

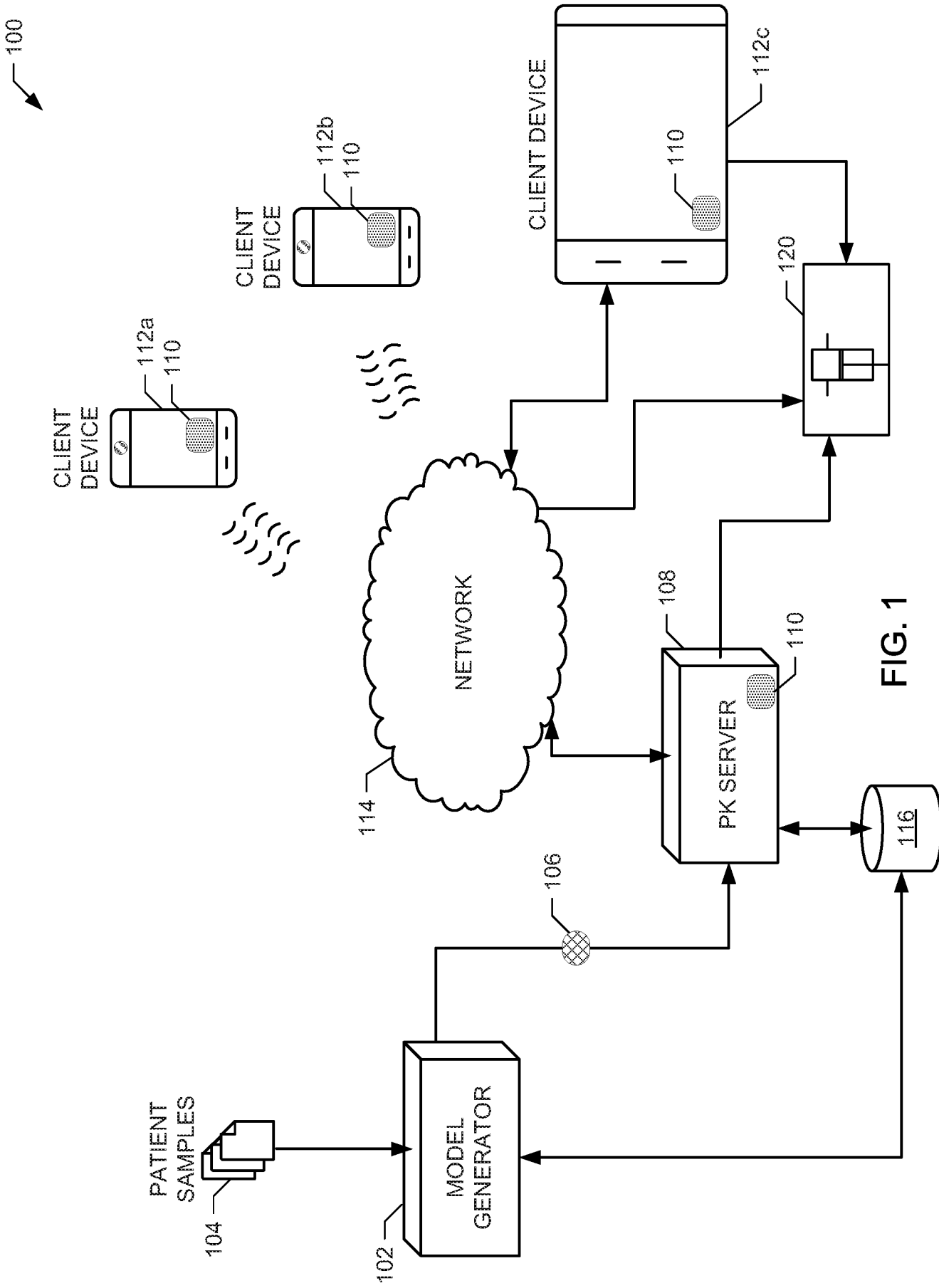


FIG. 1

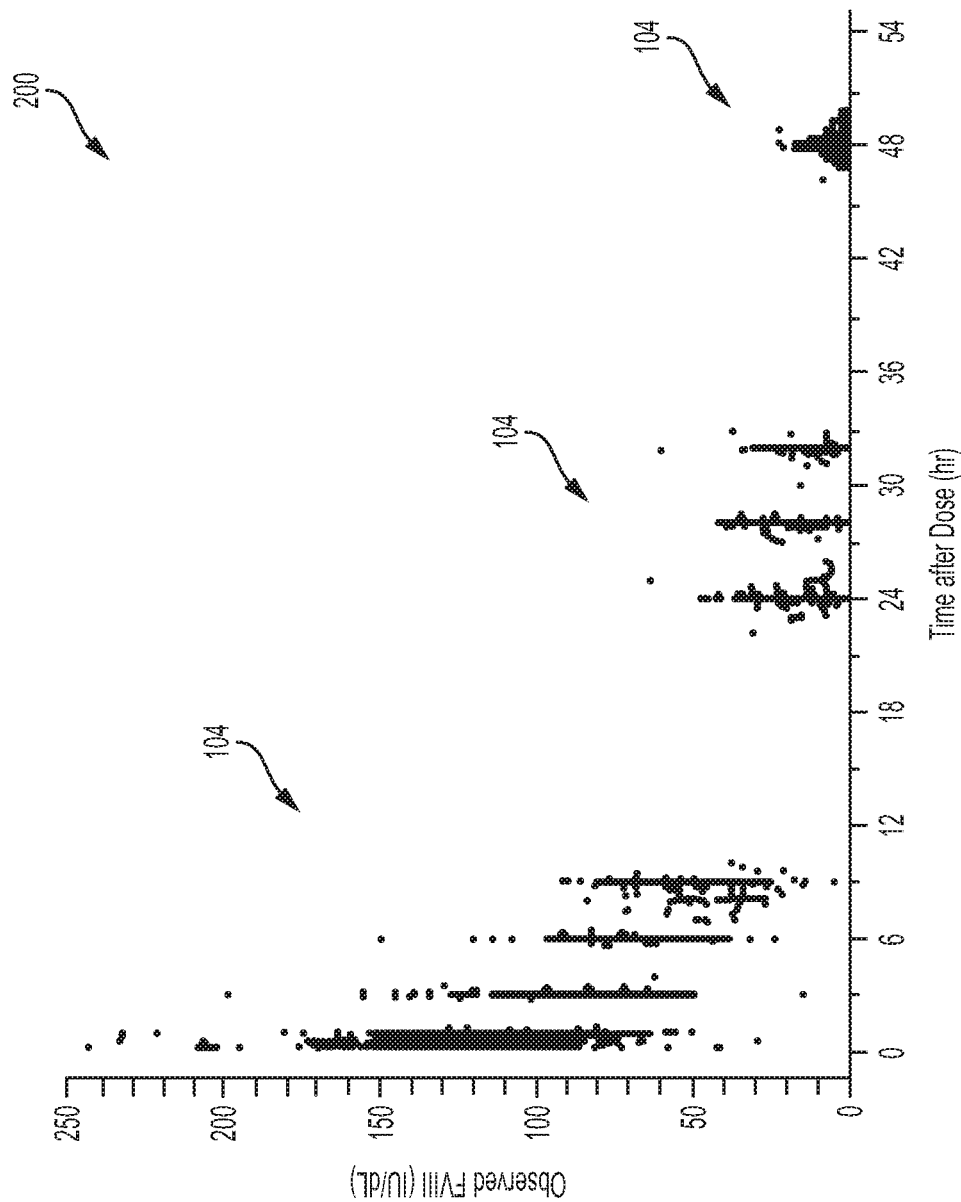


FIG. 2

300

Login

First Name *

Last Name *

Specialty *

Hospital Name

Address *

City *

State *

IL

Zip Code *

Country *

United States

Email *

Repeat Email *

Day Time Phone

Mobile Number

Fax Number

How did you hear about us?

