

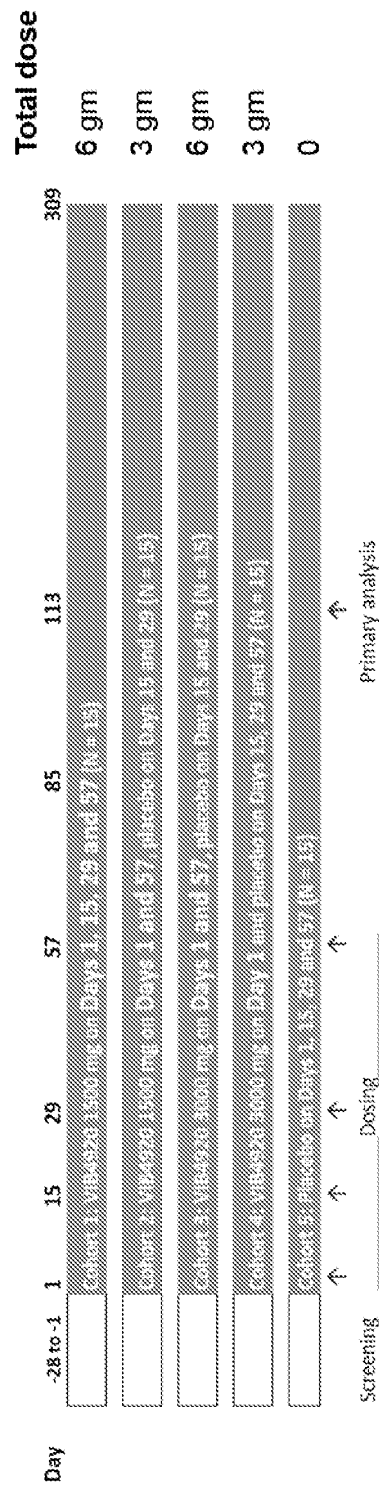


FIG. 1

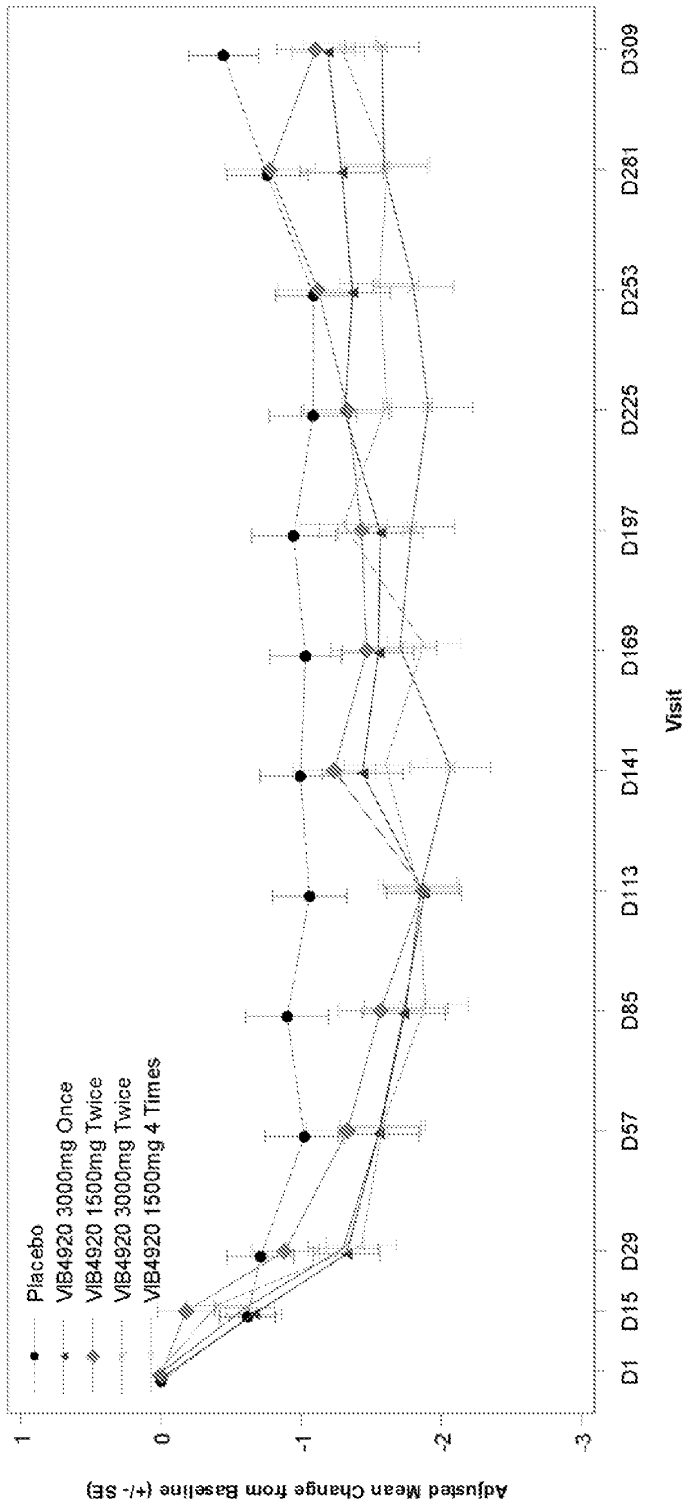


FIG. 2

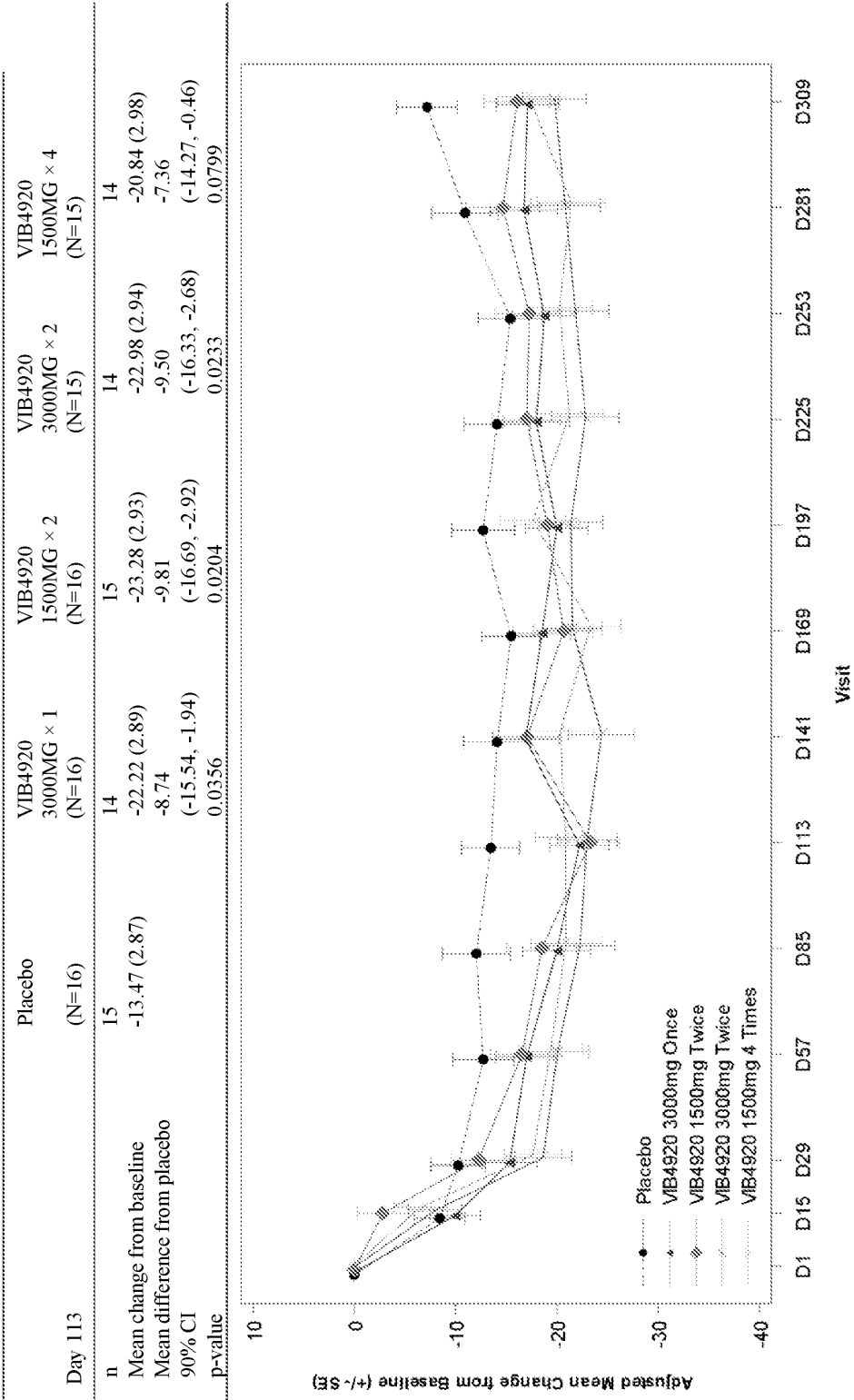


FIG. 3

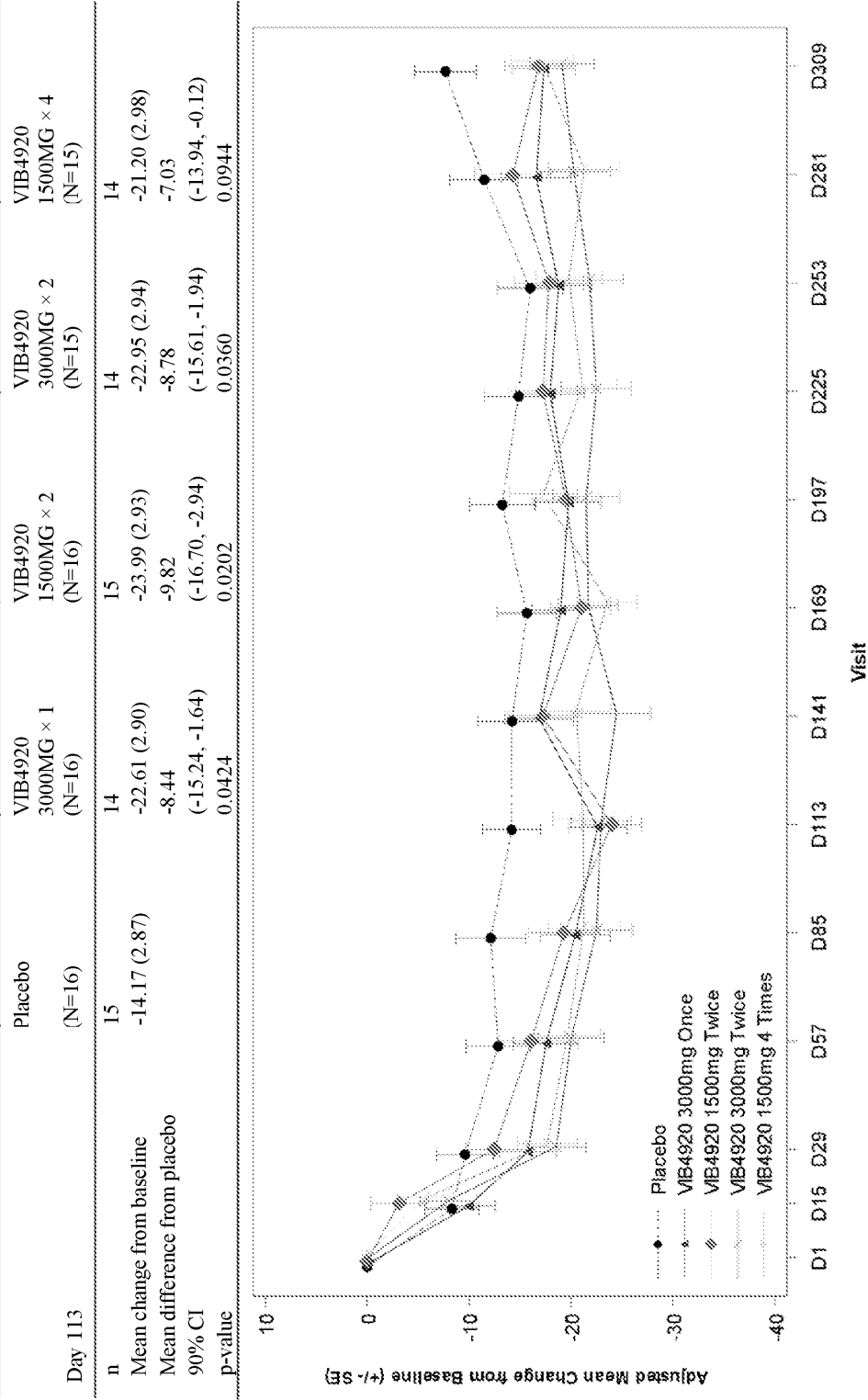


FIG. 4

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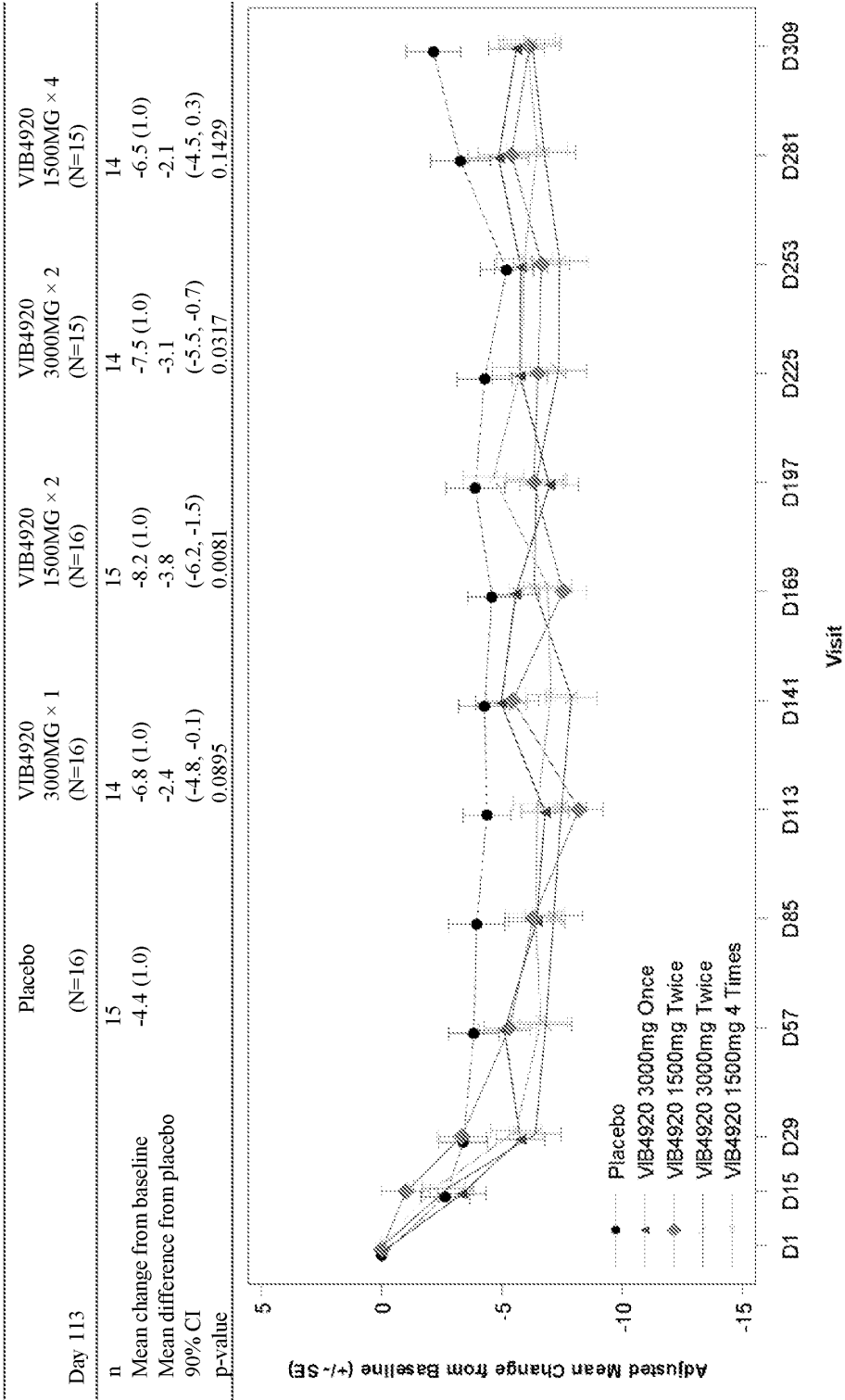


FIG. 5

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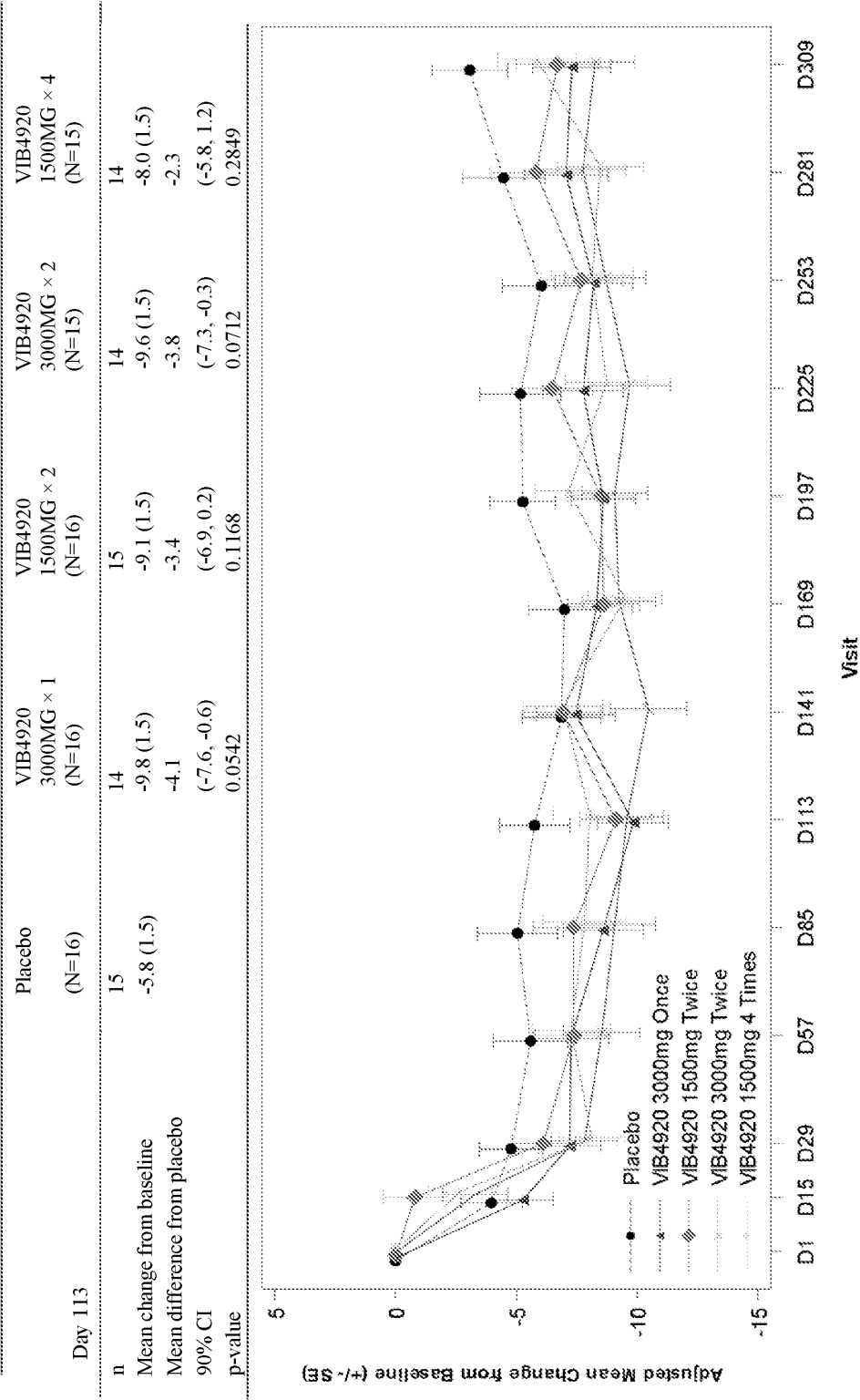


FIG. 6

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Day 113		Placebo (N=16)	VIB4920 3000MG × 1 (N=16)	VIB4920 1500MG × 2 (N=16)	VIB4920 3000MG × 2 (N=15)	VIB4920 1500MG × 4 (N=15)
n		15	14	15	14	14
Mean change from baseline		-26.1 (5.3)	-38.0 (5.4)	-40.4 (5.4)	-38.2 (5.6)	-42.1 (5.6)
Mean difference from placebo			-12.0	-14.3	-12.1	-16.0
90% CI			(-24.7, 0.8)	(-27.0, -1.6)	(-25.0, 0.8)	(-28.9, -3.1)
p-value			0.1215	0.0642	0.1213	0.0425

