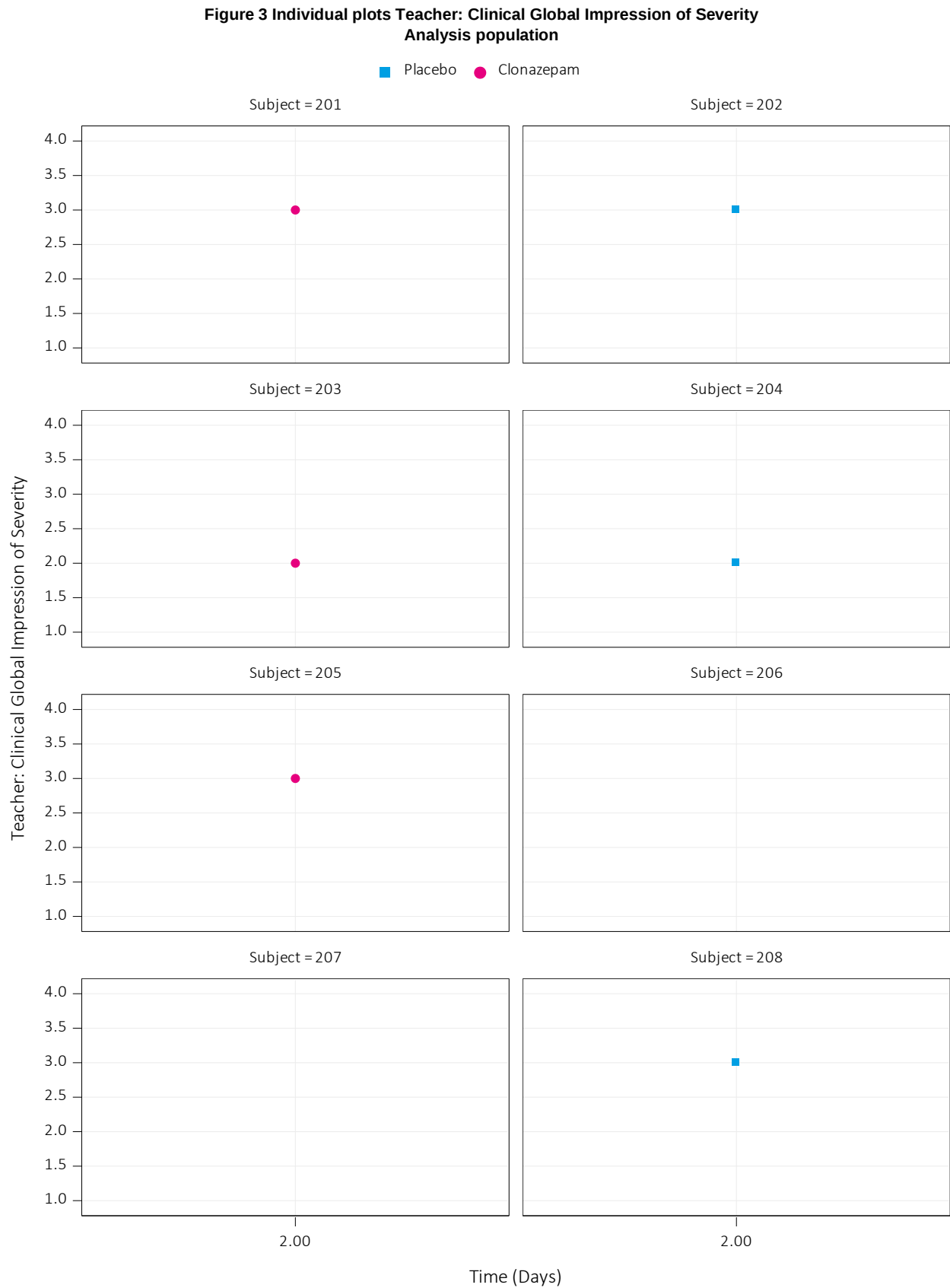
**Figure 2 Individual plots Parents: Clinical Global Impression of Improvement
Analysis population**



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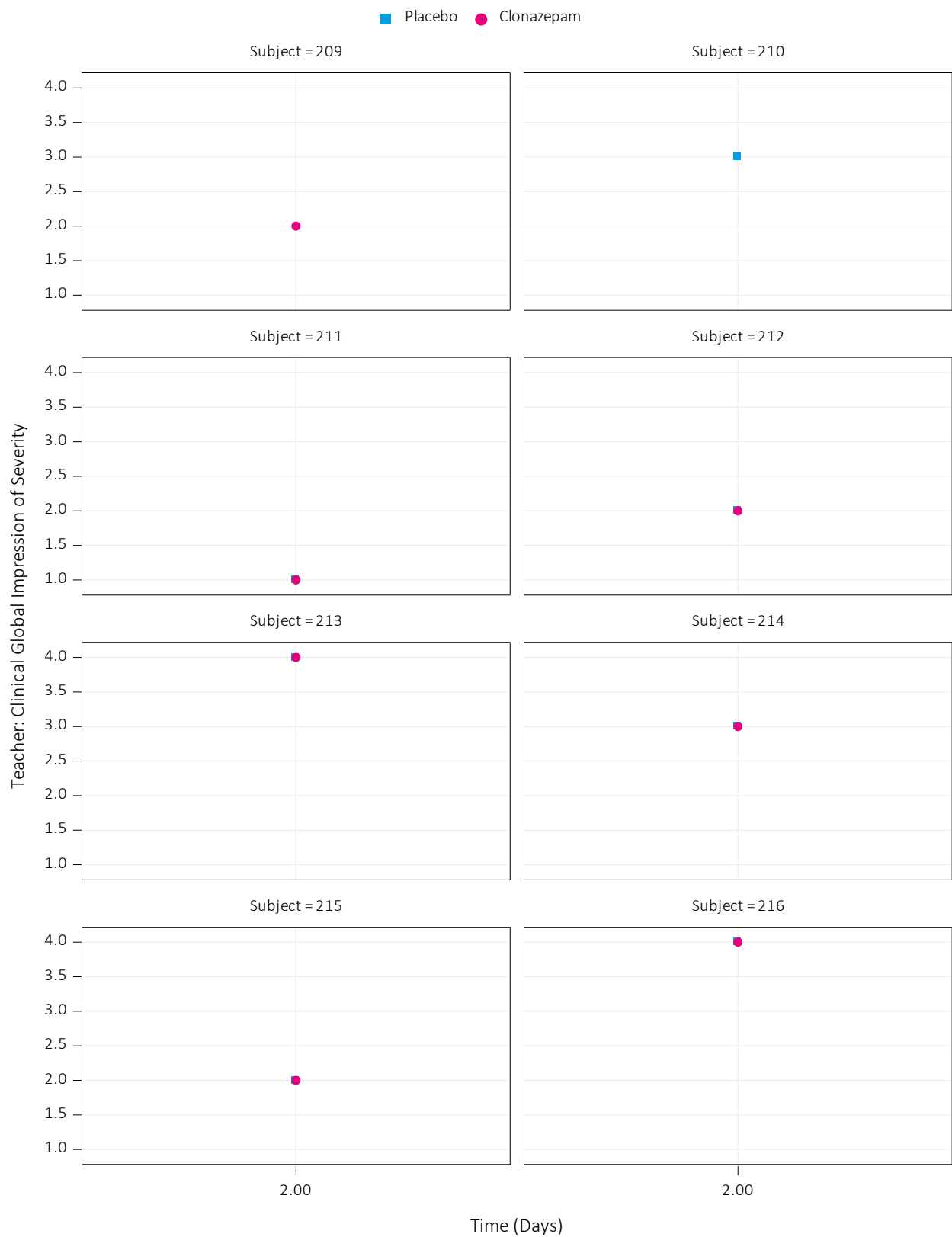
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Figure 3 Individual plots Teacher: Clinical Global Impression of Severity