Newspaper

DEA Number *

☐ I confirm that my DEA Number may be used to verify my identity and i am a registered Healthcare Professional *

☐

☒

FIG. 3

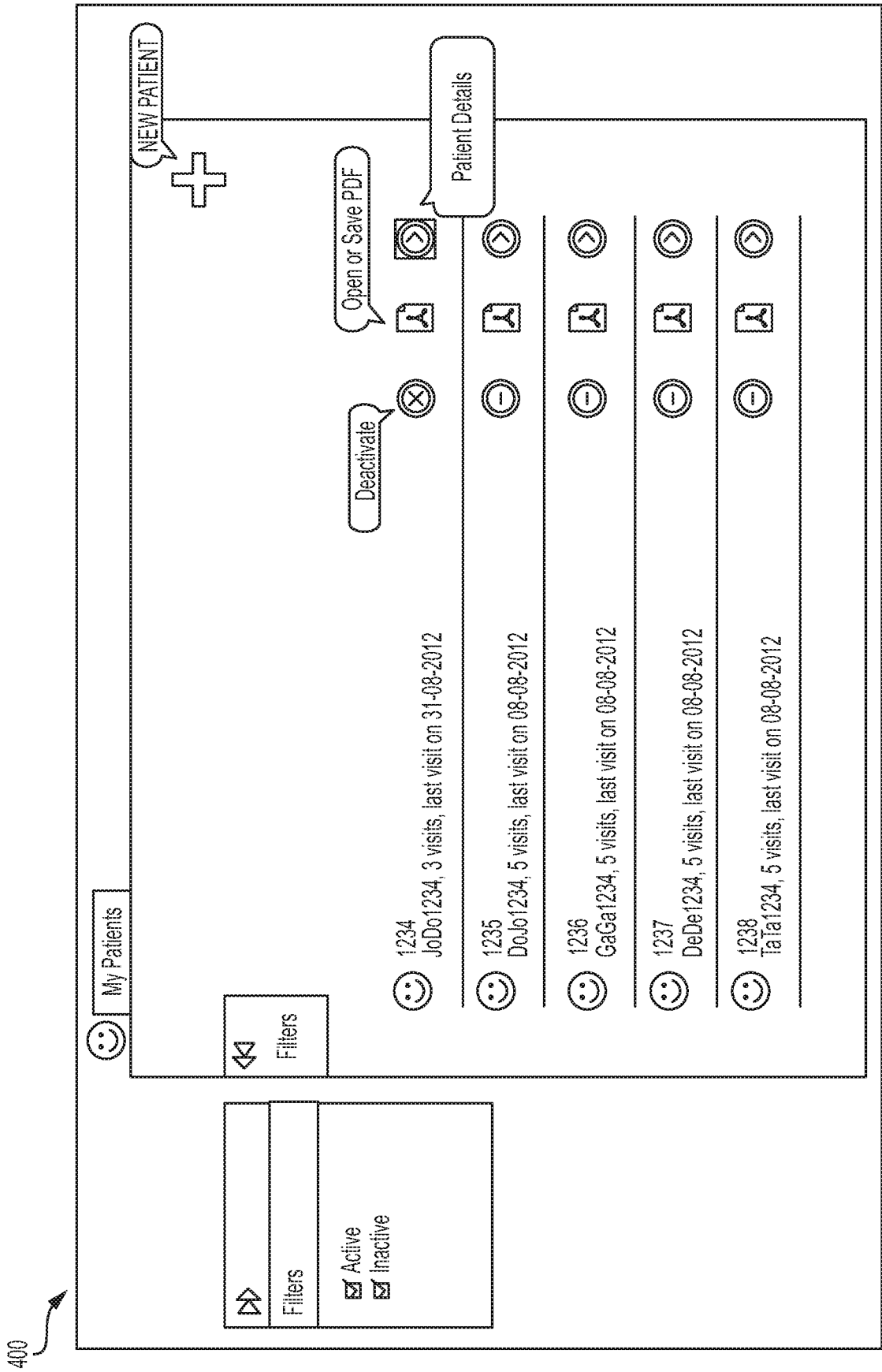


FIG. 4

500

Visit# A123-004 for Patient JoDo1234 with ID# 1234

General Patient Information			
Patient Name *	JoDo1234	Wash Out	Yes No
Body Weight *	1200 Kg	Pre-infusion Levels	IU / Kg
Please enter a correct Patient Weight In Kg			
Birth Date *	12/01/1960		
Age: 0 year 0 month			
Dose for PK infusion			IU / KG
Date of PK FVIII infusion	/ /	00:00AM	
Next Step			
Patient History			
Calculation			
Dosage			
Visit History			

FIG. 5

600

SAMPLES

Washout

Yes

No

Pre-infusion FVIII level (IU/dL)

PK Assessment	Sample	Last Edit	Collection Date & Time	Hours After	FVIII Level (IU/dL)	Assay Type	Included	Excluded	
1	1	08-May-2014 01:47	06-May-2014 16:00	6.00	57.0	One-Stage	Included	Excluded	
1	2	08-May-2014 01:47	07-May-2014 10:00	24.00	15.0	One-Stage	Included	Excluded	
1	3	08-May-2014 01:47	07-May-2014 16:00	30.00	10.0	One-Stage	Included	Excluded	

Add Sample

Reminder: PK profiles should only be generated using the same assay type.

	Bayesian Estimate
Clearance (dL/hr/kg)	0.035
Volume in steady state (dL/kg)	0.5
FVIII half-life (hours)	11.8
Time to 1% above baseline (hours)	70.0

Next Step

FIG. 6

700

General Patient Information

Patient History

Calculation

Visit#	Sample#	Collectoin Date/Time	Observed IU/dl	Theoretical IU/dl	Adjusted IU/dl	
A123-001	01	20-12-2011 8:00am	5	4.86	4.66	ON
A123-002	01	01-08-2012 8:00am	5	4.86	4.66	ON
A123-002	02	05-08-2012 8:00am	5	4.83	4.50	ON

Add sample

	Theoretical	Adjusted	Offset
Clearance (L/hr)			
vdBeta			
cMax / Peak (IU/dL)			
FV/III half-life (hours)			
Time to 1% (hours)			

Calculate

Next Step

Dosage

Visit History

FIG. 7

700

General Patient Information

Patient History

Calculation

Visit#	Sample#	Collection Date/Time	Observed IU/dL	Theoretical IU/dL	Adjusted IU/dL	
A123-001	01	20-12-2011 8:00am	5	4.86	4.66	ON
A123-002	01	01-08-2012 8:00am	5	4.86	4.66	ON
A123-002	02	05-08-2012 8:00am	5	4.83	4.50	OFF
A123-004	01	05-08-2012 8:00am	10	9.86	9.66	ON

Add sample

	Theoretical	Adjusted	Offset
Clearance (L/hr)	0.68	0.67	0.01
vdBeta	11.86	11.86	0
cMax / Peak (IU/dL)	20.65	16.86	3.79
FVIII half-life (hours)	12.07	12.18	-0.11
Time to 1% (hours)	1.93	1.82	0.11