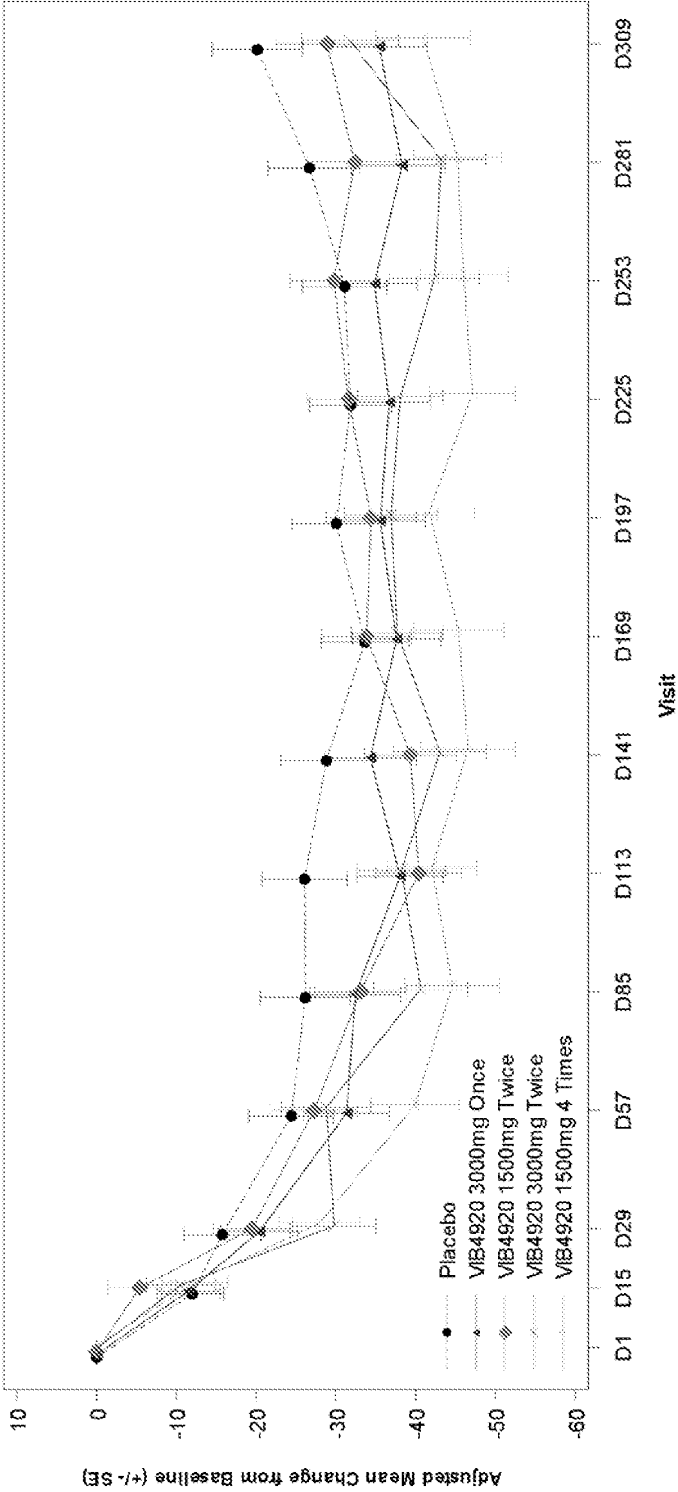


FIG. 7

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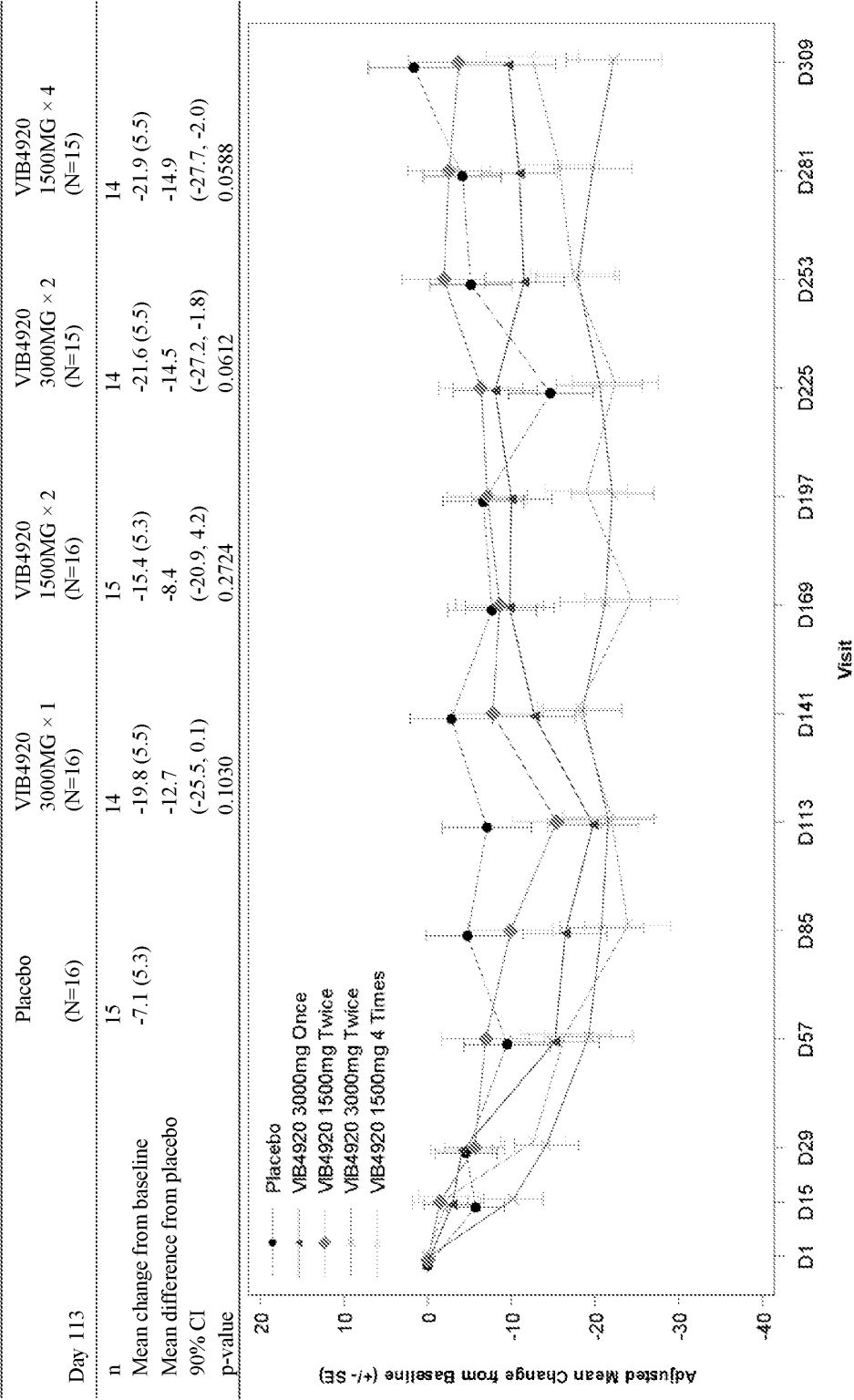


FIG. 8

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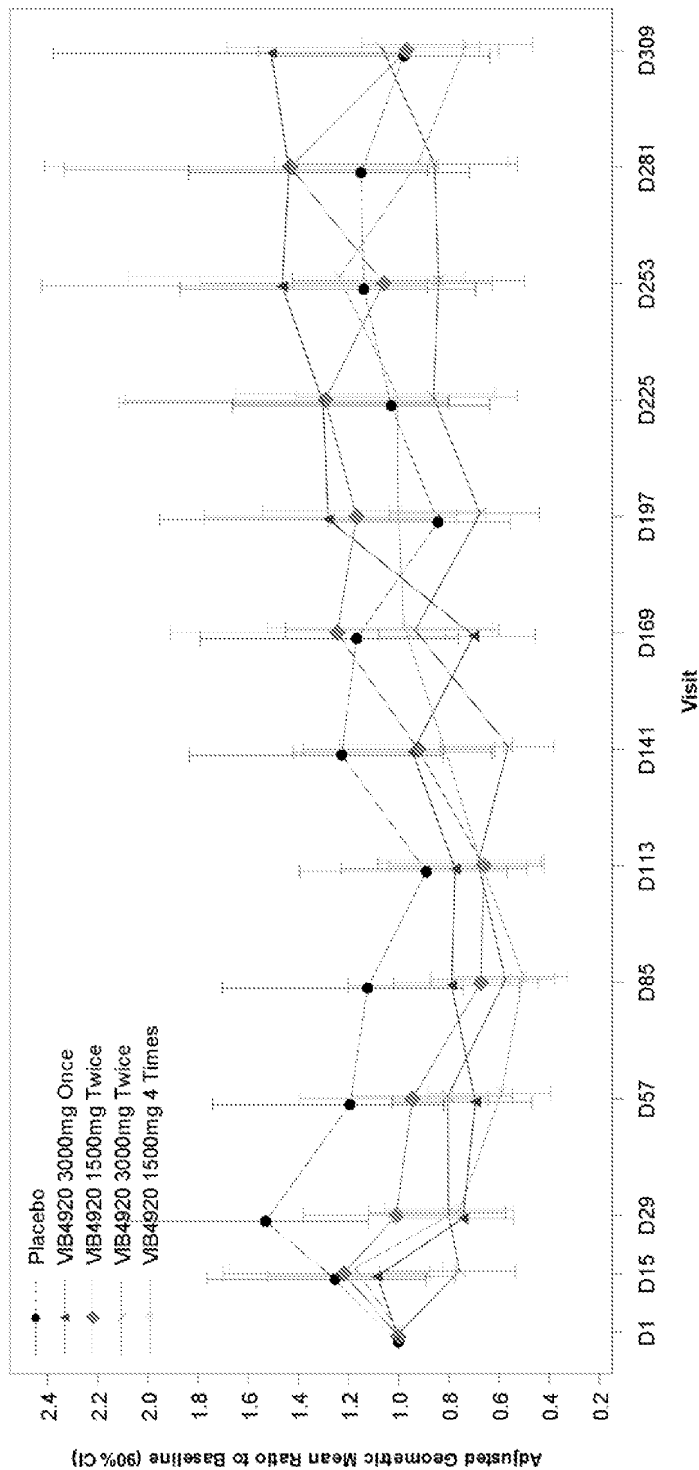