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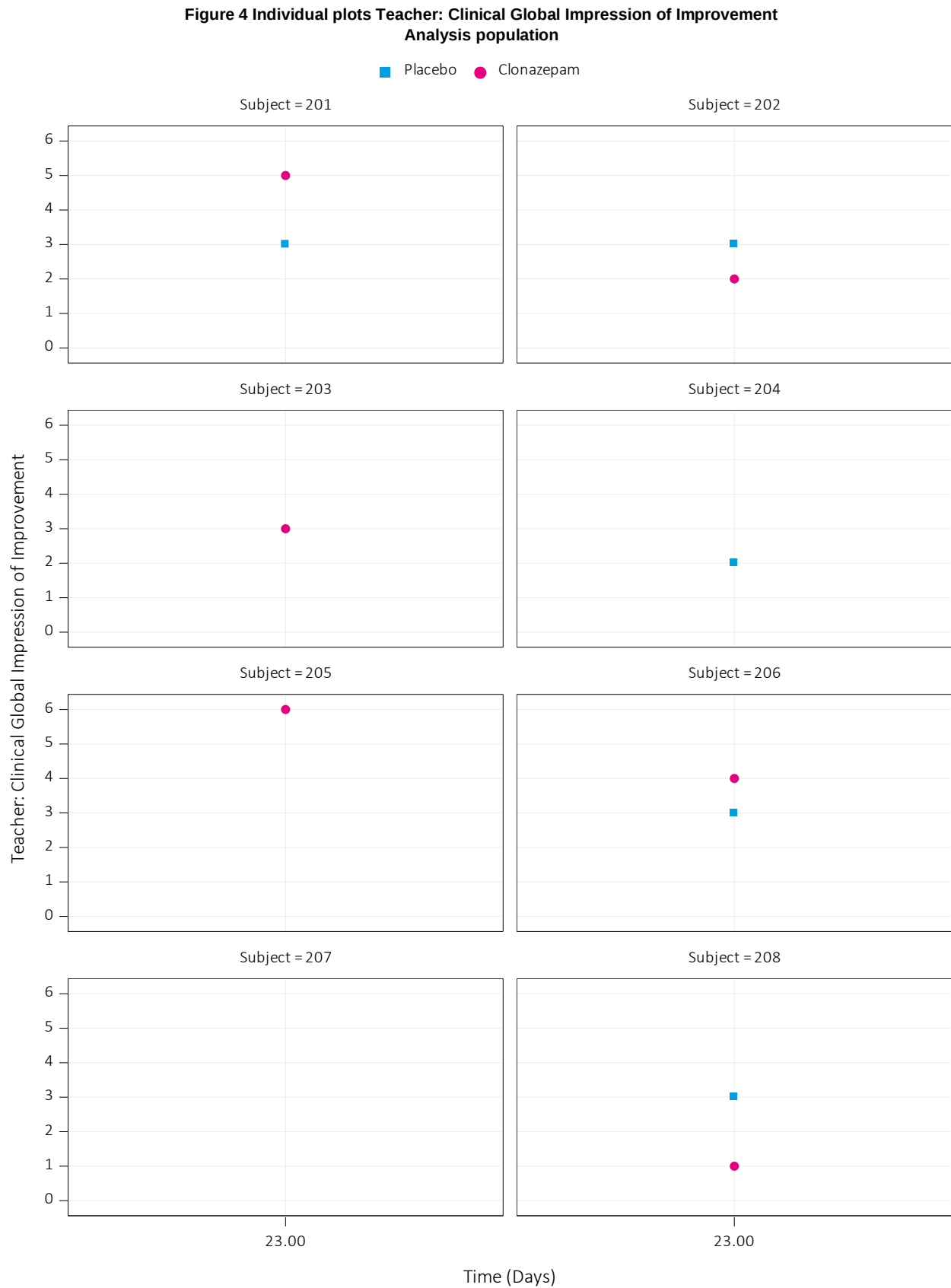
**Figure 3 Individual plots Teacher: Clinical Global Impression of Severity
Analysis population**



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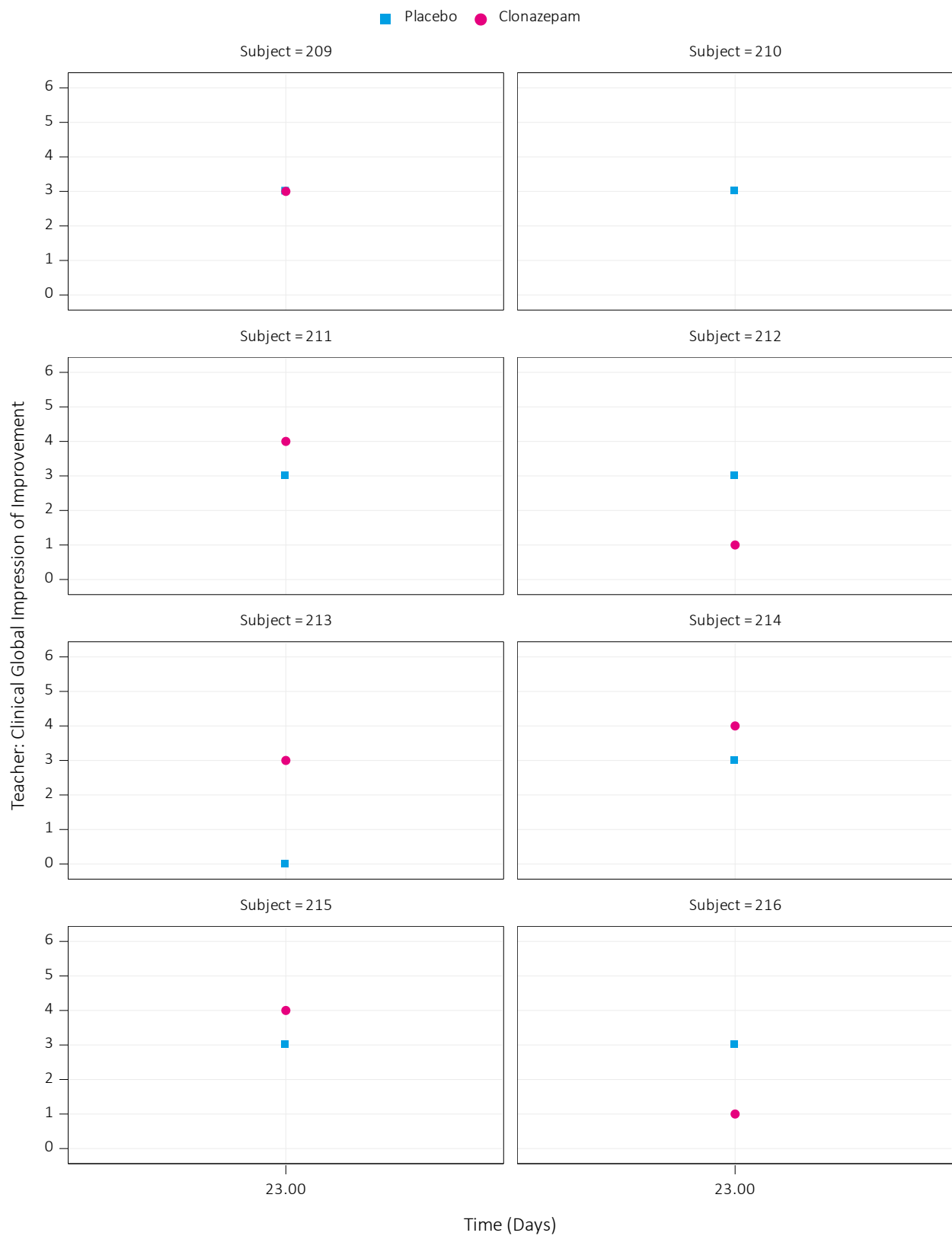
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Figure 4 Individual plots Teacher: Clinical Global Impression of Improvement