Calculate

Next Step

Dosage

Visit History

FIG. 8

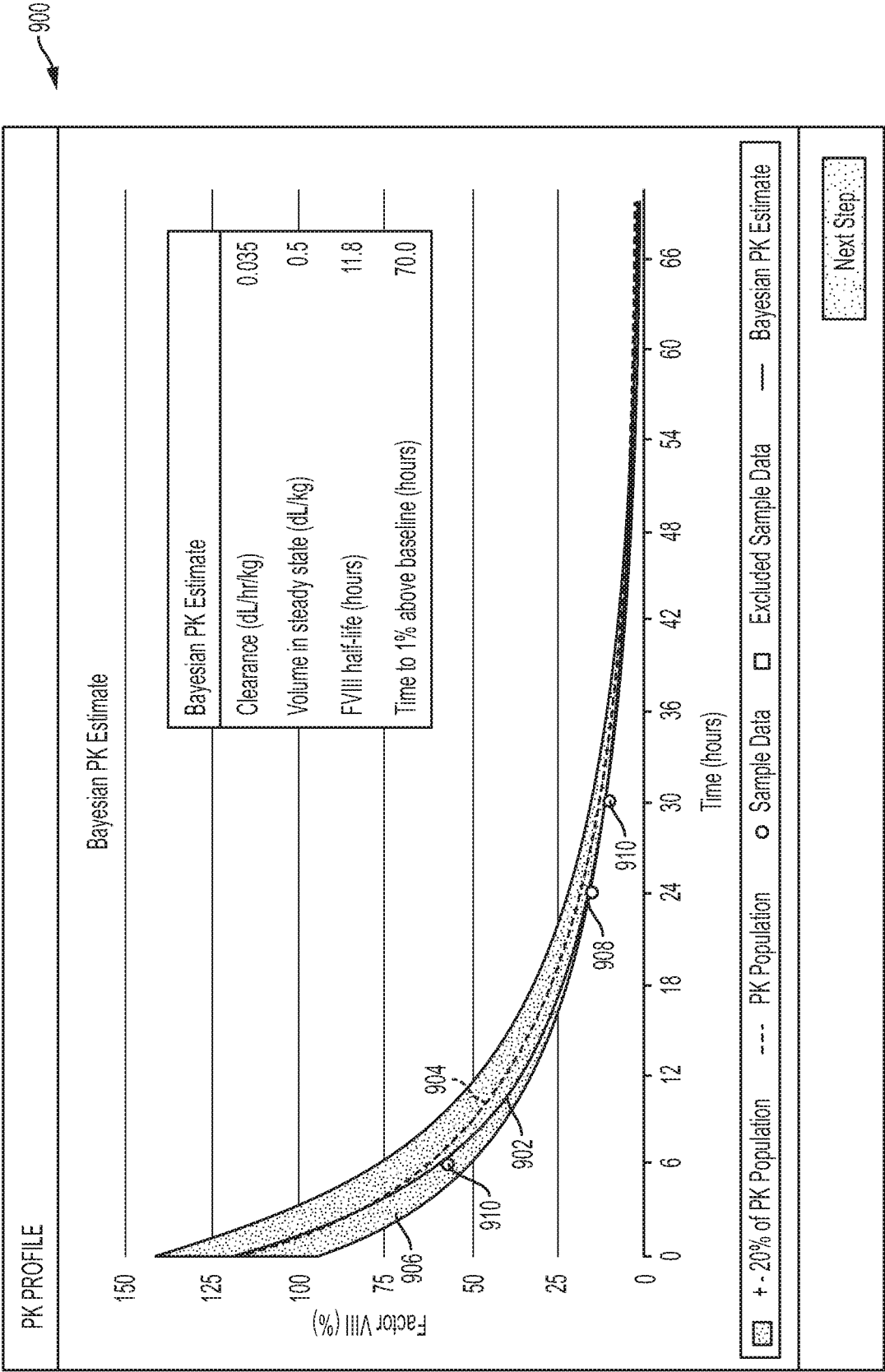


FIG. 9

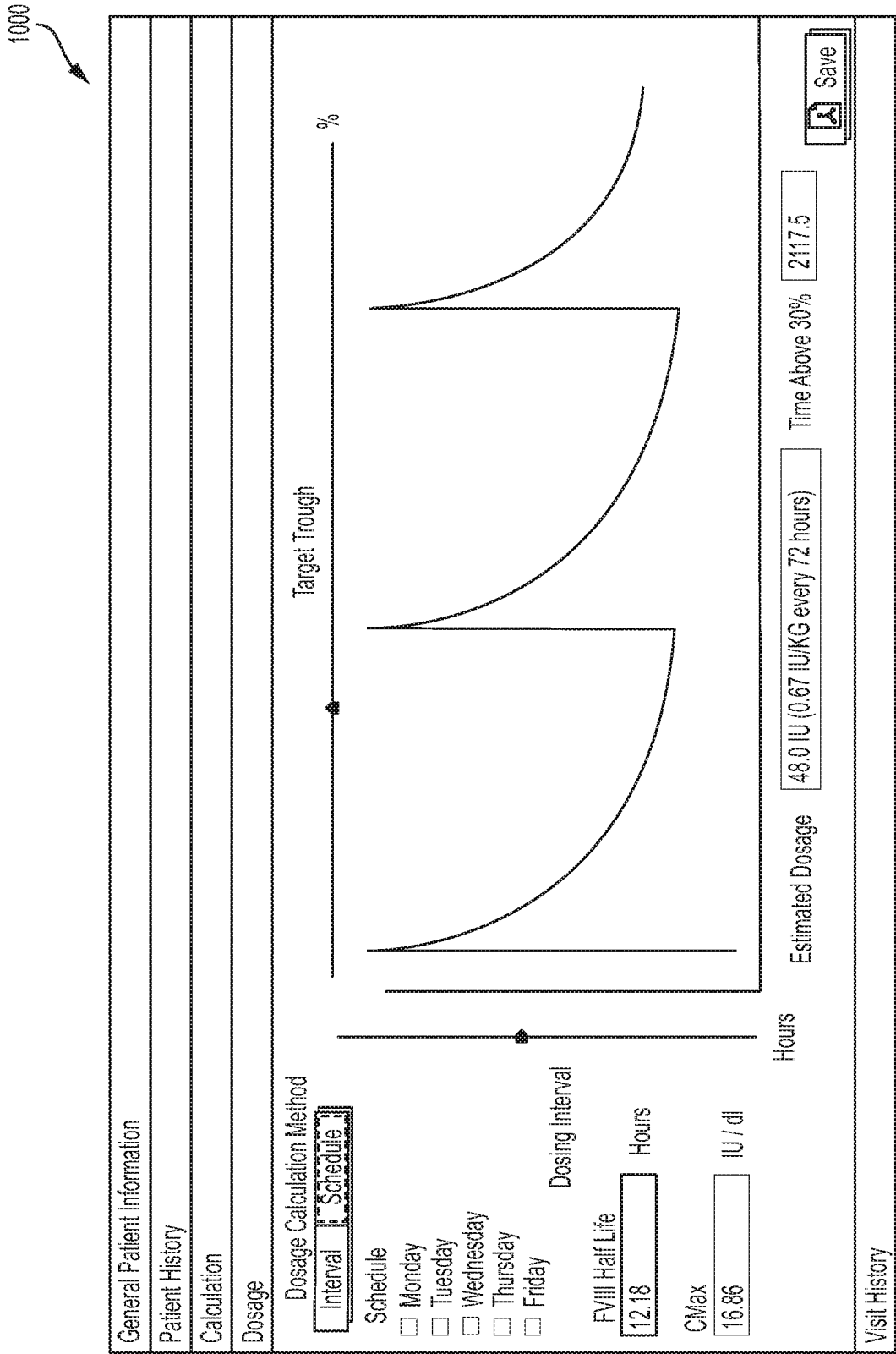


FIG. 10

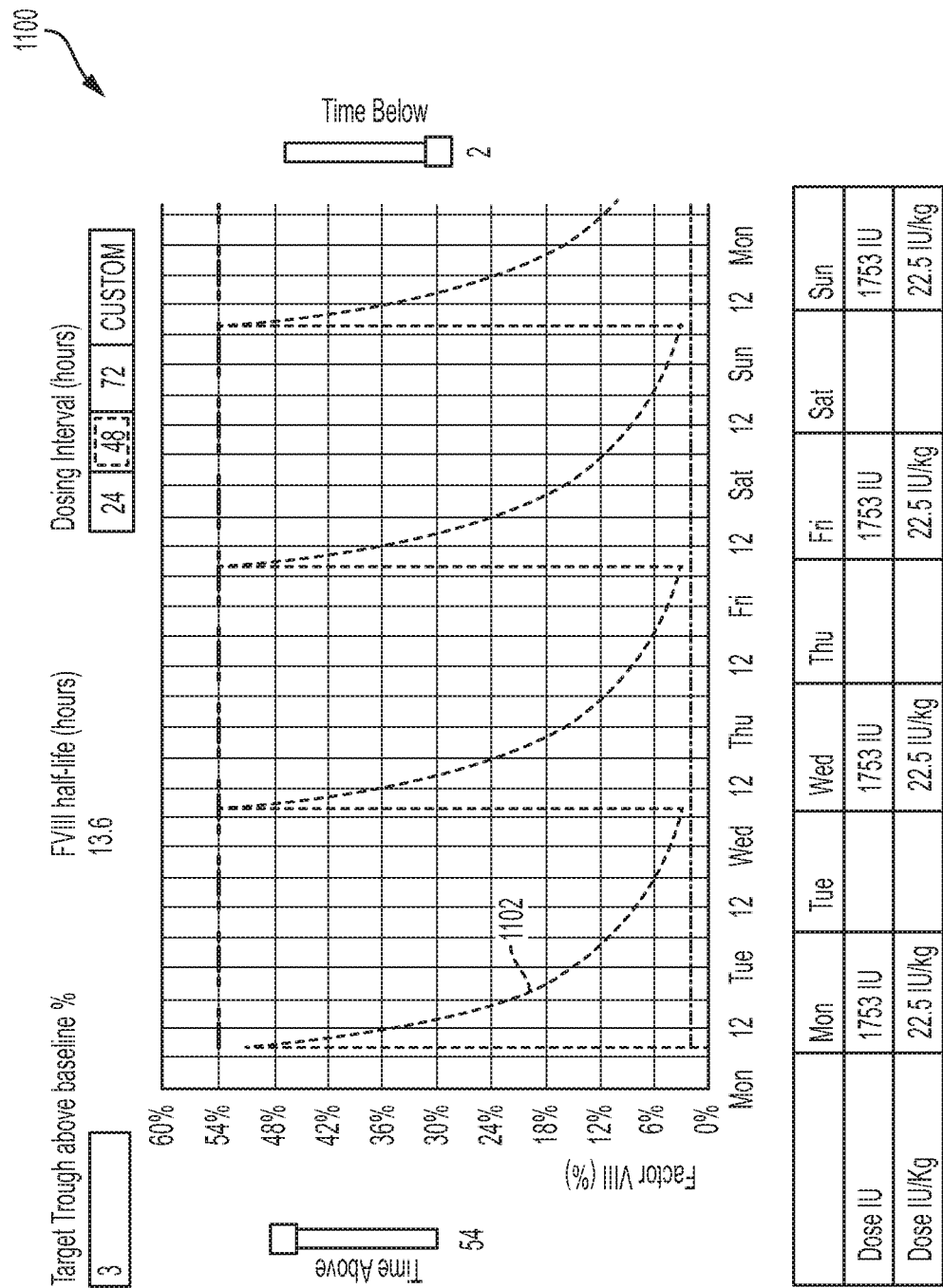


FIG. 11

1200

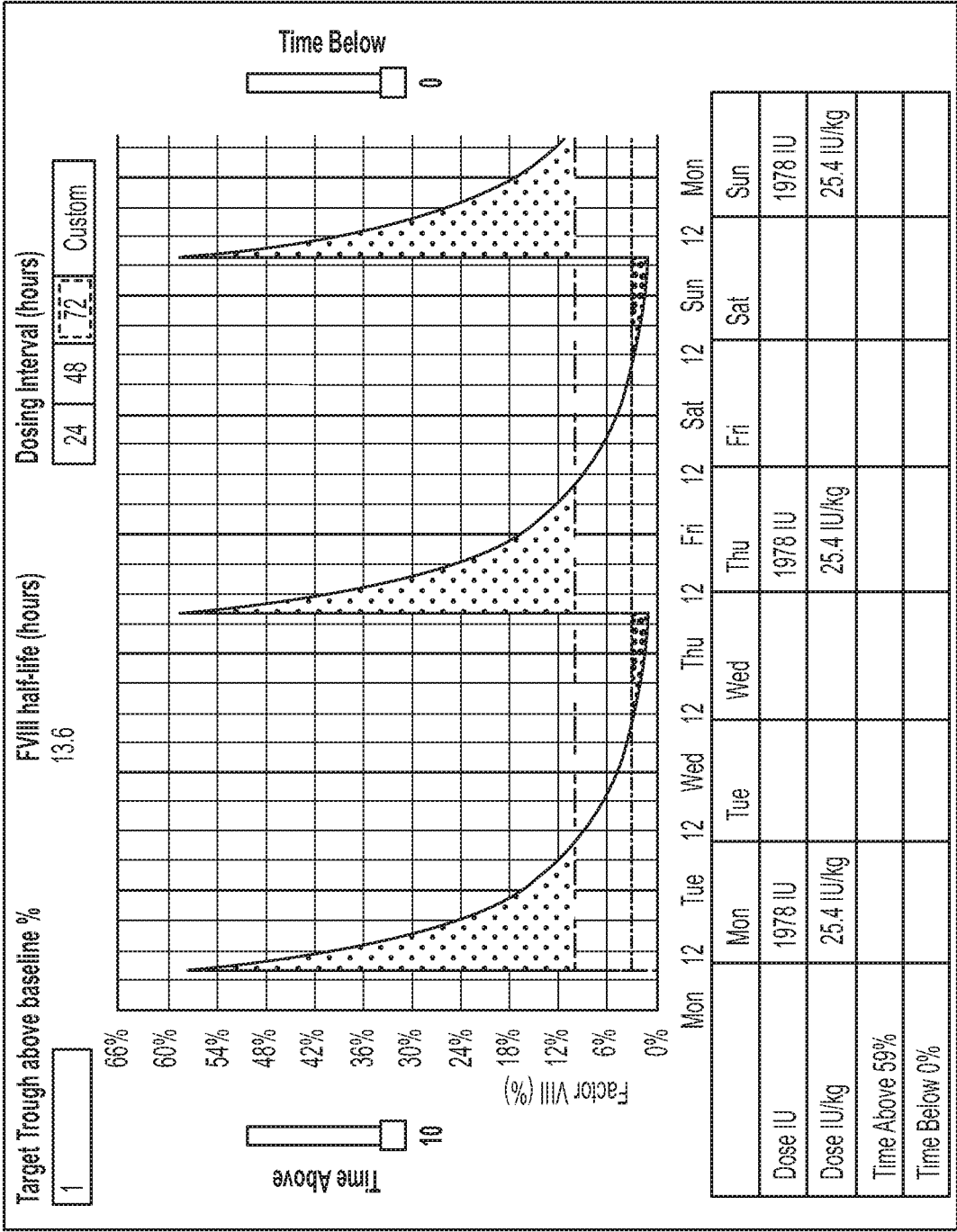


FIG. 12

1300

DETAILS FOR VISIT A123-002

Patient Name JoDo1234	Current Body Weight 80 Kg	Wash Out ☒ YES ☐ NO	Dose for PK Infusion
Birth Date * 12/01/1960	Age: 53 years 2 months 1 day	Pre-Infusion Levels 	Date of PK FVIII Infusion / / 00:00AM

Visit#		Collectoin Date/Time	Observed IU/dl	Theoretical IU/dl	Adjusted IU/dl
A123-002	01	01-08-2012 8:00am	5	4.86	4.66
A123-002	02	05-08-2012 8:00am	5	4.83	4.50

	Theoretical	Adjusted	Offset
Clearance (L/hr)	0.68	0.67	0.01
vdBeta	11.86	11.86	0
cMaz / Peak (IU/dL)	20.65	16.86	3.79
FVIII half-life (hours)	12.07	12.18	-0.11
Time to 1% (hours)	1.93	1.82	0.11