FIG. 9

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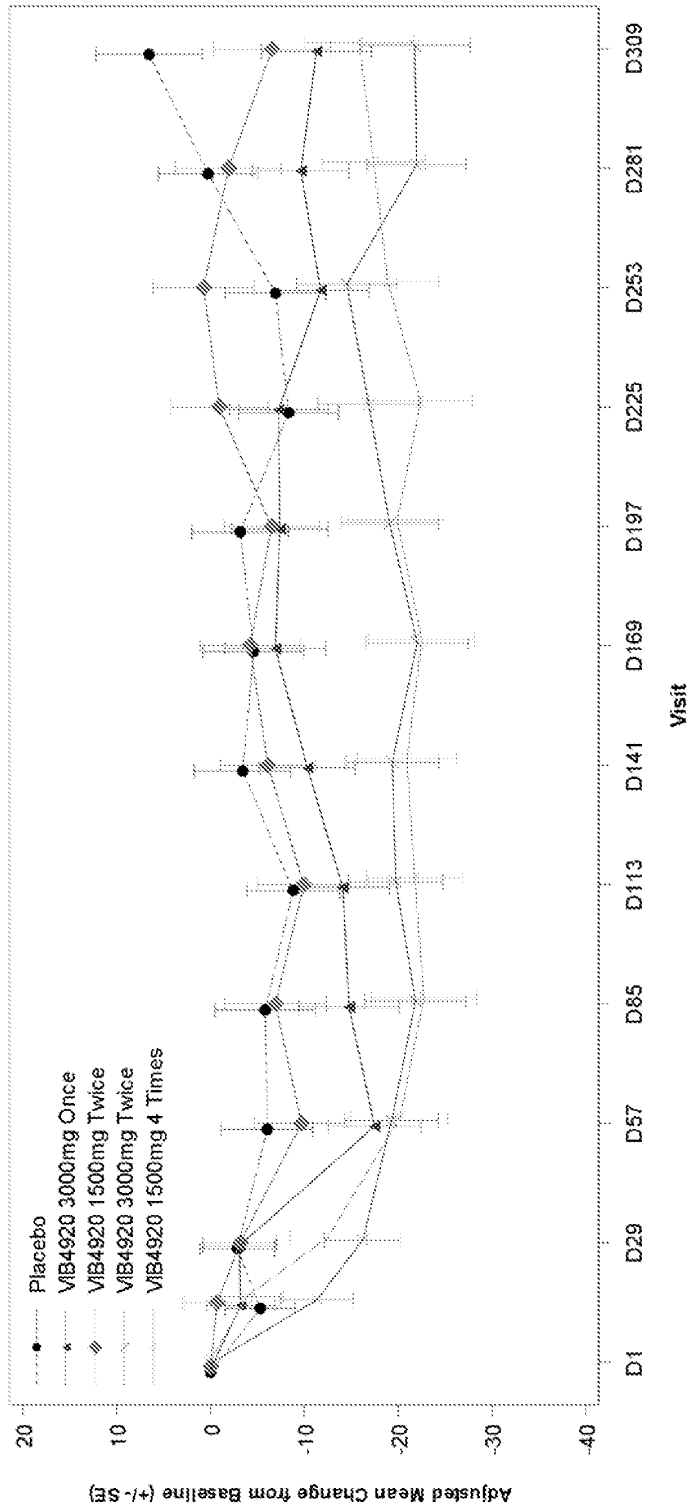


FIG. 10

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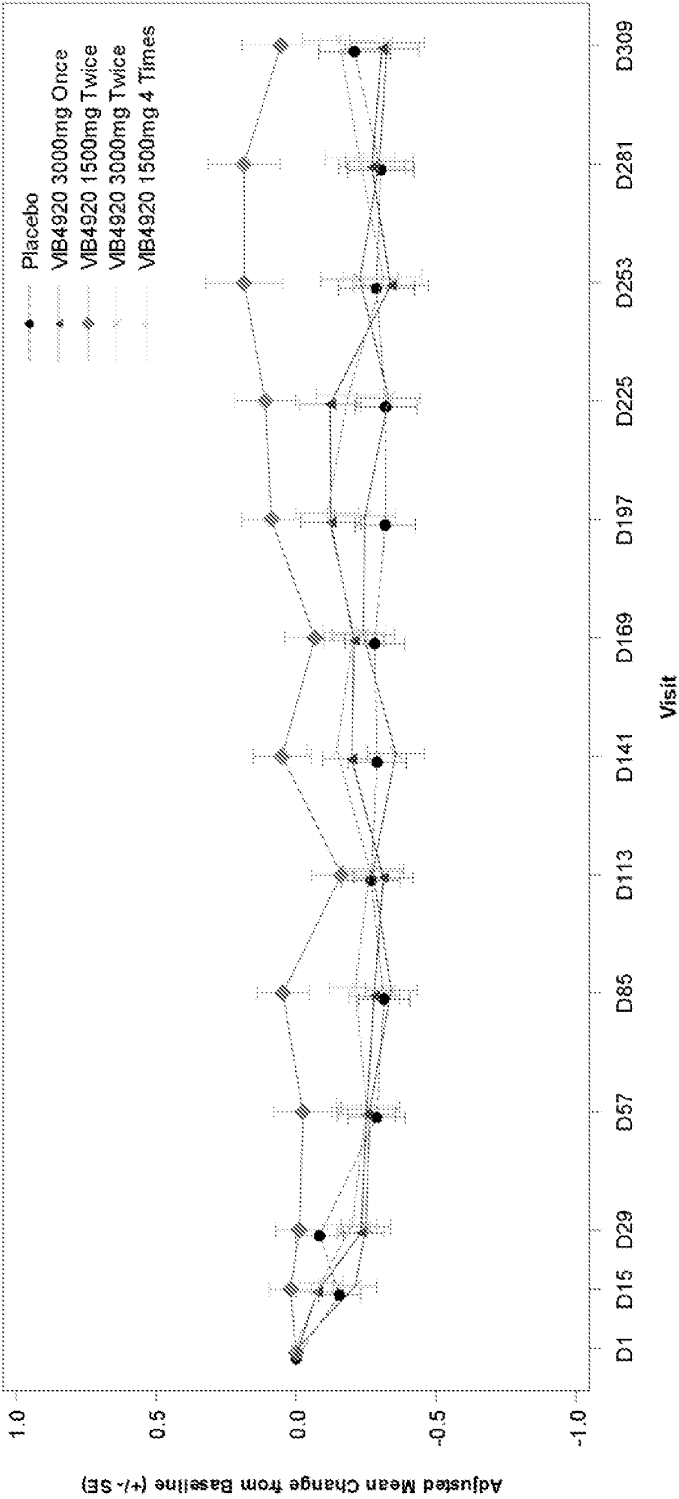


FIG. 11

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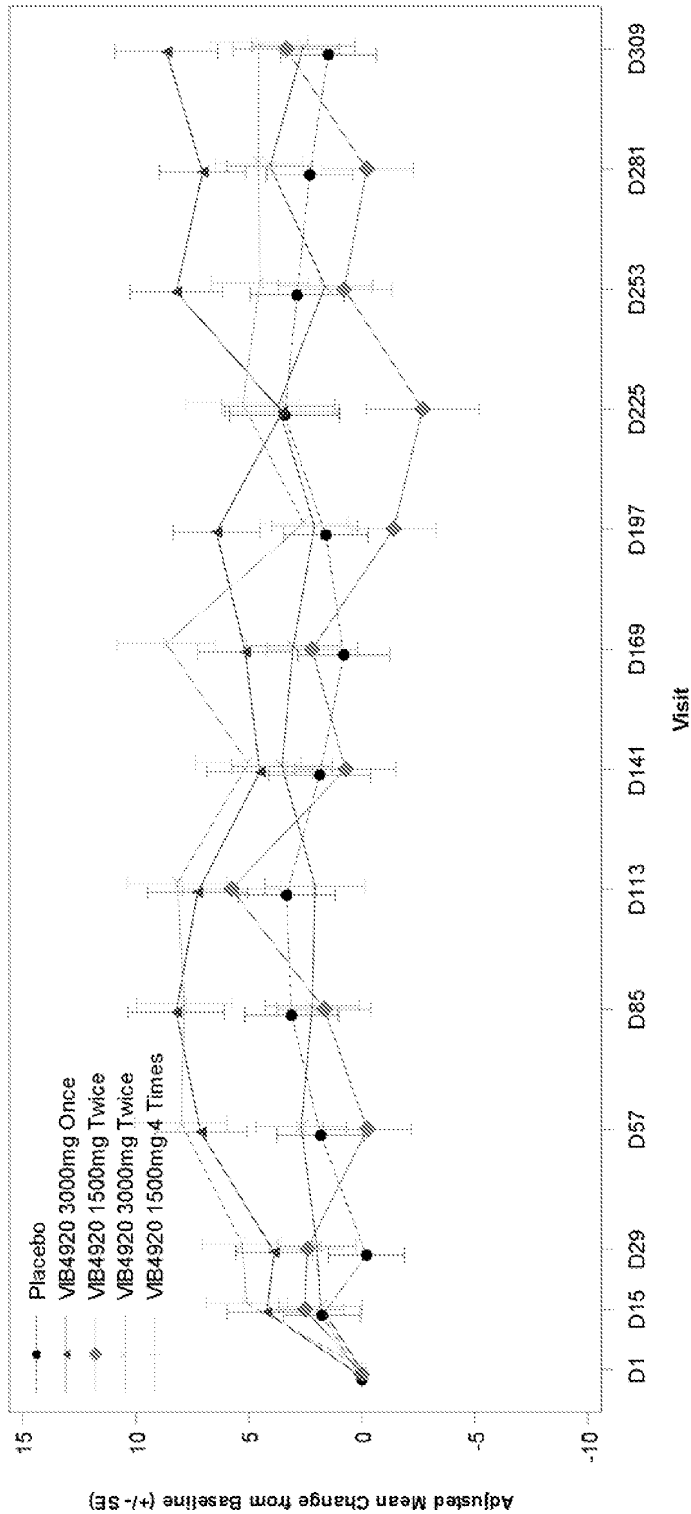