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**Figure 4 Individual plots Teacher: Clinical Global Impression of Improvement
Analysis population**



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Summary pharmacodynamic response tables and graphs

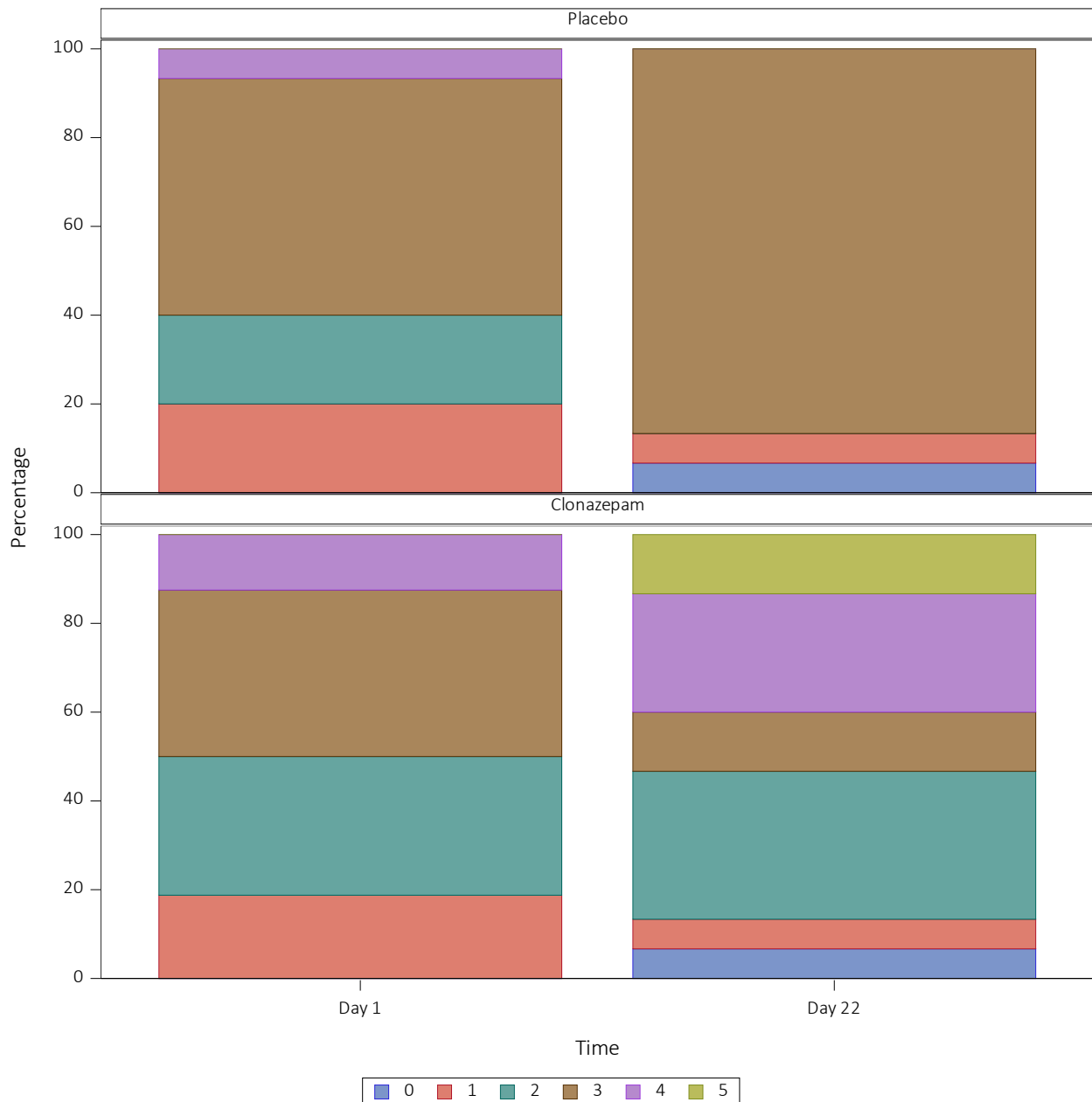
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Summaries 1 Parents: Clinical Global Impression (CGI) Score**Frequency table: Parents: Clinical Global Impression (CGI) Score**
Analysis population

Treatment	Time	0	1	2	3	4	5
Placebo	Day 1	0 (0%)	3 (20%)	3 (20%)	8 (53%)	1 (7%)	0 (0%)
	Day 22	1 (7%)	1 (7%)	0 (0%)	13 (87%)	0 (0%)	0 (0%)
Clonazepam	Day 1	0 (0%)	3 (19%)	5 (31%)	6 (38%)	2 (13%)	0 (0%)
	Day 22	1 (7%)	1 (7%)	5 (33%)	2 (13%)	4 (27%)	2 (13%)

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Figure 8.4.1.1.1 Summary Parents: Clinical Global Impression (CGI) Score
Analysis population**CGI Severity score on day 1, CGI Improvement Score on day 22**