Dosage Calculation Method
☒ Interval ☐ Schedule

Schedule
☐ Monday
☐ Tuesday
☐ Wednesday
☐ Thursday
☐ Friday

Dosing Interval

FVIII Half Life
 12.18 Hours

CMaz
 16.86 IU / dl

Estimated Dosage 48.0 IU (0.67 IU/KG every 72 hours) Time Above 30% 2117.5

FIG. 13

1400

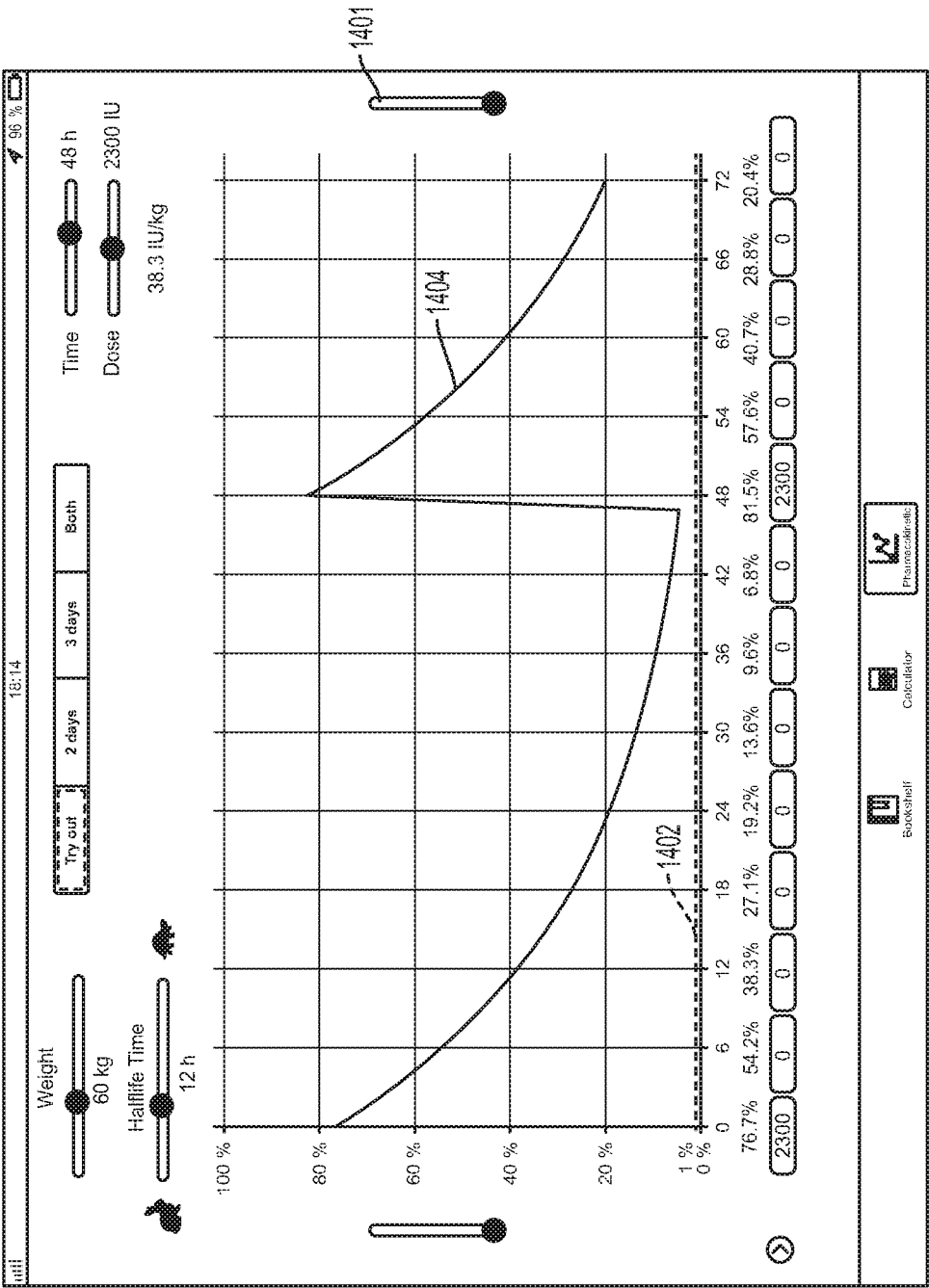


FIG. 14

1400

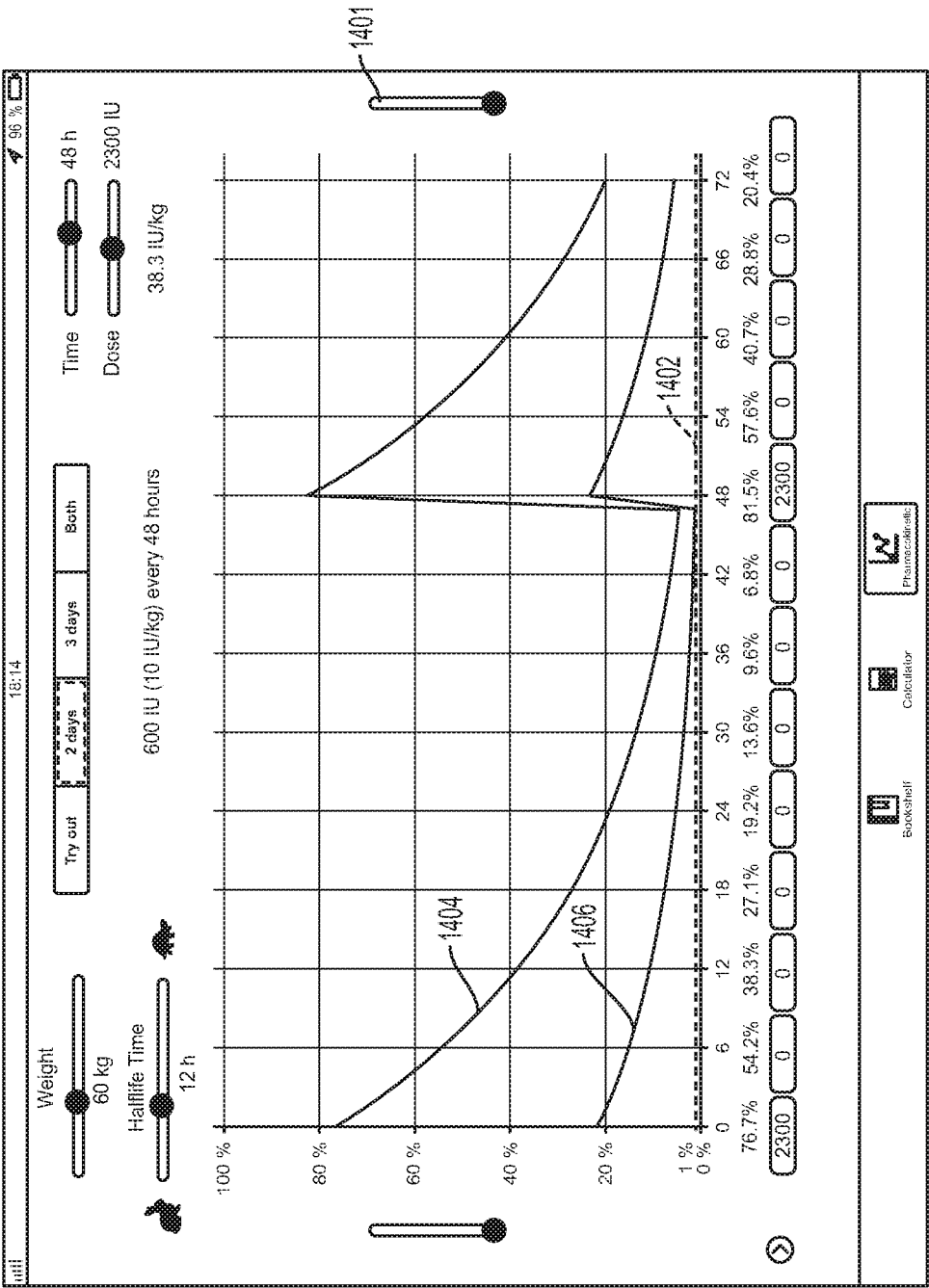


FIG. 15

1400

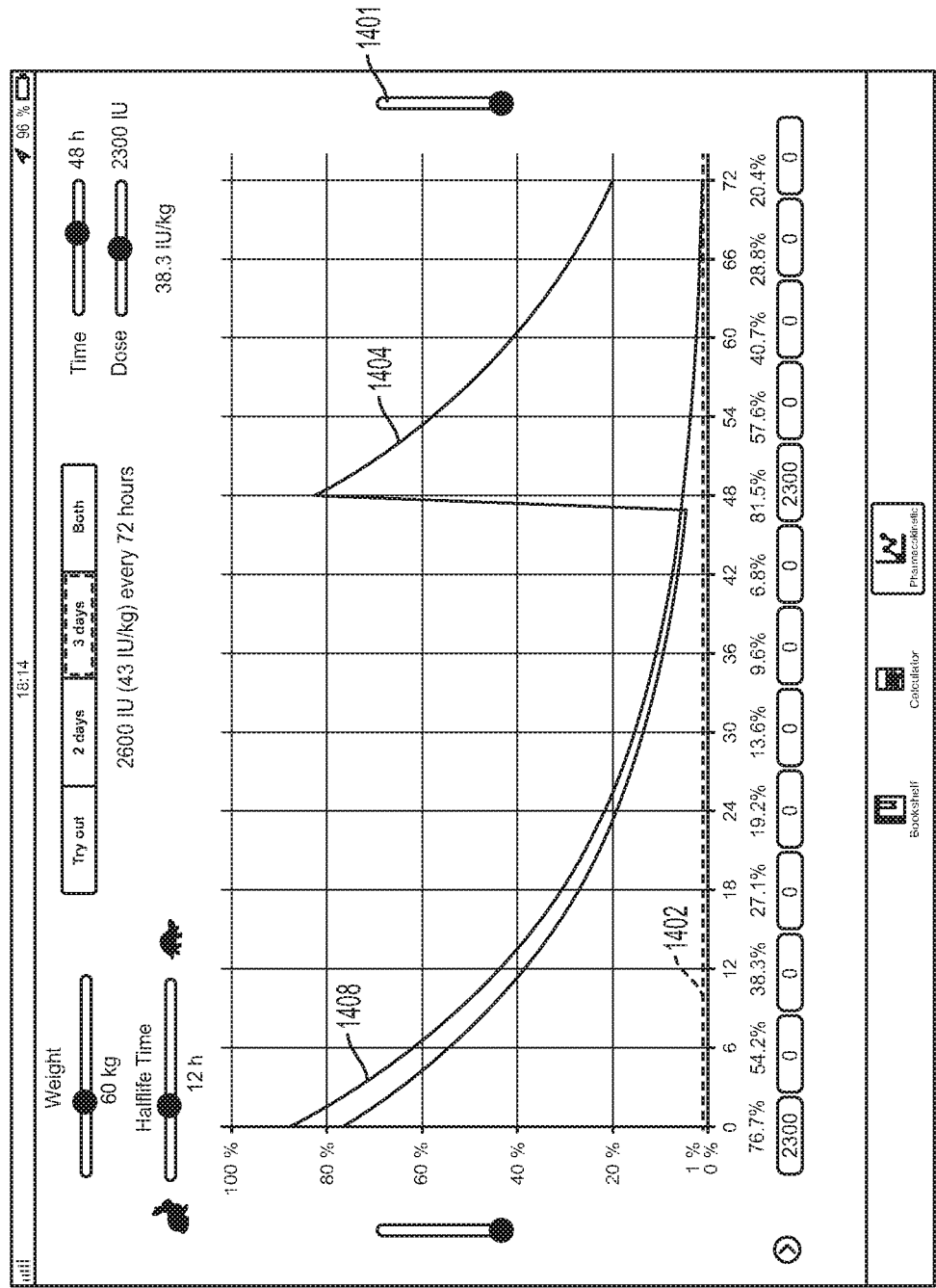


FIG. 16

1400