FIG. 12

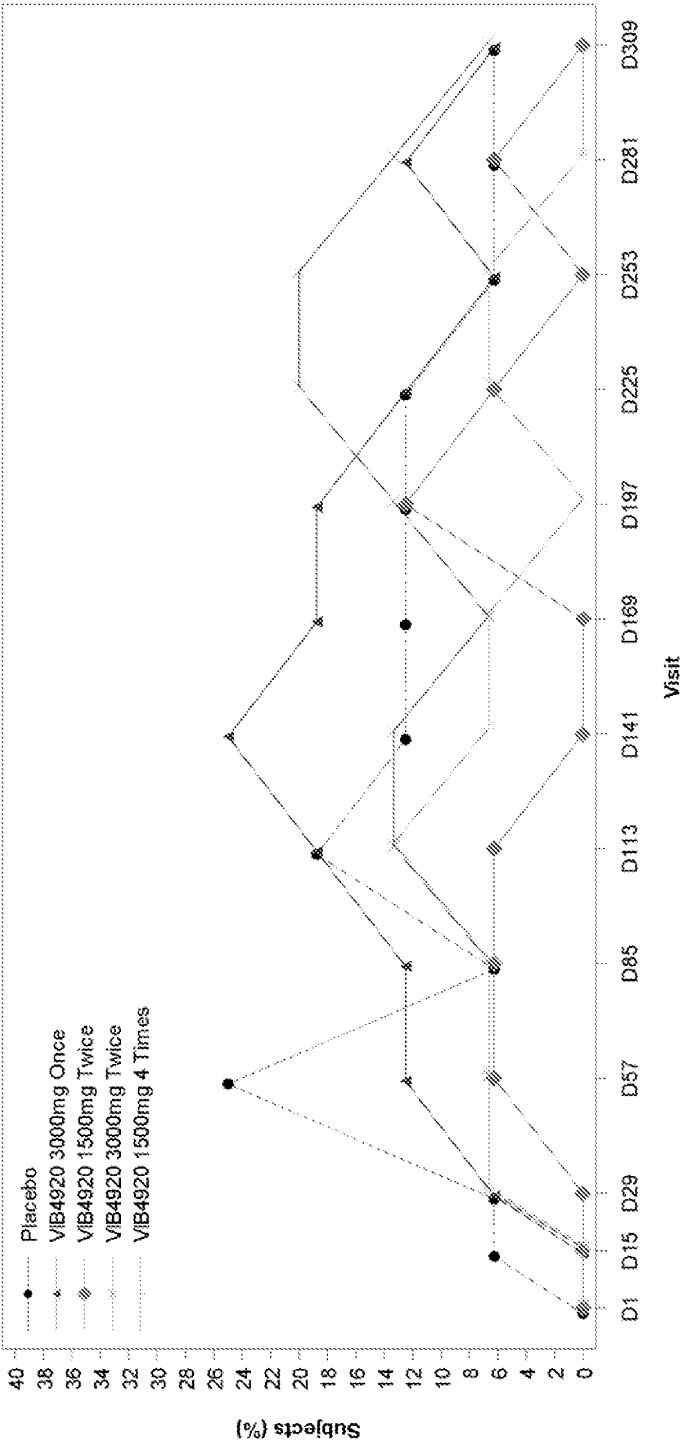


FIG. 13

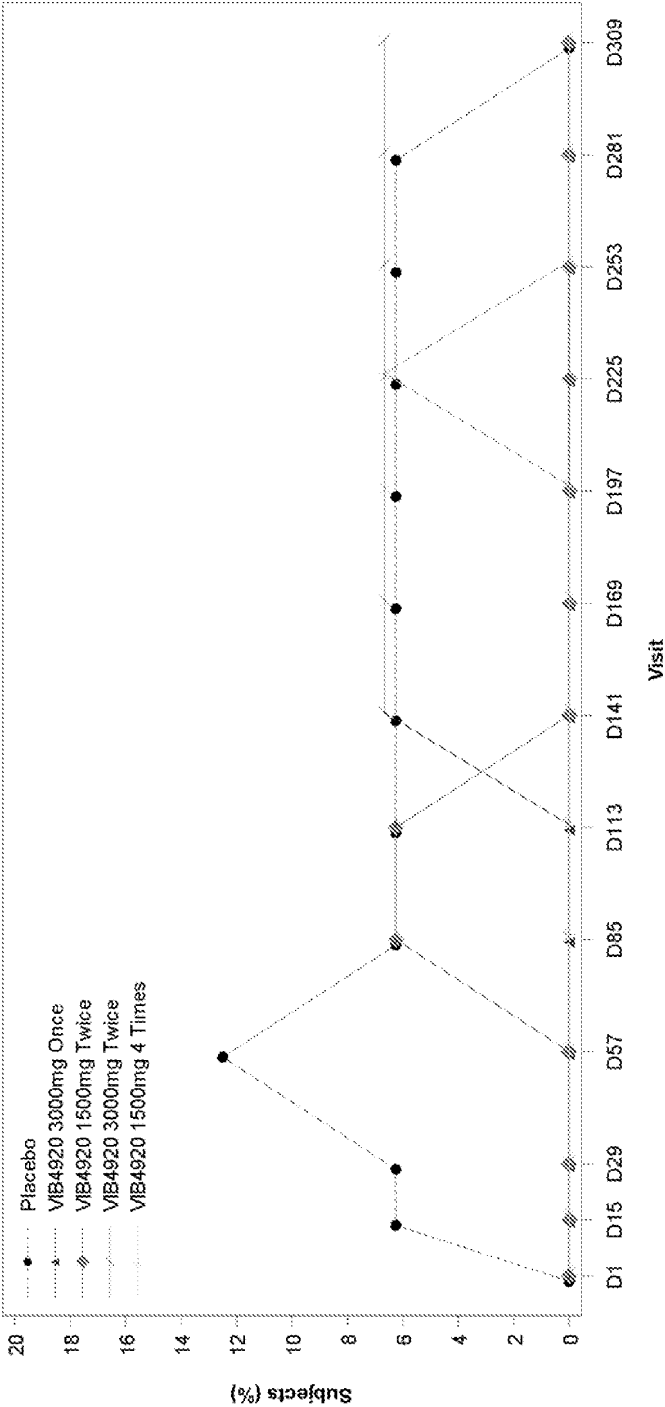


FIG. 14

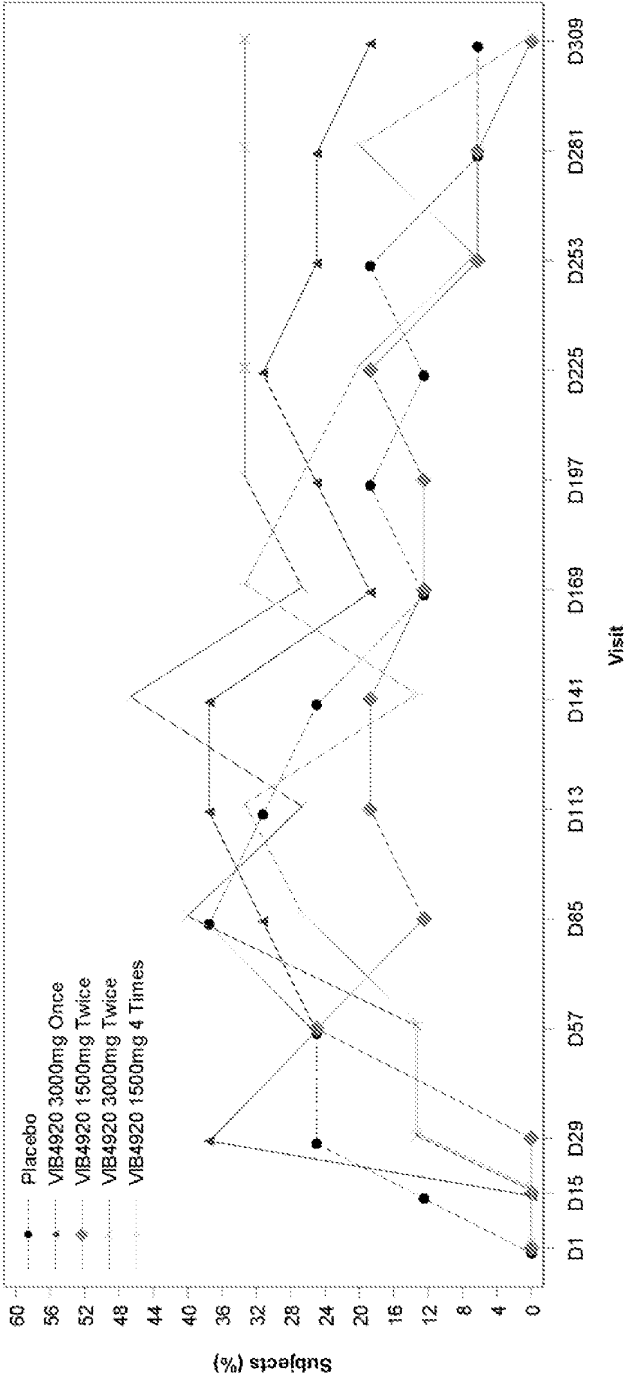


FIG. 15

	Placebo (N=16)	VIB4920 3000MG × 1 (N=16)	VIB4920 1500MG × 2 (N=16)	VIB4920 3000MG × 2 (N=15)	VIB4920 1500MG × 4 (N=15)
Rescued	1 (6.3%)	1 (6.3%)	1 (6.3%)	3 (20%)	1 (6.7%)
Censored	15 (93.8%)	15 (93.8%)	15 (93.8%)	12 (80%)	14 (93.3%)
Hazard Ratio		1.04 (0.10, 10.60)	1.04 (0.10, 10.60)	3.01 (0.45, 20.09)	1.04 (0.10, 10.60)
90% CI		0.9805	0.9805	0.3407	0.9805
p-value					