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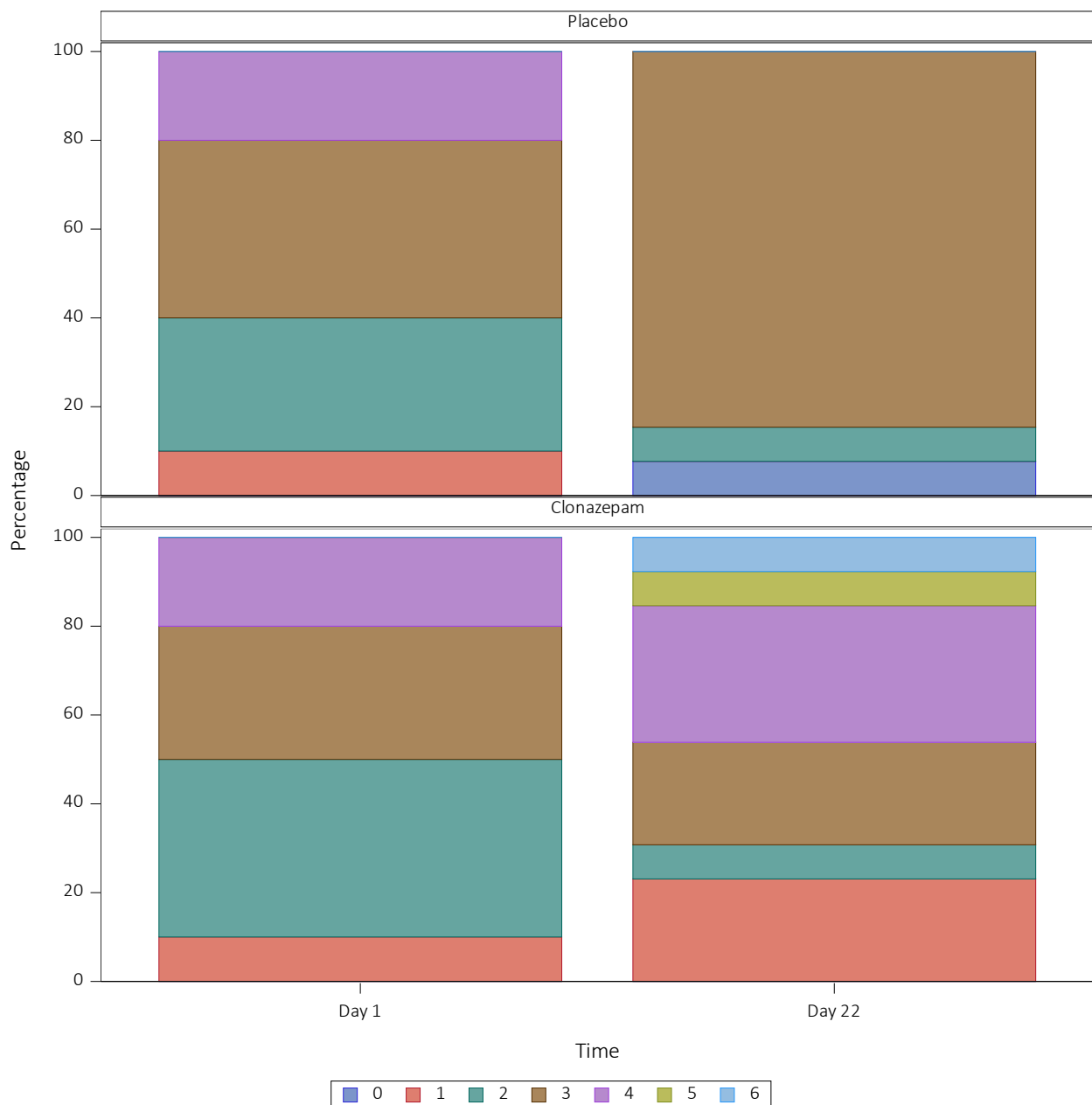
Page 1 of 1

Summaries 2 Teacher: Clinical Global Impression (CGI) Score**Frequency table: Teacher: Clinical Global Impression (CGI) Score
Analysis population**

Treatment	Time	0	1	2	3	4	5	6
Placebo	Day 1 Severity	0 (0%)	1 (10%)	3 (30%)	4 (40%)	2 (20%)	0 (0%)	0 (0%)
	Day 22 Improvement	1 (8%)	0 (0%)	1 (8%)	11 (85%)	0 (0%)	0 (0%)	0 (0%)
Clonazepam	Day 1	0 (0%)	1 (10%)	4 (40%)	3 (30%)	2 (20%)	0 (0%)	0 (0%)
	Day 22	0 (0%)	3 (23%)	1 (8%)	3 (23%)	4 (31%)	1 (8%)	1 (8%)

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**Figure 8.4.1.1.2 Summary Teacher: Clinical Global Impression (CGI) Score
Analysis population****CGI Severity score on day 1, CGI Improvement Score on day 22**

Source: g:\2019\CHDR1939-B\PMX\Reports\FPD\2022_09_26\Programs\CHDR1939B_FPD_General_RTF_v1.sas
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Analysis results pharmacodynamic response table, LSM table and graphs

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Analysis results 1 Parents: Clinical Global Impression (CGI) Improvement**Analysis results 1 Parents: Clinical Global Impression (CGI) Improvement
Analysis population**

Effect/Contrast	P-value	First LSM of contrast	Second LSM of contrast	Estimate of the difference	95% CI	
					Lower	Upper
CGI Severity score	0.4738					
Treatment	0.6635					
Period	0.6635					
Clonazepam - Placebo	0.6635	2.9	2.7	0.2	-0.8	1.2

Treatment	LSM	95% CI		LSM change from baseline	95% CI	
		Lower	Upper		Lower	Upper
Placebo	2.7	2.0	3.3	0.2	-0.4	0.9
Clonazepam	2.9	2.2	3.5	0.4	-0.2	1.1

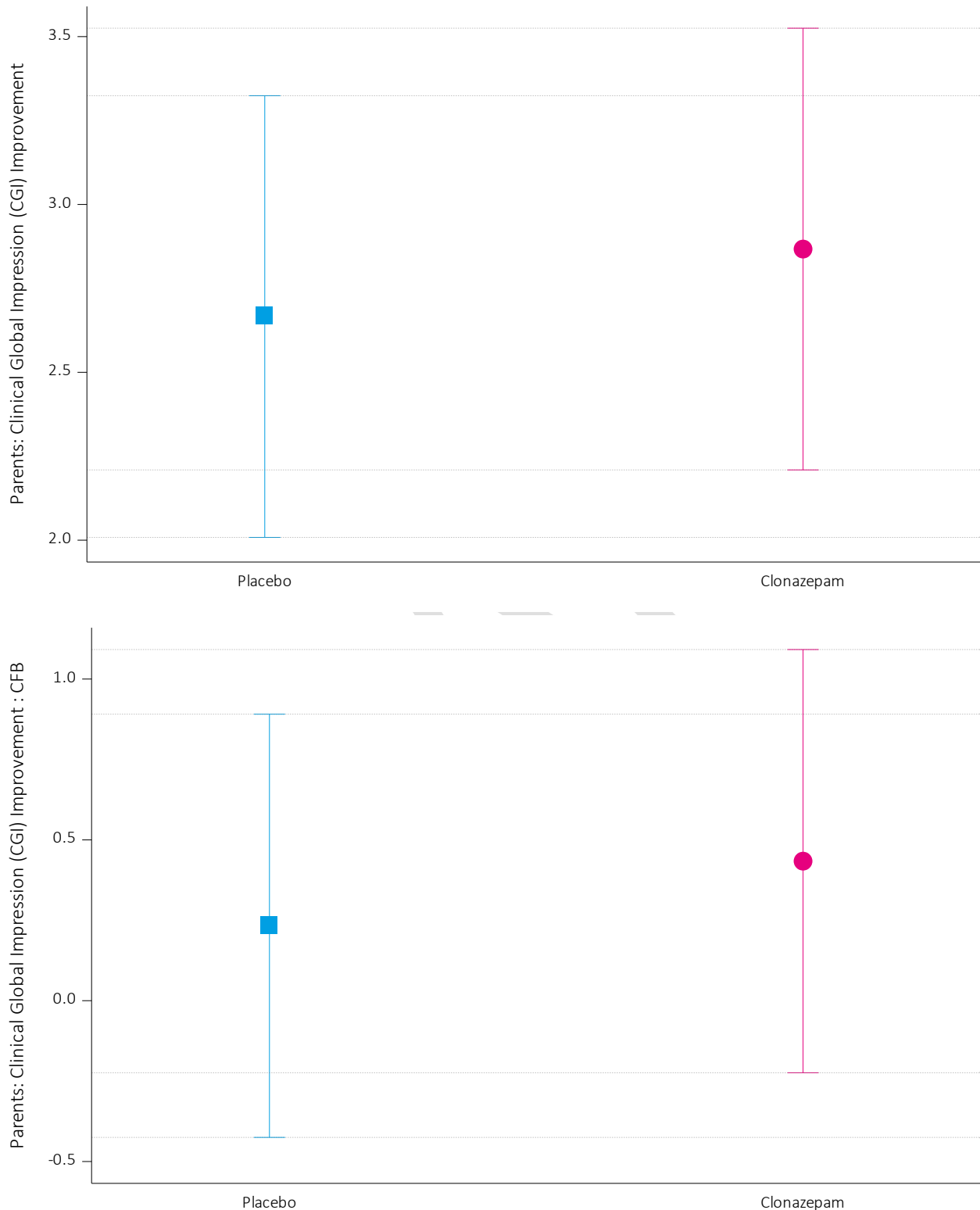
CGI Improvement score as response, CGI Severity score as covariate, period as occasion