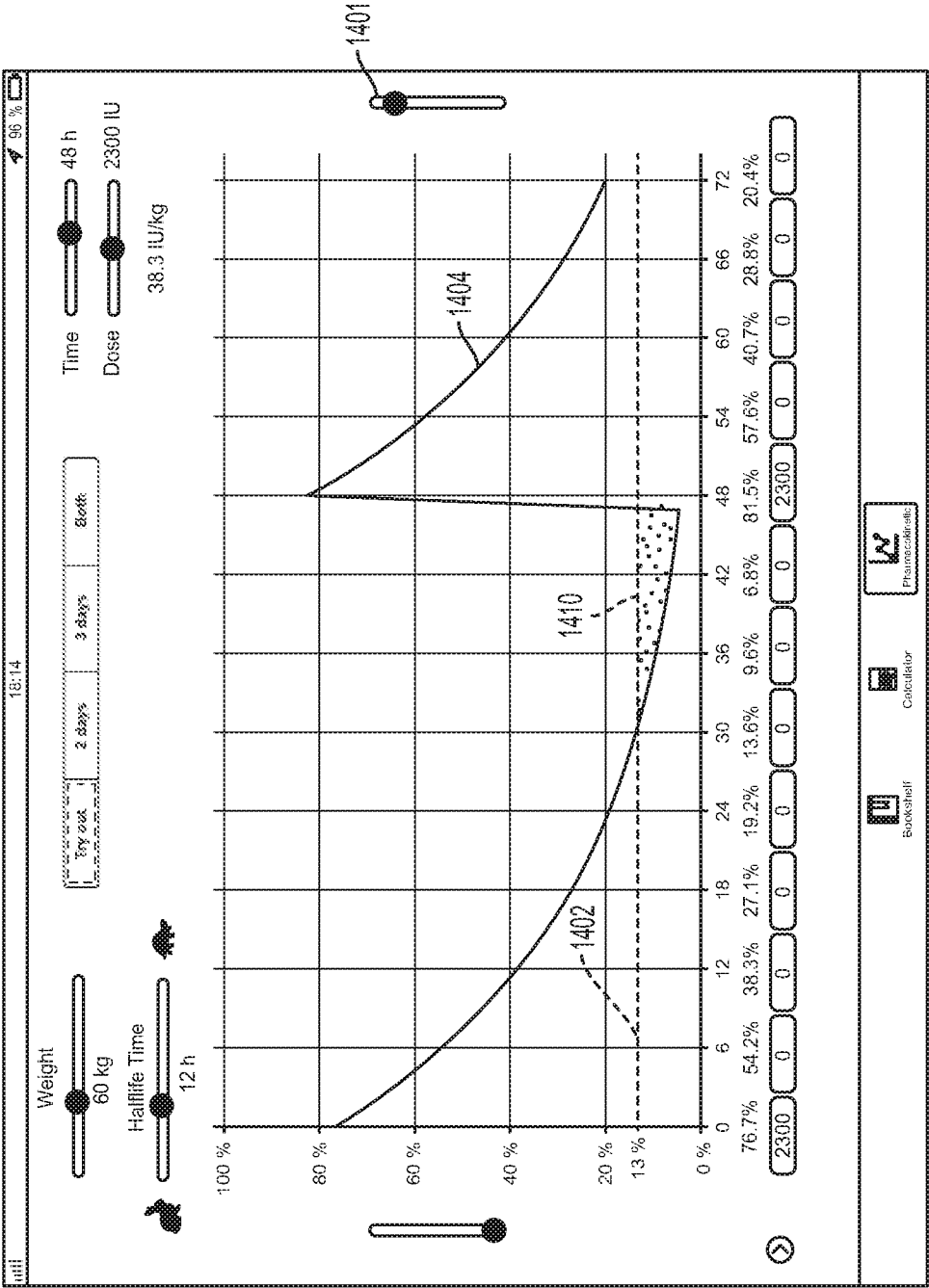


FIG. 17

1800

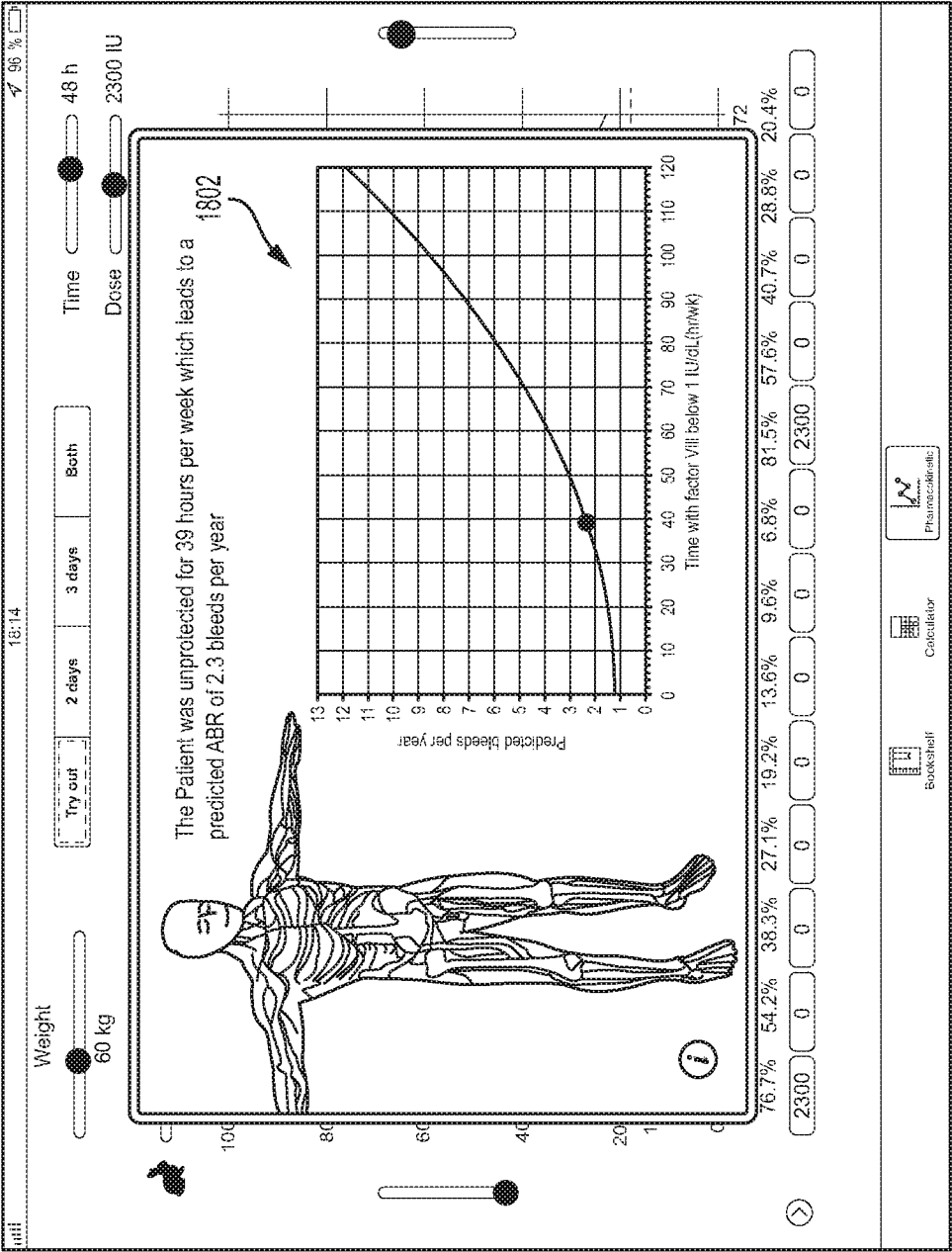


FIG. 18

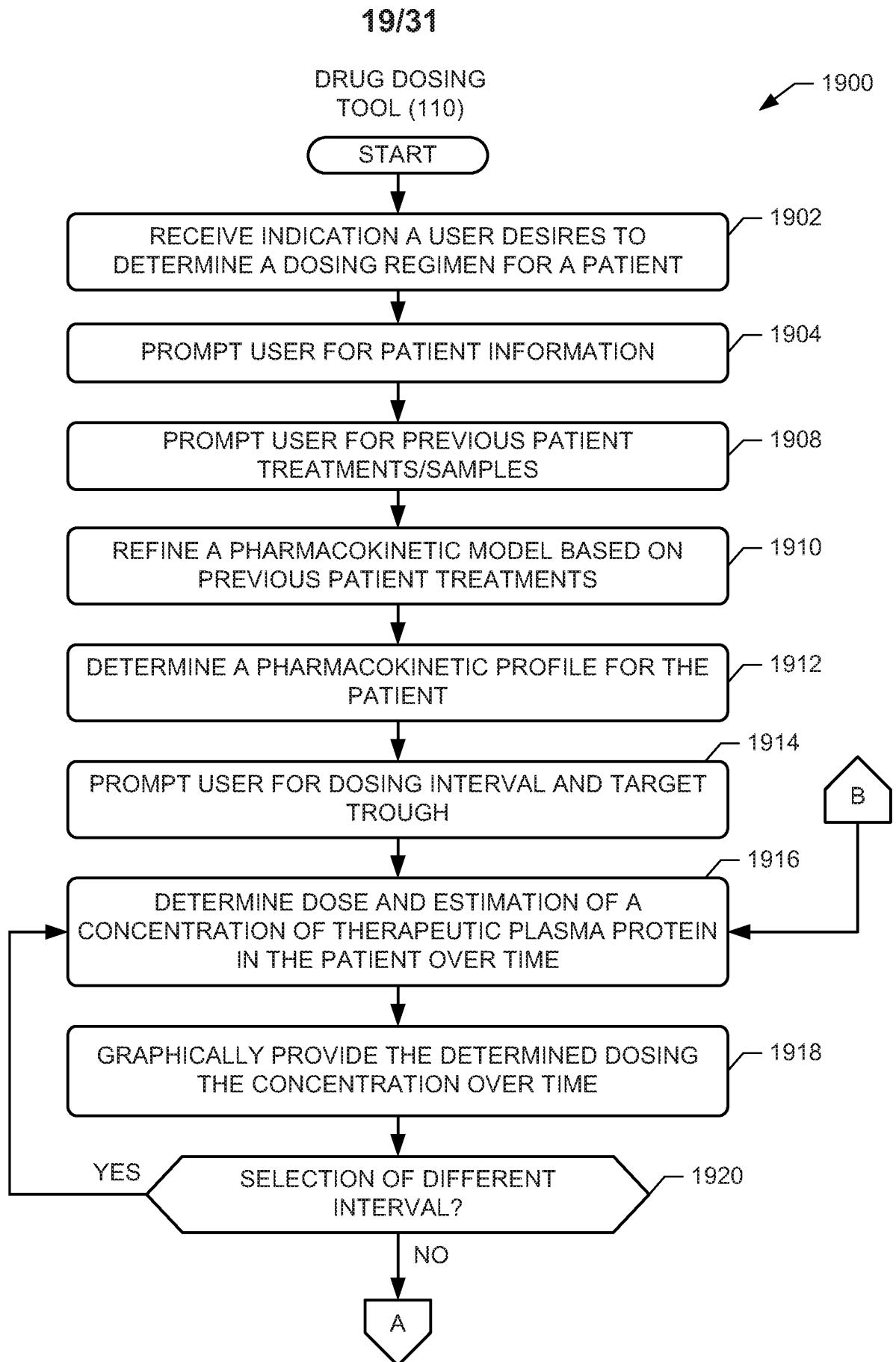


FIG. 19

20/31

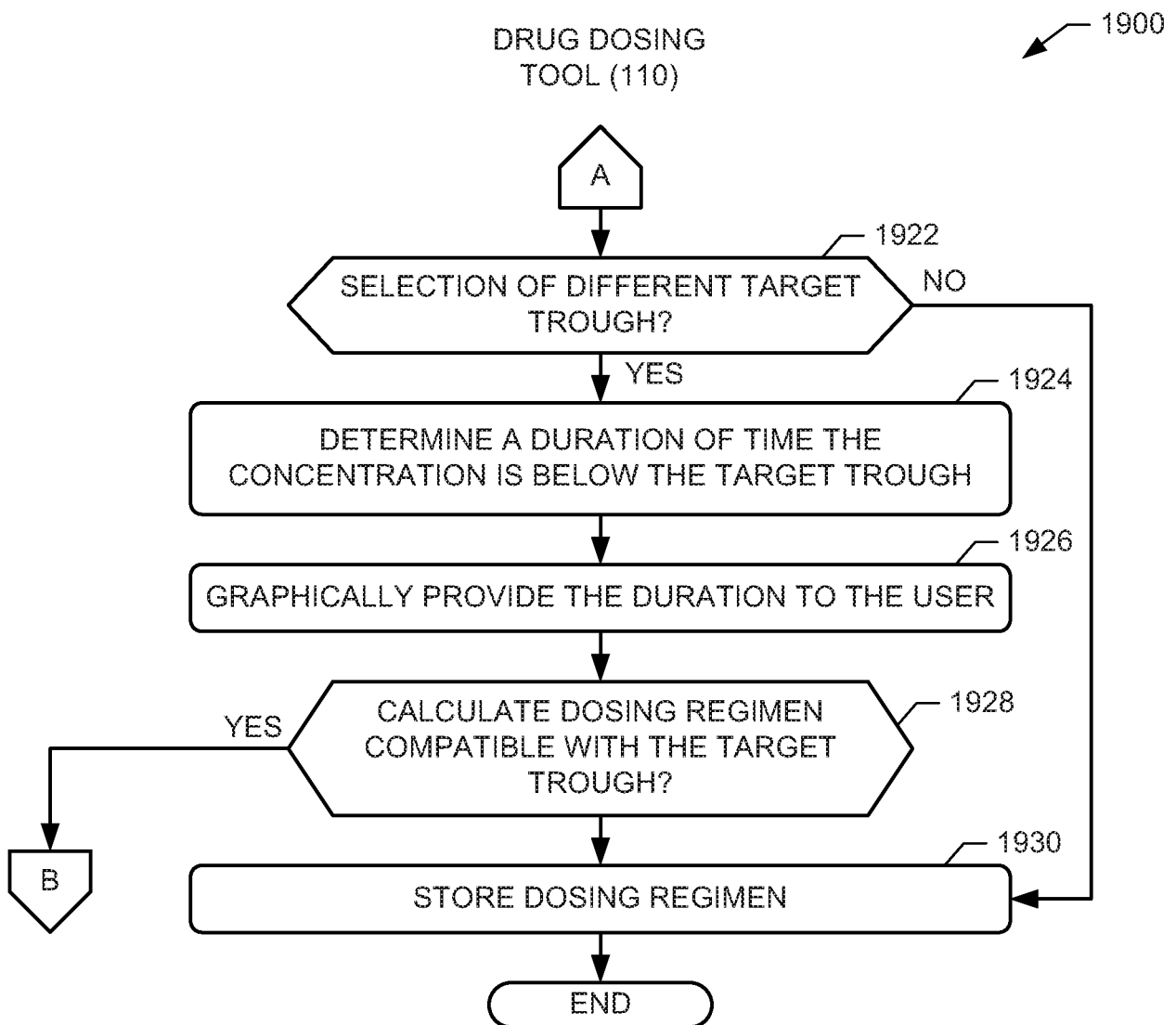
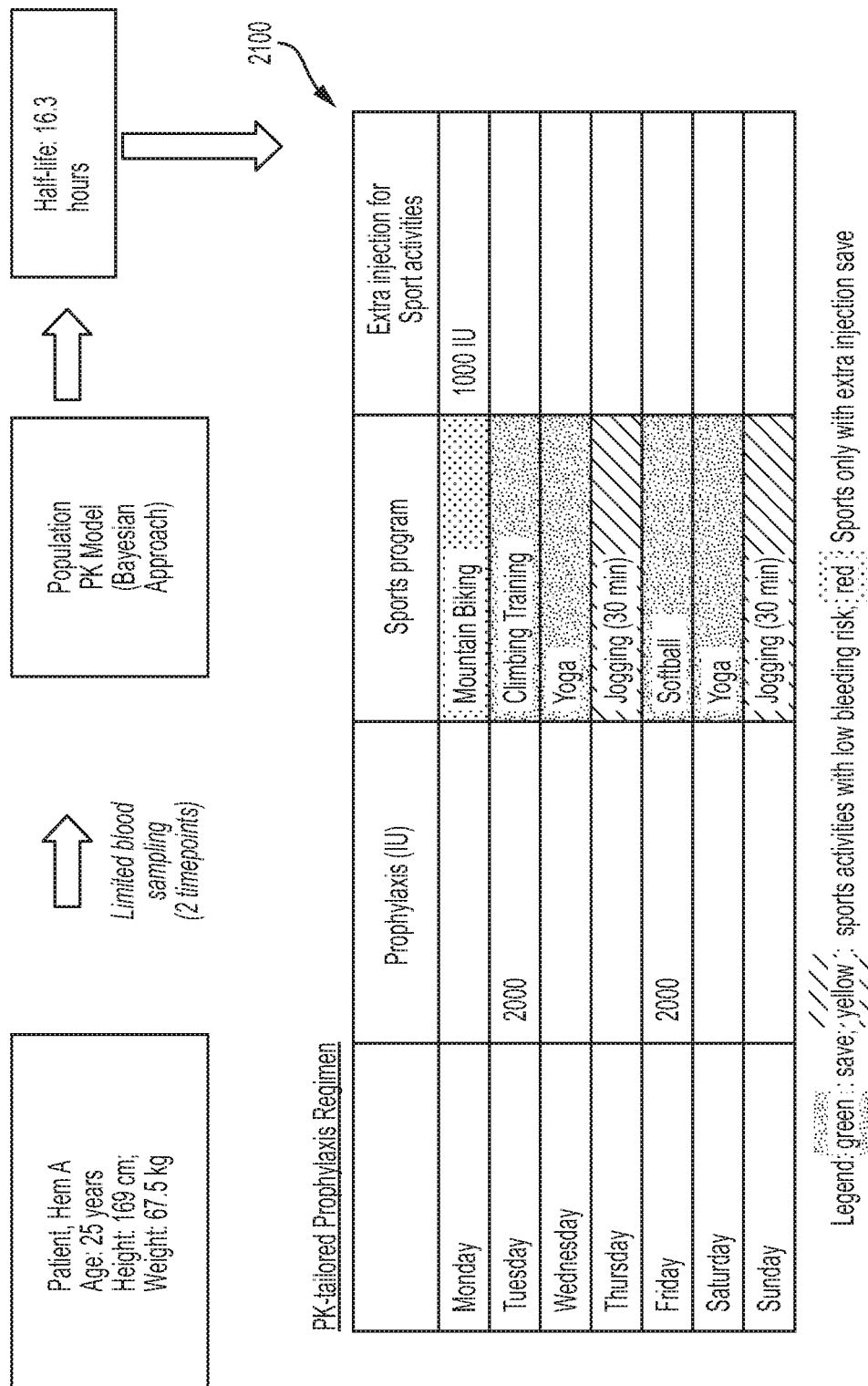