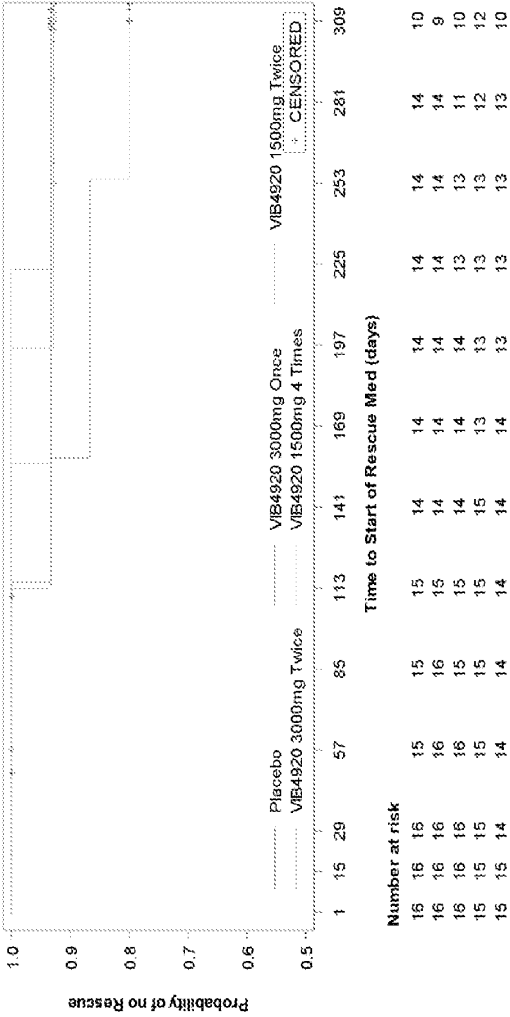


FIG. 16

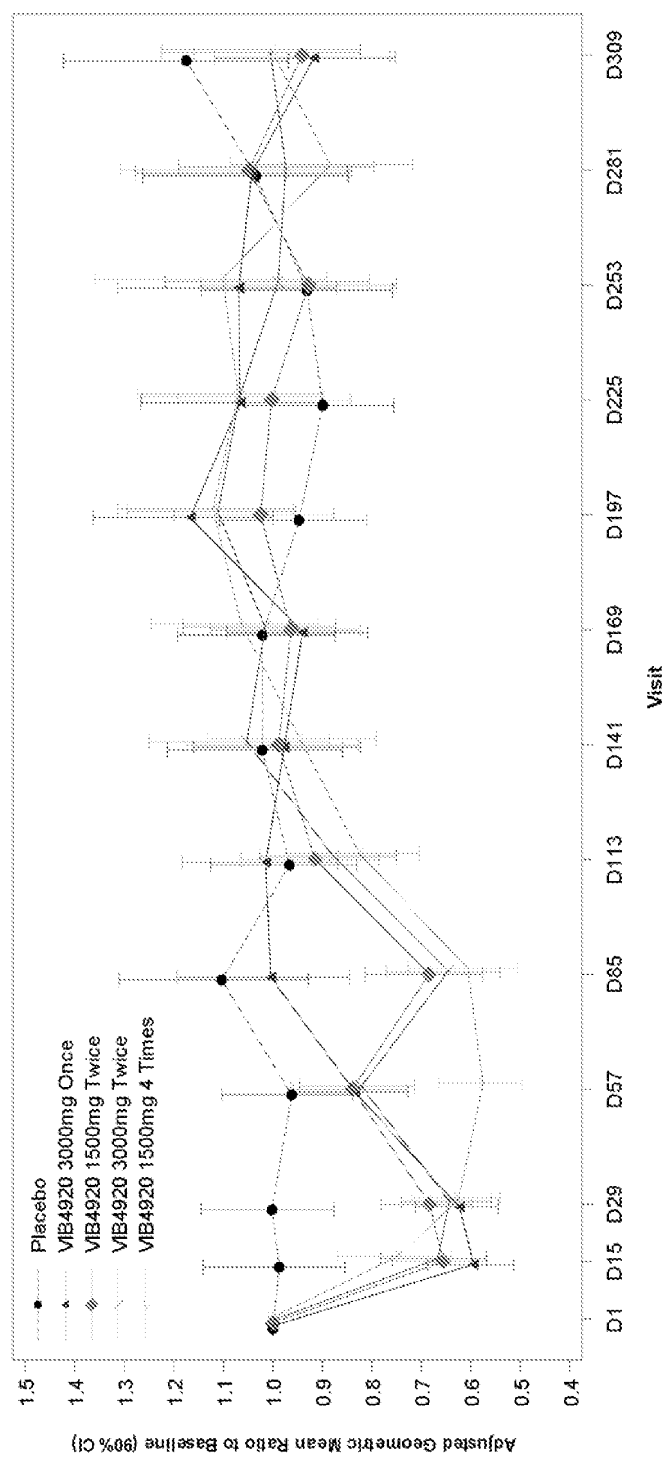


FIG. 17

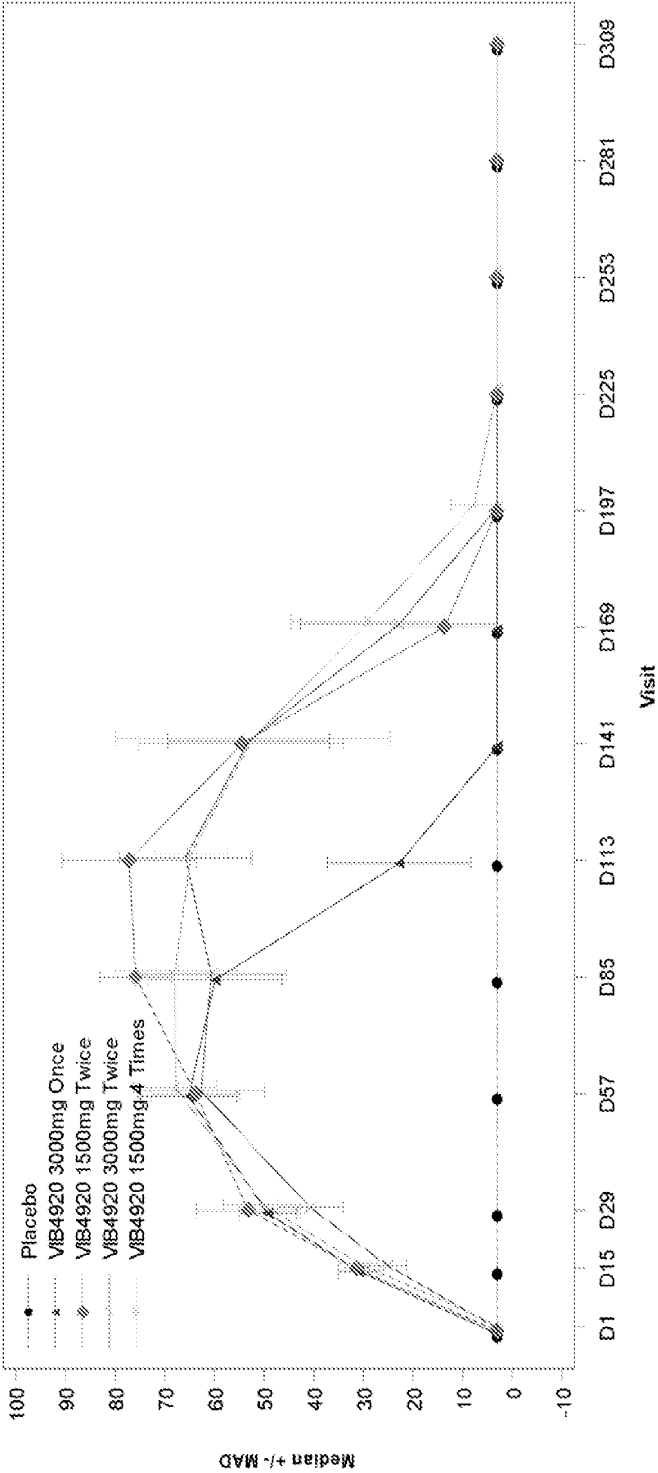


FIG. 18

Day 113									
Placebo		VIB4920 3000MG x 1		VIB4920 1500MG x 2		VIB4920 3000MG x 2		VIB4920 1500MG x 4	
(N=16)		(N=16)		(N=16)		(N=15)		(N=15)	
n	15	14	14	15	14	14	14	14	14
Geo. mean ratio to BSL	1.2	0.77	0.77	0.74	0.72	0.6	0.57	0.47	
Ratio of geo. mean vs P80		0.64	0.62	0.62	0.48	0.43	0.38	0.38	
90% CI		(0.52, 0.79)	(0.50, 0.76)	(0.48, 0.74)	(0.38, 0.59)				
p-value		0.0007	0.0003	0.0001	<0.0001				

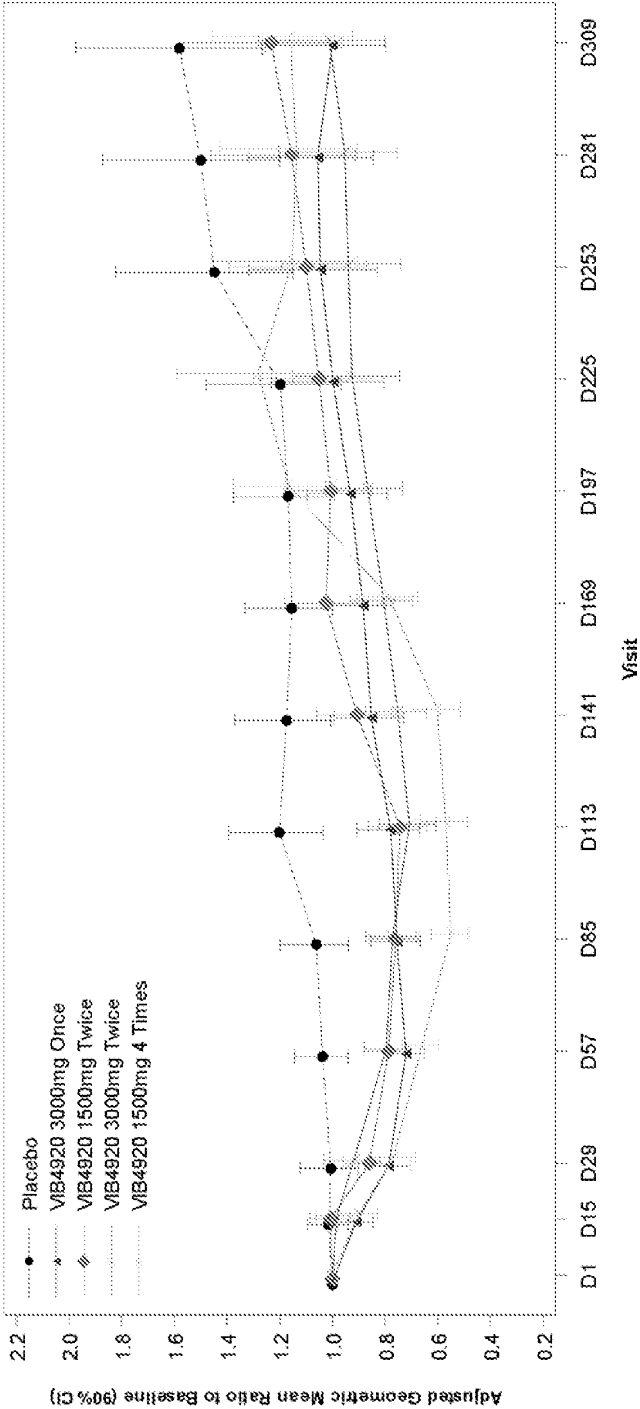


FIG. 19

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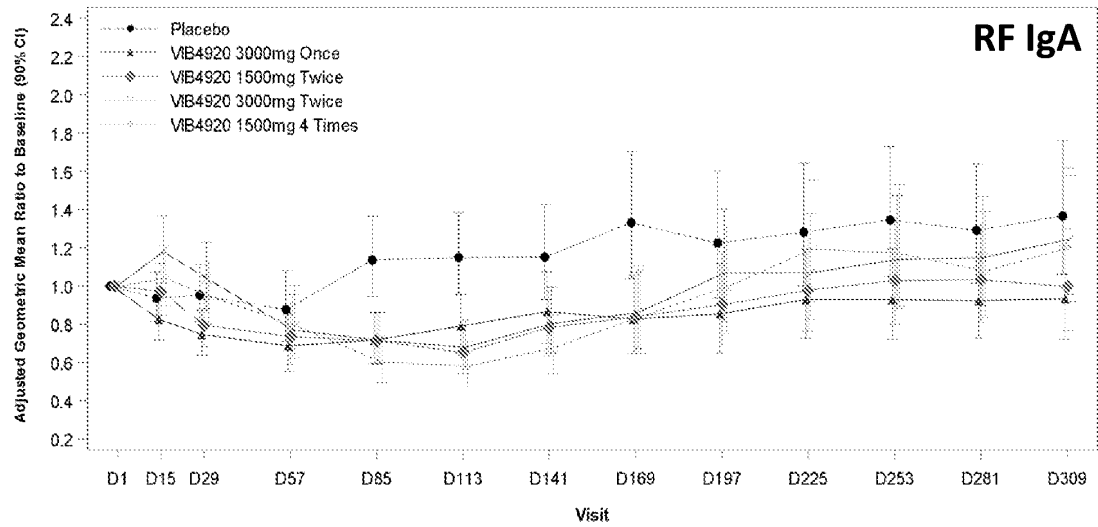