Source: g:\2019\CHDR1939-B\PMX\Reports\FPD\2022_09_26\Programs\CHDR1939B_FPD_General_RTF_v1.sas
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**Estimated means 1 Parents: Clinical Global Impression (CGI) Improvement
Analysis population**

Estimated means (95% CI error bars)



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Analysis results 2 Teacher: Clinical Global Impression (CGI) Improvement**Analysis results 2 Teacher: Clinical Global Impression (CGI) Improvement**
Analysis population

Effect/Contrast	P-value	First LSM of contrast	Second LSM of contrast	Estimate of the difference	95% CI	
					Lower	Upper
CGI Severity score	0.3484					
Treatment	0.2617					
Period	0.8018					
Clonazepam - Placebo	0.2617	3.3	2.5	0.8	-0.7	2.3

Treatment	LSM	95% CI		LSM change from baseline	95% CI	
		Lower	Upper		Lower	Upper
Placebo	2.5	1.6	3.5	-0.1	-1.1	0.9
Clonazepam	3.3	2.4	4.3	0.7	-0.3	1.7

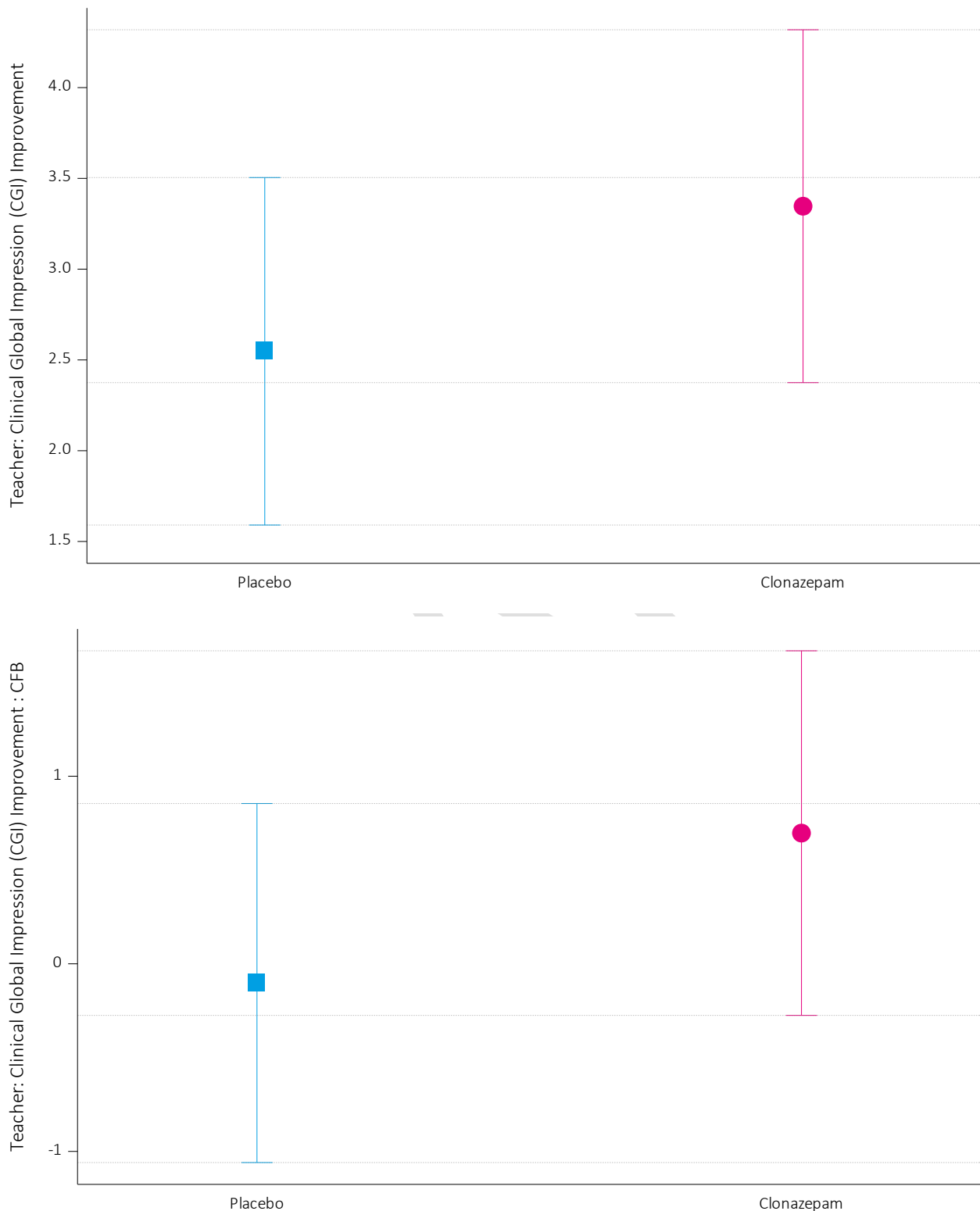
CGI Improvement score as response, CGI Severity score as covariate, period as occasion

Source: g:\2019\CHDR1939-B\PMX\Reports\FPD\2022_09_26\Programs\CHDR1939B_FPD_General_RTF_v1.sas
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**Estimated means 2 Teacher: Clinical Global Impression (CGI) Improvement
Analysis population**

Estimated means (95% CI error bars)



Source: g:\2019\CHDR1939-B\PMX\Reports\FPD\2022_09_26\Programs\CHDR1939B_FPD_General_RTF_v1.sas
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Unedited SAS output

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SAS output 1 Parents: Clinical Global Impression (CGI) Improvement

Proc MIXED analysis of Parents: Clinical Global Impression (CGI) Improvement
Analysis population

1

Model Information

Data Set	WORK.MIX
Dependent Variable	CGITest
Covariance Structure	Variance Components
Estimation Method	REML
Residual Variance Method	Profile
Fixed Effects SE Method	Kenward-Roger
Degrees of Freedom Method	Kenward-Roger

Class Level Information

Class	Levels	Values
by1	2	Placebo Clonazepam
Occasion	2	1 2
SubjectNr	16	201 202 203 204 205 206 207 208 209 210 211 212 213 214 215 216

Dimensions

Covariance Parameters	2
Columns in X	6
Columns in Z	16
Subjects	1
Max Obs per Subject	30

Number of Observations

Number of Observations Read	31
Number of Observations Used	30
Number of Observations Not Used	1

Iteration History

Iteration	Evaluations	-2 Res Log Like	Criterion
0	1	95.53220864	
1	2	95.53110331	0.00000000

Convergence criteria met.

Covariance Parameter Estimates

Cov Parm	Estimate
SubjectNr	0.01430
Residual	1.5176

Fit Statistics

-2 Res Log Likelihood	95.5
AIC (Smaller is Better)	99.5
AICC (Smaller is Better)	100.1
BIC (Smaller is Better)	101.1

Proc MIXED analysis of Parents: Clinical Global Impression (CGI) Improvement
Analysis population

2

Null Model Likelihood Ratio Test

DF	Chi-Square	Pr > ChiSq
1	0.00	0.9735

Type 3 Tests of Fixed Effects

Effect	Num DF	Den DF	F Value	Pr > F
by1	1	12.8	0.20	0.6635
Occasion	1	12.8	0.20	0.6635
preCGITest	1	15.7	0.54	0.4738

Estimates

Label	Estimate	Standard Error	DF	t Value	Pr > t	Alpha	Lower	Upper
Clonazepam - Placebo	0.2010	0.4512	12.8	0.45	0.6635	0.05	-0.7755	1.1775

Least Squares Means

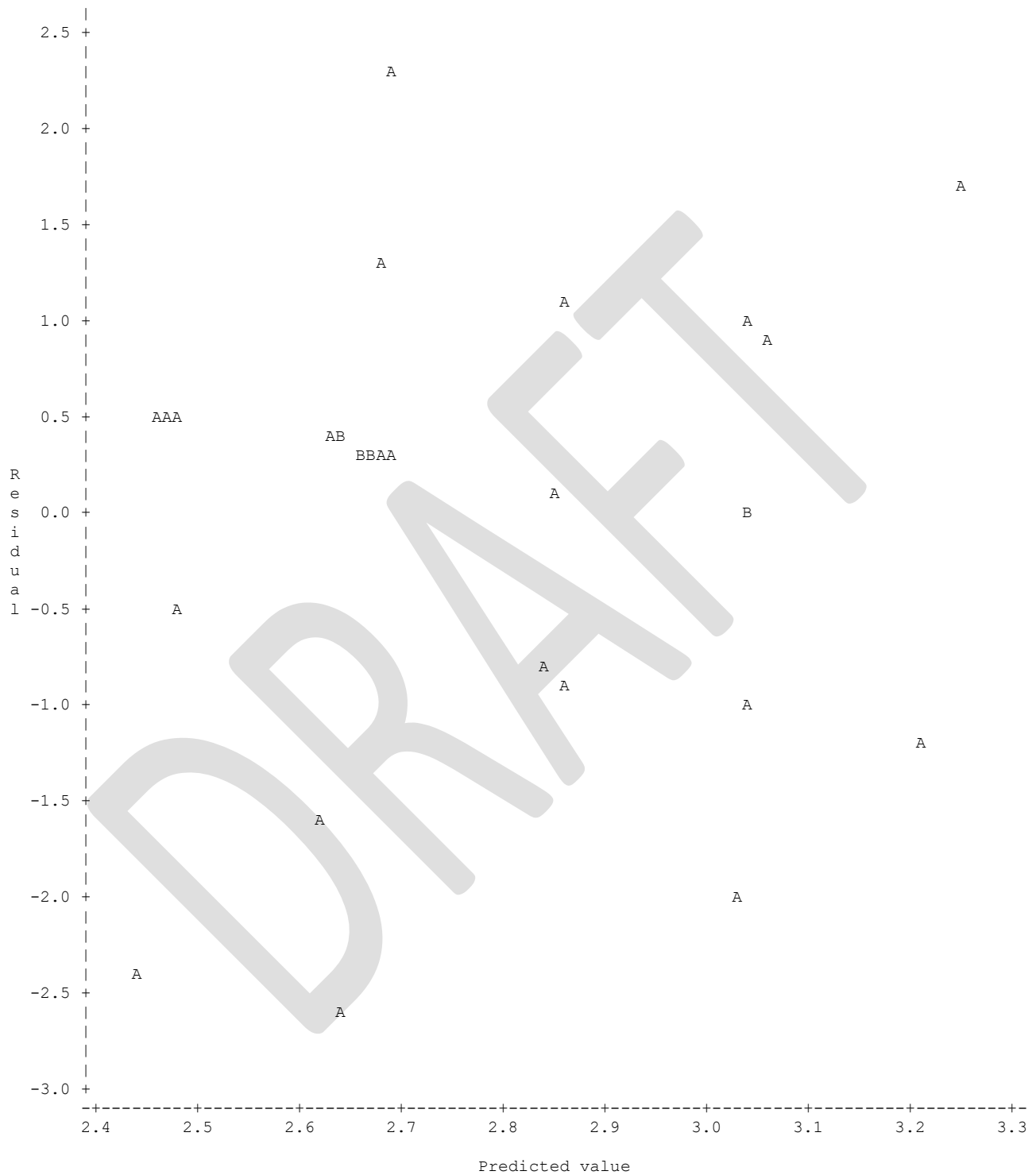
Effect	Treatment	Estimate	Standard Error	DF	t Value	Pr > t	Alpha	Lower	Upper
by1	Placebo	2.6662	0.3200	26	8.33	<.0001	0.05	2.0083	3.3240
by1	Clonazepam	2.8672	0.3200	26	8.96	<.0001	0.05	2.2093	3.5250

Proc MIXED analysis of Parents: Clinical Global Impression (CGI) Improvement
Analysis population

3

Residuals vs Predicted values

Plot of Resid*Pred. Legend: A = 1 obs, B = 2 obs, etc.