FIG. 20A

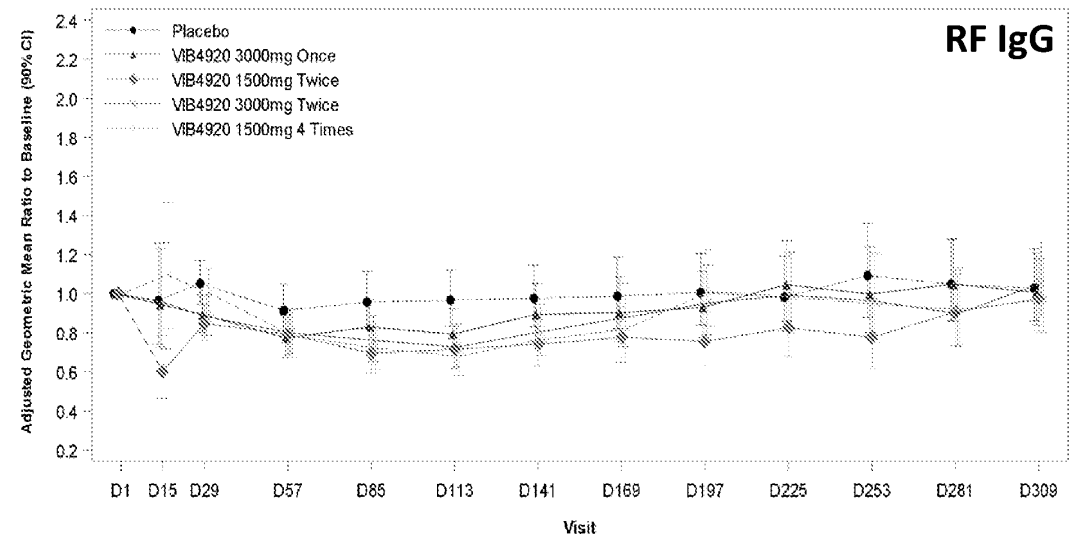


FIG. 20B

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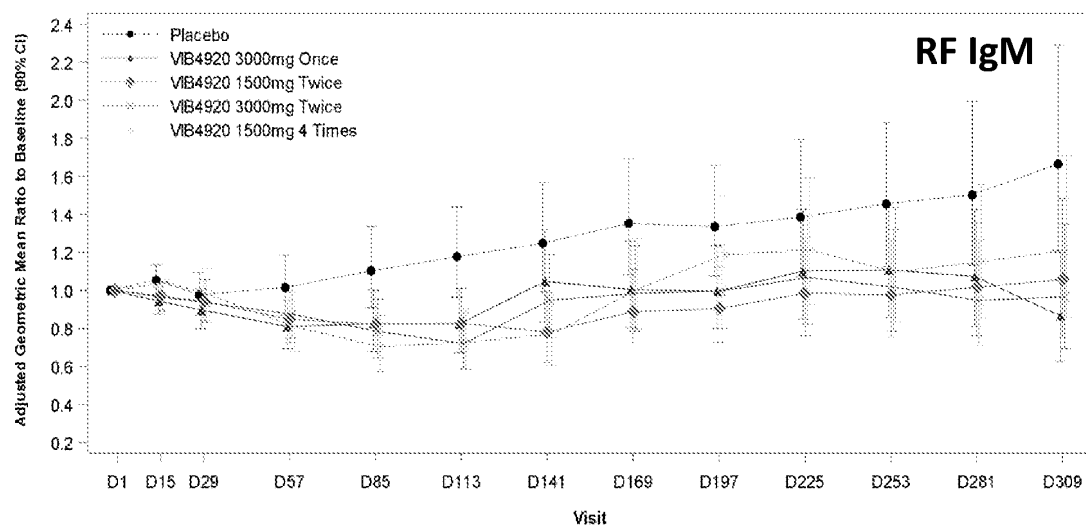


FIG. 20C

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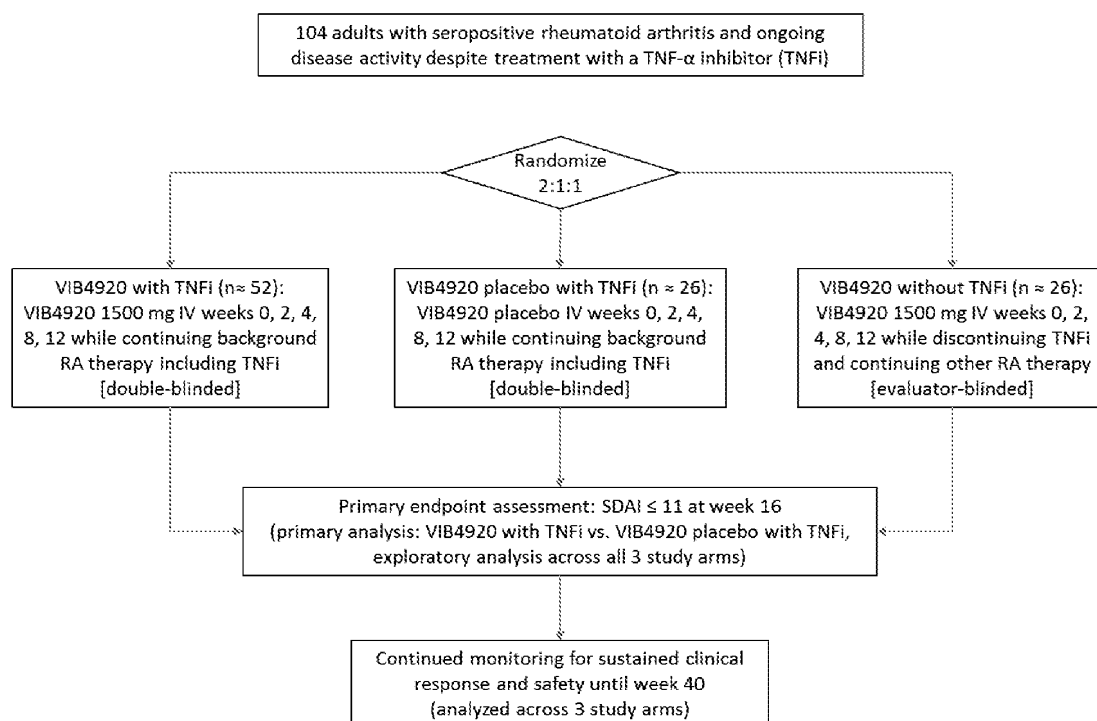


FIG. 21

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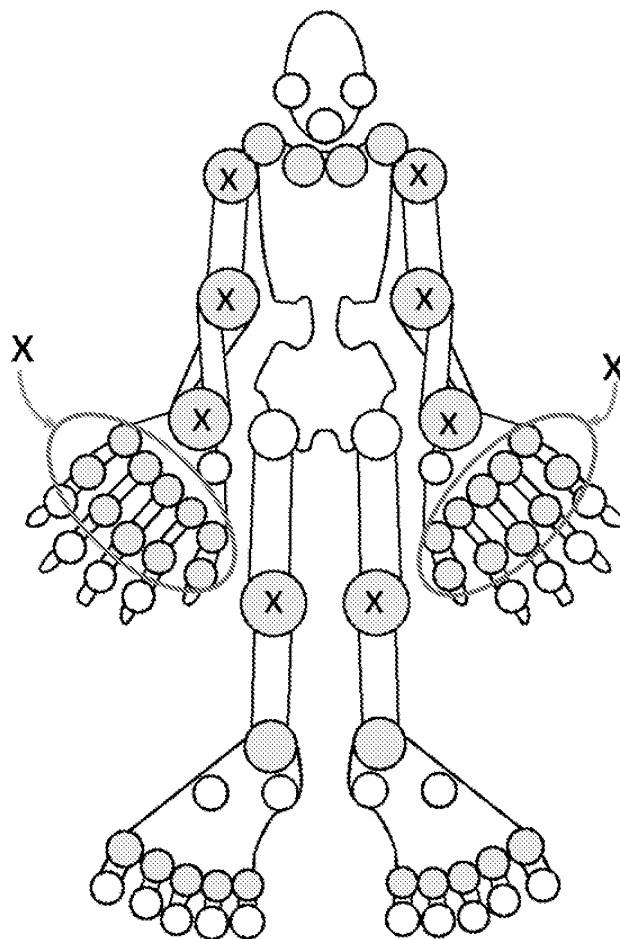


FIG. 22

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1. PLEASE CHECK THE ONE BEST ANSWER FOR YOUR ABILITIES AT THIS TIME.

OVER THE LAST WEEK, WERE YOU ABLE TO:	NEITHER DATE FORGETTABLE	WITH SOME DIFFICULTY	WITH MODERATE DIFFICULTY	EASIER TO DO
a. Drive yourself, including tying shoelaces and doing buttons?	0	1	2	3
b. Get in and out of bed?	0	1	2	3
c. Lift a full cup or glass to your mouth?	0	1	2	3
d. Walk outdoors on flat ground?	0	1	2	3
e. Wash and dry your entire body?	0	1	2	3
f. Bend down to pick up clothing from the floor?	0	1	2	3
g. Turn regular faucets on and off?	0	1	2	3
h. Get in and out of a car, bus, train, or airplane?	0	1	2	3
i. Walk two miles or three kilometers, if you wish?	0	1	2	3
j. Participate in recreational activities and sports as you would like, if you wish?	0	1	2	3
k. Get a good night's sleep?	0	3.1	2.2	3.3
l. Deal with feelings of anxiety or being nervous?	0	3.1	2.2	3.3
m. Deal with feelings of depression or feeling blue?	0	3.1	2.2	3.3

1. RAPID3 (0-10):

3.0-5 5.0-5.5
5.6-7 7.0-7.5
7.6-8 8.0-8.5
8.6-9 9.0-9.5
9.6-10 10.0-10.5

2. PAIN (0-10):

2.00 2.25 2.50 2.75 3.00 3.25 3.50 3.75 4.00 4.25 4.50 4.75 5.00 5.25 5.50 5.75 6.00 6.25 6.50 6.75 7.00 7.25 7.50 7.75 8.00 8.25 8.50 8.75 9.00 9.25 9.50 9.75 10.00

3. PAIN AS BAD AS IT COULD BE

2. HOW MUCH PAIN HAVE YOU HAD BECAUSE OF YOUR CONDITION OVER THE PAST WEEK?
PLEASE INDICATE BELOW HOW SEVERE YOUR PAIN HAS BEEN:

NO PAIN

0 0.5 1.0 1.5 2.0 2.5 3.0 3.5 4.0 4.5 5.0 5.5 6.0 6.5 7.0 7.5 8.0 8.5 9.0 9.5 10

PAIN AS BAD AS IT COULD BE

3. CONSIDERING ALL THE WAYS IN WHICH ILLNESS AND HEALTH CONDITIONS MAY AFFECT YOU
AT THIS TIME, PLEASE INDICATE BELOW HOW YOU ARE DOING:

VERY WELL

0 0.5 1.0 1.5 2.0 2.5 3.0 3.5 4.0 4.5 5.0 5.5 6.0 6.5 7.0 7.5 8.0 8.5 9.0 9.5 10

VERY POORLY

CONVERSION TABLE

New Remission (NR): 1.0-3.0 3.1-3.3
Low Severity (LS): 4.1-5.5 5.6-6.0
Moderate Severity (MS): 7.0-8.5 8.6-9.0 9.1-9.5 10.0-10.5 11.0-11.5 12.0-12.5

High Anxiety (HA): 13.0-13.5 14.0-14.5 15.0-15.5 16.0-16.5 17.0-17.5 18.0-18.5 19.0-19.5 20.0-20.5
21.0-21.5 22.0-22.5 23.0-23.5 24.0-24.5 25.0-25.5 26.0-26.5 27.0-27.5 28.0-28.5 29.0-29.5 30.0-30.5

HOW TO CALCULATE RAPID 3 SCORES

1. Ask the patient to complete questions 1, 2, and 3 while in the waiting room prior to his/her visit.

2. For question 1, add up the scores in questions A-J only (questions K-M have been added to be informative, but are not scored formally). Use the formula in the box on the right to calculate the formal score (0-10). For example, a patient whose answers total 19 would score a 6.5. Enter this score as an evaluation of the patient's functional status (FNC).