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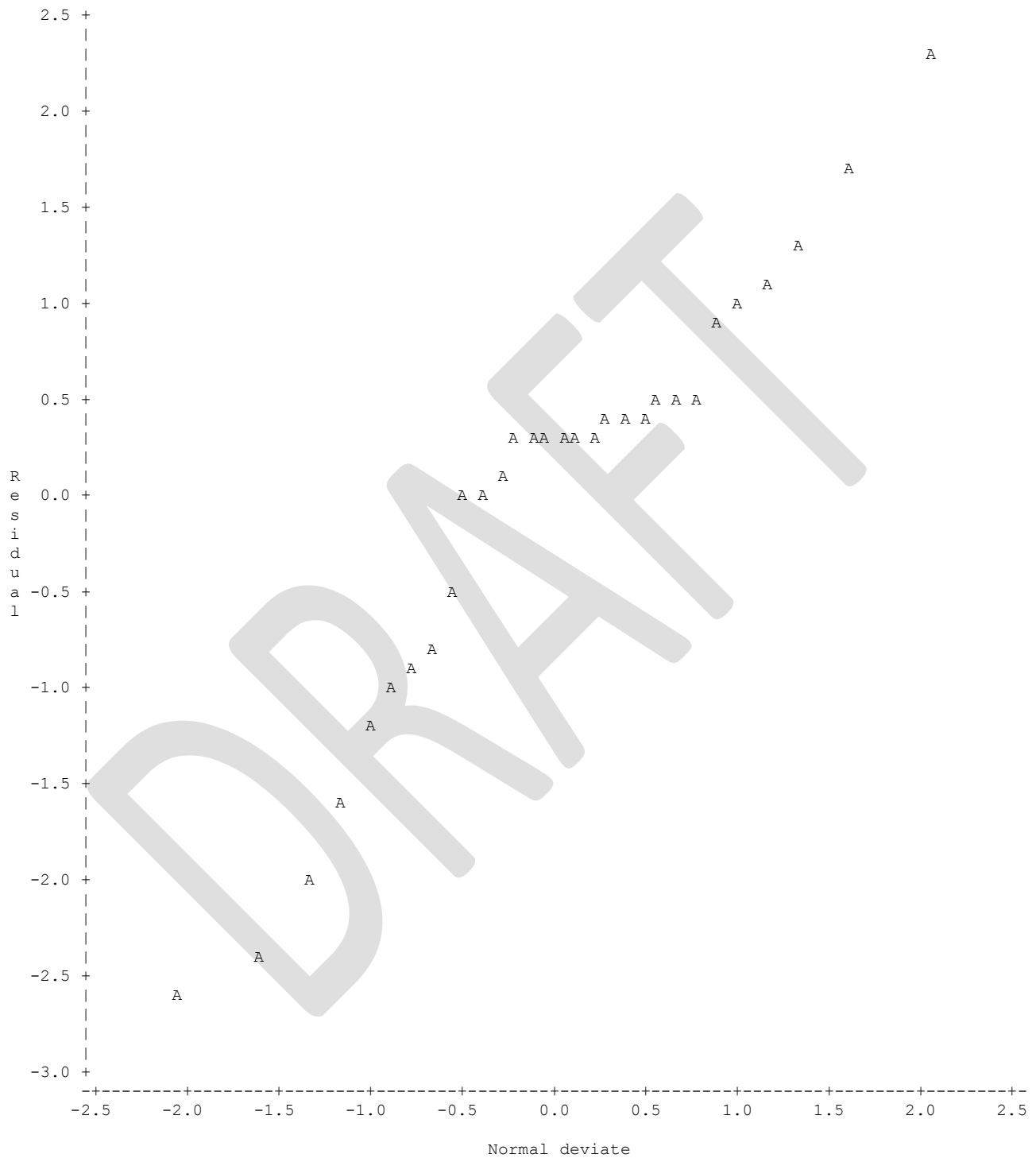
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Proc MIXED analysis of Parents: Clinical Global Impression (CGI) Improvement
Analysis population

4

Normal probability plot for residuals CGITest
(Shapiro-Wilk Test Statistic = 0.935 : P-value=0.0659)

Plot of Resid*rankres. Legend: A = 1 obs, B = 2 obs, etc.



NOTE: 1 obs had missing values.

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SAS output 2 Teacher: Clinical Global Impression (CGI) Improvement

Proc MIXED analysis of Teacher: Clinical Global Impression (CGI) Improvement
Analysis population

1

Model Information

Data Set	WORK.MIX
Dependent Variable	CGITest_T
Covariance Structure	Variance Components
Estimation Method	REML
Residual Variance Method	Profile
Fixed Effects SE Method	Kenward-Roger
Degrees of Freedom Method	Kenward-Roger

Class Level Information

Class	Levels	Values
by1	2	Placebo Clonazepam
Occasion	2	1 2
SubjectNr	14	201 202 203 204 205 208 209 210 211 212 213 214 215 216

Dimensions

Covariance Parameters	2
Columns in X	6
Columns in Z	14
Subjects	1
Max Obs per Subject	20

Number of Observations

Number of Observations Read	31
Number of Observations Used	20
Number of Observations Not Used	11

Iteration History

Iteration	Evaluations	-2 Res Log Like	Criterion
0	1	64.16577482	
1	2	63.99293861	0.00000021
2	1	63.99293492	0.00000000

Convergence criteria met.

Estimated G matrix is not positive definite.

Covariance Parameter Estimates

Cov Parm	Estimate
SubjectNr	-0.3870
Residual	2.2232

Fit Statistics

-2 Res Log Likelihood	64.0
AIC (Smaller is Better)	68.0
AICC (Smaller is Better)	68.9
BIC (Smaller is Better)	69.3

Proc MIXED analysis of Teacher: Clinical Global Impression (CGI) Improvement
Analysis population

2

Null Model Likelihood Ratio Test

DF	Chi-Square	Pr > ChiSq
1	0.17	0.6776

Type 3 Tests of Fixed Effects

Effect	Num DF	Den DF	F Value	Pr > F
by1	1	9.84	1.42	0.2617
Occasion	1	9.86	0.07	0.8018
preCGITest_T	1	5.05	1.07	0.3484

Estimates

Label	Estimate	Standard Error	DF	t Value	Pr > t	Alpha	Lower	Upper
Clonazepam - Placebo	0.7993	0.6712	9.84	1.19	0.2617	0.05	-0.6997	2.2983