3. For question 2, enter the raw score (0-10) in the box on the right as an evaluation of the patient's pain tolerance (PTN).

4. For question 3, enter the raw score (0-10) in the box on the right as an evaluation of the patient's global estimate (PVE).

5. Add the total score (0-30) from questions 1, 2, and 3 and enter them as the patient's RAPID 3 cumulative score. Use the final conversion table to simplify the patient's weighted RAPID 3 score. For example, a patient who scores 11 on the cumulative RAPID 3 scale would score a weighted 3.7. A patient who scores between 0-1.0 is defined as near remission (NR); 1.1-2.0 as low severity (LS); 2.1-6.0 as moderate severity (MS); and 6.1-10.0 as high severity (HS).

FIG. 23A

Disease state	RAPID3 score
Remission	<3
Low disease severity	3.1 – 6.0
Moderate disease severity	6.1 – 12
High disease severity	12.1-30

FIG. 23B

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The STANFORD HEALTH ASSESSMENT QUESTIONNAIRE®
Stanford University School of Medicine, Division of Immunology & Rheumatology

HAQ Disability Index:

In this section we are interested in learning how your illness affects your ability to function in daily life. Please feel free to add any comments on the back of this page.

Please check the response which best describes your usual abilities **OVER THE PAST WEEK:**

	<u>Without ANY</u> <u>difficulty</u> ⁰	<u>With SOME</u> <u>difficulty</u> ¹	<u>With MUCH</u> <u>difficulty</u> ²	<u>UNABLE</u> <u>to do</u> ³
DRESSING & GROOMING				
Are you able to:				
-Dress yourself, including tying shoelaces and doing buttons?	☐	☐	☐	☐
-Shampoo your hair?	☐	☐	☐	☐
ARISING				
Are you able to:				
-Stand up from a straight chair?	☐	☐	☐	☐
-Get in and out of bed?	☐	☐	☐	☐
EATING				
Are you able to:				
-Cut your meat?	☐	☐	☐	☐
-Lift a full cup or glass to your mouth?	☐	☐	☐	☐
-Open a new milk carton?	☐	☐	☐	☐
WALKING				
Are you able to:				
-Walk outdoors on flat ground?	☐	☐	☐	☐
-Climb up five steps?	☐	☐	☐	☐

Please check any **AIDS OR DEVICES** that you usually use for any of these activities:

- | | |
|-------------------------------------|---|
| ☐ Cane | ☐ Devices used for dressing (button hook, zipper pull, long-handled shoe horn, etc.) |
| ☐ Walker | ☐ Built up or special utensils |
| ☐ Crutches | ☐ Special or built up chair |
| ☐ Wheelchair | ☐ Other (Specify: _____) |

Please check any categories for which you usually need **HELP FROM ANOTHER PERSON:**

- | | |
|--|----------------------------------|
| ☐ Dressing and Grooming | ☐ Eating |
| ☐ Arising | ☐ Walking |

FIG. 24

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Please check the response which best describes your usual abilities **OVER THE PAST WEEK:**

	Without ANY difficulty ³	With SOME difficulty ¹	With MUCH difficulty ²	UNABLE to do ²
HYGIENE				
Are you able to:				
-Wash and dry your body?	☐	☐	☐	☐
-Take a tub bath?	☐	☐	☐	☐
-Get on and off the toilet?	☐	☐	☐	☐
REACH				
Are you able to:				
-Reach and get down a 5-pound object (such as a bag of sugar) from just above your head?	☐	☐	☐	☐
-Bend down to pick up clothing from the floor?	☐	☐	☐	☐
GRIP				
Are you able to:				
-Open car doors?	☐	☐	☐	☐
-Open jars which have been previously opened?	☐	☐	☐	☐
-Turn faucets on and off?	☐	☐	☐	☐
ACTIVITIES				
Are you able to:				
-Run errands and shop?	☐	☐	☐	☐
-Get in and out of a car?	☐	☐	☐	☐
-Do chores such as vacuuming or yardwork	☐	☐	☐	☐

Please check any **AIDS OR DEVICES** that you usually use for any of these activities:

- | | |
|--|--|
| ☐ Raised toilet seat | ☐ Bathtub bar |
| ☐ Bathtub seat | ☐ Long-handled appliances for reach |
| ☐ Jar opener (for jars previously opened) | ☐ Long-handled appliances in bathroom |
| | ☐ Other (Specify: _____) |

Please check any categories for which you usually need **HELP FROM ANOTHER PERSON:**

- | | |
|----------------------------------|--|
| ☐ Hygiene | ☐ Gripping and opening things |
| ☐ Reach | ☐ Errands and chores |

We are also interested in learning whether or not you are affected by pain because of your illness.

How much pain have you had because of your illness **IN THE PAST WEEK:**

PLACE A VERTICAL (|) MARK ON THE LINE TO INDICATE THE SEVERITY OF THE PAIN

No Pain	Severe Pain
0	100

Considering all the ways that your arthritis affects you, rate how you are doing on the following scale by placing a vertical mark on the line.

Very Well	Very Poor
0	100

FIG. 24 Continued

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PROMIS-29 Profile v2.1

Please respond to each question or statement by marking one box per row.

Physical Function		Without any difficulty	With a little difficulty	With some difficulty	With much difficulty	Unable to do
PF11	Are you able to do chores such as vacuuming or yard work?	☐ 5	☐ 4	☐ 3	☐ 2	☐ 1
PF12	Are you able to go up and down stairs at a normal pace?	☐ 5	☐ 4	☐ 3	☐ 2	☐ 1
PF13	Are you able to go for a walk of at least 15 minutes?	☐ 5	☐ 4	☐ 3	☐ 2	☐ 1
PF14	Are you able to run errands and shop?	☐ 5	☐ 4	☐ 3	☐ 2	☐ 1
Anxiety						
In the past 7 days...		Never	Rarely	Sometimes	Often	Always
EDAN11	I felt fearful	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
EDAN12	I found it hard to focus on anything other than my anxiety	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
EDAN13	My worries overwhelmed me	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
EDAN14	I felt uneasy	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
Depression						
In the past 7 days...		Never	Rarely	Sometimes	Often	Always
EDDE11	I felt worthless	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
EDDE12	I felt helpless	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
EDDE13	I felt depressed	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
EDDE14	I felt hopeless	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
Fatigue						
During the past 7 days...		Not at all	A little bit	Somewhat	Quite a bit	Very much
FI1	I feel fatigued	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
FI2	I have trouble starting things because I am tired	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5

FIG. 25

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PROMIS-29 Profile v2.1

<u>Fatigue</u>						
In the past 7 days...		Not at all	A little bit	Somewhat	Quite a bit	Very much
FATFPM1	How run-down did you feel on average? ...	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
FATFPM2	How fatigued were you on average?	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
<u>Sleep Disturbance</u>						
In the past 7 days...		Very poor	Poor	Fair	Good	Very good
SLEEP12	My sleep quality was	☐ 5	☐ 4	☐ 3	☐ 2	☐ 1
In the past 7 days...		Not at all	A little bit	Somewhat	Quite a bit	Very much
SLEEP13	My sleep was refreshing	☐ 5	☐ 4	☐ 3	☐ 2	☐ 1
SLEEP20	I had a problem with my sleep	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
SLEEP41	I had difficulty falling asleep	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
<u>Ability to Participate in Social Roles and Activities</u>						
		Never	Rarely	Sometimes	Usually	Always
GRPERF11 _CAPS	I have trouble doing all of my regular leisure activities with others	☐ 5	☐ 4	☐ 3	☐ 2	☐ 1
GRPERF16 _CAPS	I have trouble doing all of the family activities that I want to do	☐ 5	☐ 4	☐ 3	☐ 2	☐ 1
GRPERF23 _CAPS	I have trouble doing all of my usual work (include work at home)	☐ 5	☐ 4	☐ 3	☐ 2	☐ 1
GRPERF40 _CAPS	I have trouble doing all of the activities with friends that I want to do	☐ 5	☐ 4	☐ 3	☐ 2	☐ 1
<u>Pain Interference</u>						
In the past 7 days...		Not at all	A little bit	Somewhat	Quite a bit	Very much
PAIN15B	How much did pain interfere with your day to day activities?	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
PAIN15C2	How much did pain interfere with work around the home?	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
PAIN15C3	How much did pain interfere with your ability to participate in social activities?	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5
PAIN15C4	How much did pain interfere with your household chores?	☐ 1	☐ 2	☐ 3	☐ 4	☐ 5

FIG. 25 Continued

	% improvement from baseline needed to achieve:		
	ACR20 response	ACR50 response	ACR70 response
Tender joint count	At least 20%	At least 50%	At least 70%
Swollen joint count	At least 20%	At least 50%	At least 70%
Patient global assessment			
Physician global assessment			
Assessment of functional ability (HAQ-DI)	At least 20% in 3 out of these 5 measures	At least 50% in 3 out of these 5 measures	At least 70% in 3 out of these 5 measures
Pain assessment			
C-reactive protein			

FIG. 26

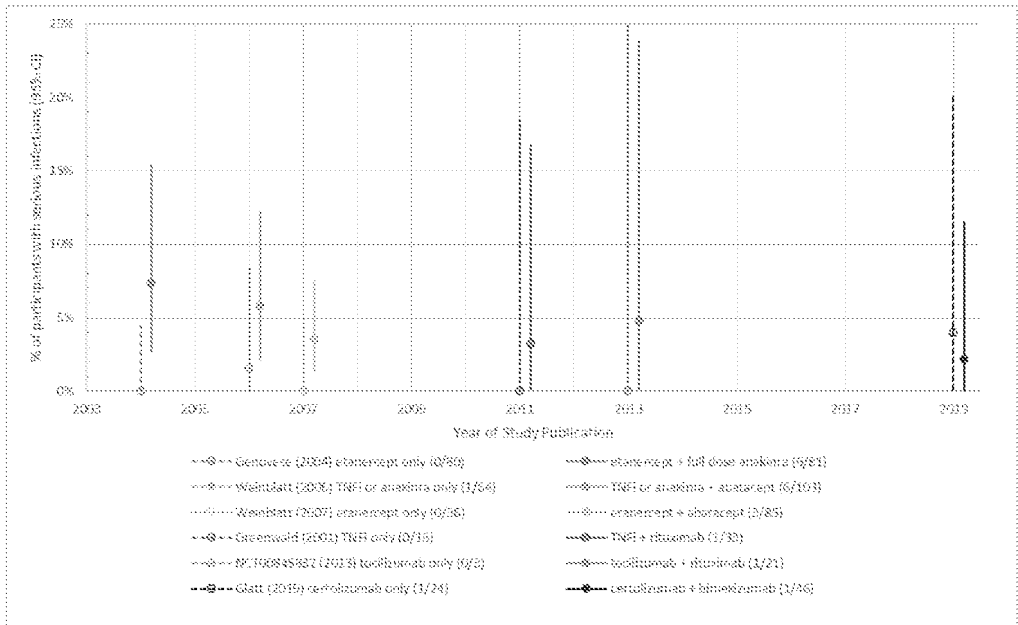


FIG. 27