Least Squares Means

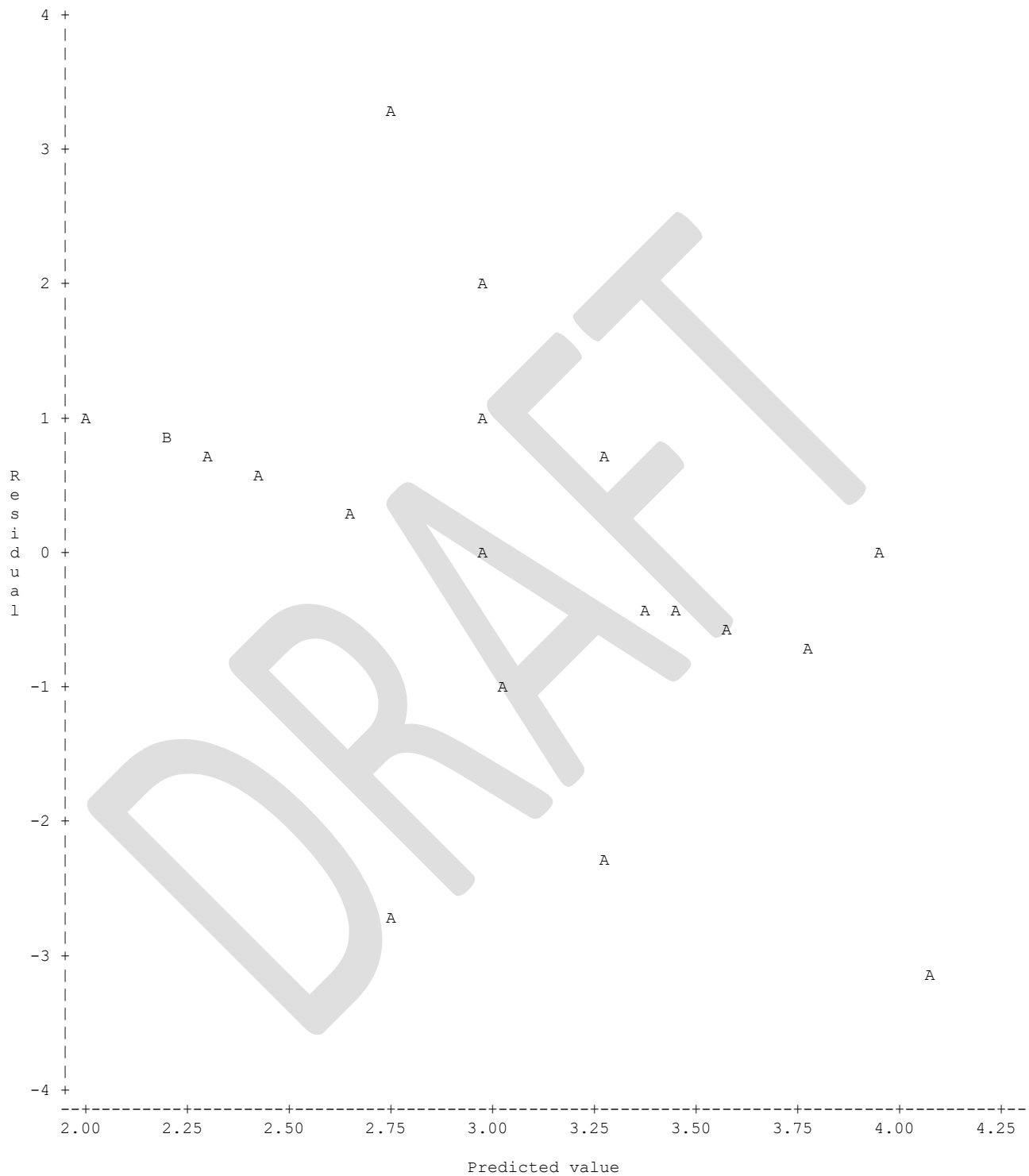
Effect	Treatment	Estimate	Standard Error	DF	t Value	Pr > t	Alpha	Lower	Upper
by1	Placebo	2.5476	0.4514	16	5.64	<.0001	0.05	1.5906	3.5045
by1	Clonazepam	3.3468	0.4584	16	7.30	<.0001	0.05	2.3750	4.3187

Proc MIXED analysis of Teacher: Clinical Global Impression (CGI) Improvement
Analysis population

3

Residuals vs Predicted values

Plot of Resid*Pred. Legend: A = 1 obs, B = 2 obs, etc.



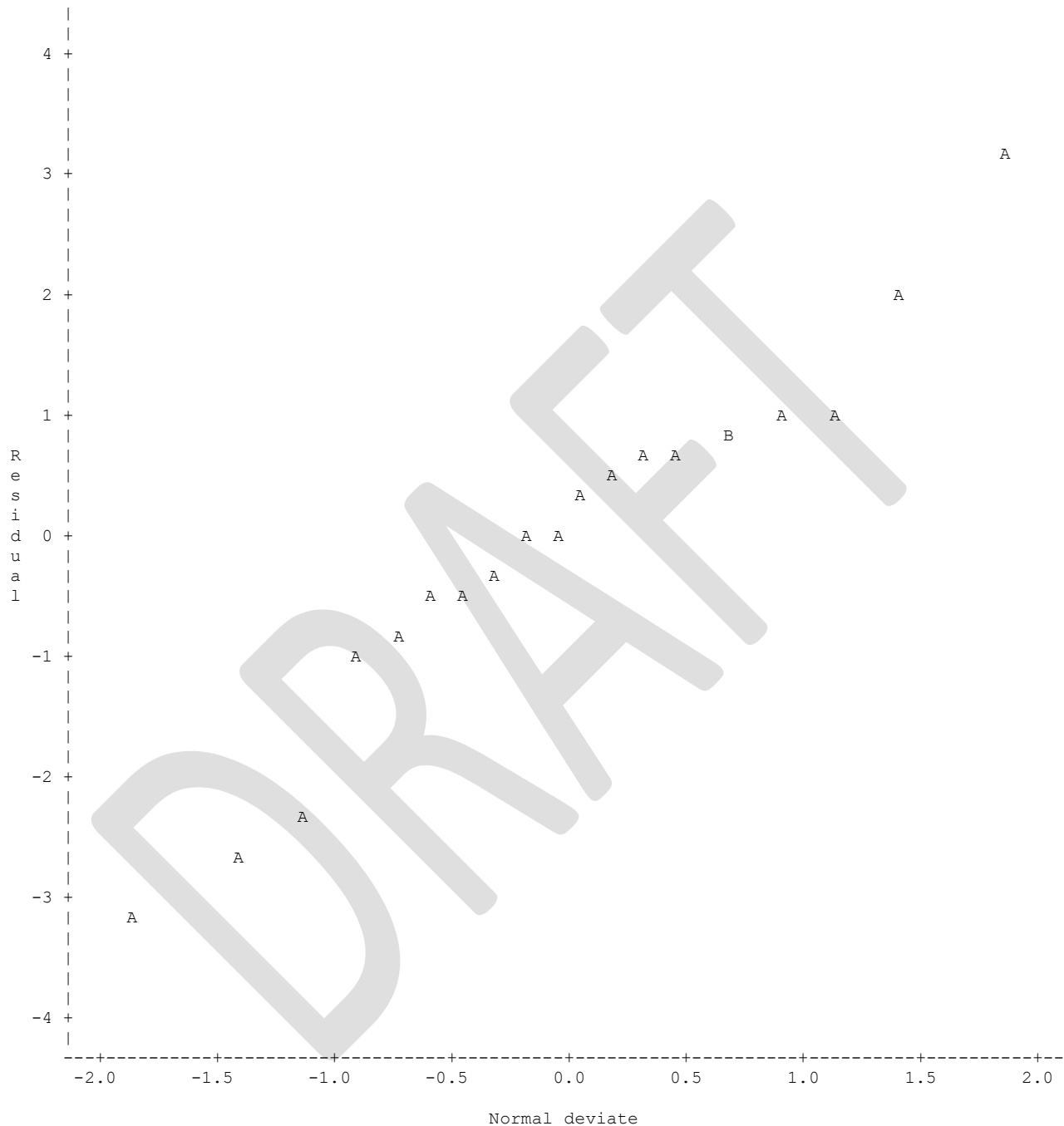
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Proc MIXED analysis of Teacher: Clinical Global Impression (CGI) Improvement
Analysis population

4

Normal probability plot for residuals CGITest_T
(Shapiro-Wilk Test Statistic = 0.952 : P-value=0.3920)

Plot of Resid*rankres. Legend: A = 1 obs, B = 2 obs, etc.



NOTE: 11 obs had missing values.