FIG. 20



25

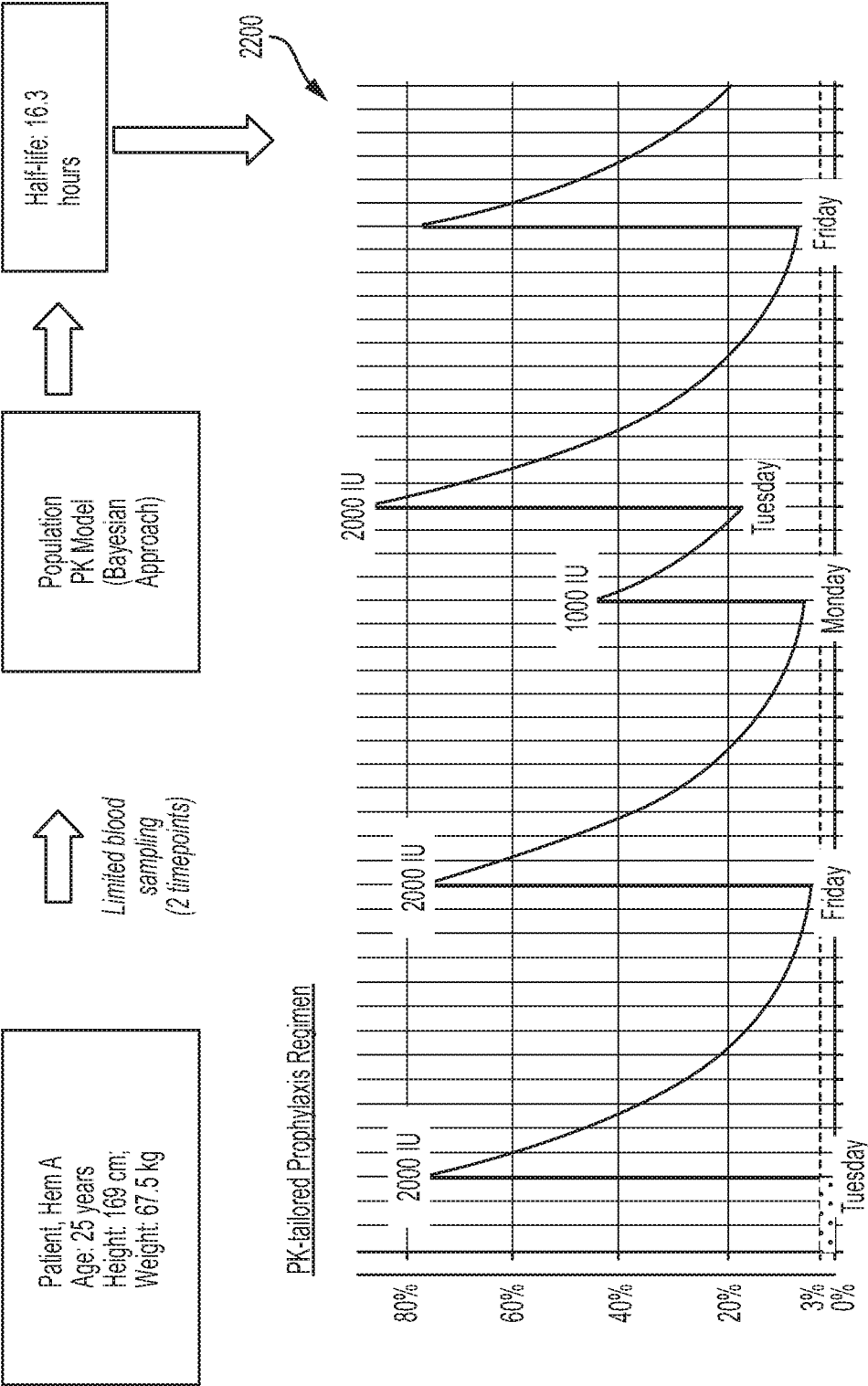


FIG. 22

23/31

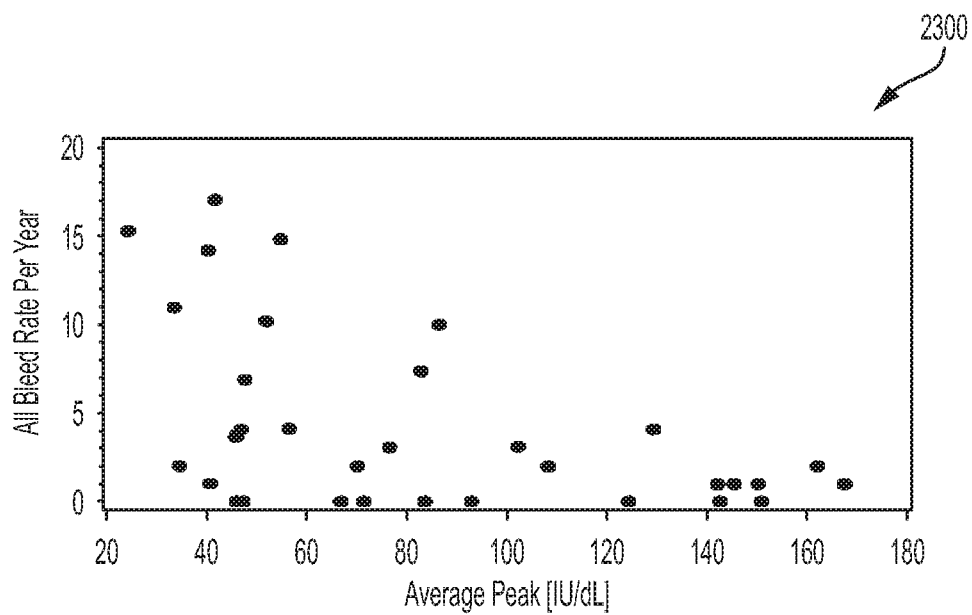


FIG. 23

2400

	All Bleeds		Joint Bleeds	
	Coeff.	P-value	Coeff.	P-value
Average Peak [IU/dL]*	-0.0174	0.0004	-0.0164	0.0013

FIG. 24

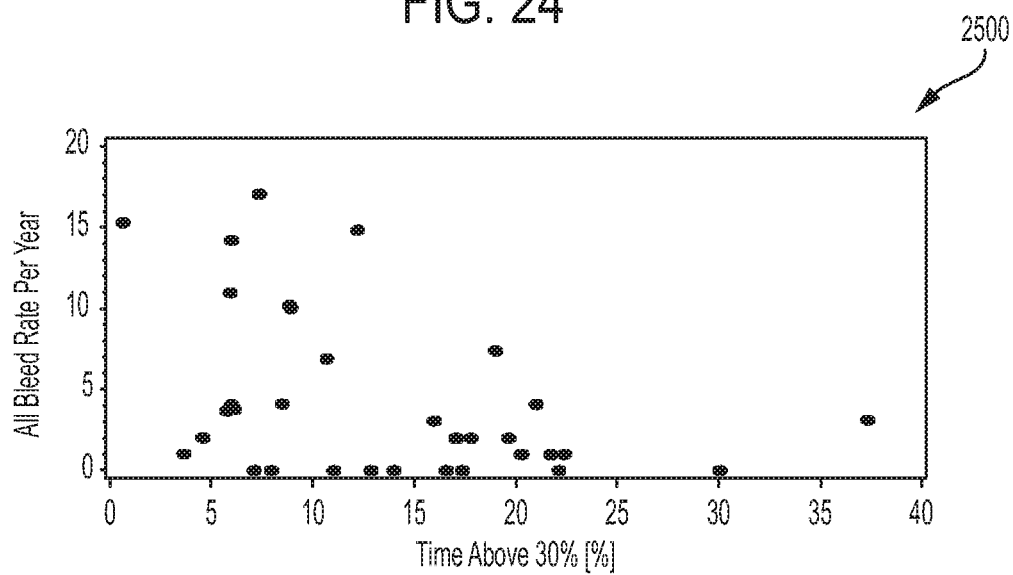


FIG. 25

24/31

2600

	All Bleeding		Joint Bleeding	
	Coeff.	P-value	Coeff.	P-value
Time Spent > 10%*		NS		NS
Time Spent > 20%		NS	-0.0578	S
Time Spent > 30%	-0.0786	S	-0.0832	S
Time Spent > 40%	-0.0911	S	-0.0951	S

FIG. 26

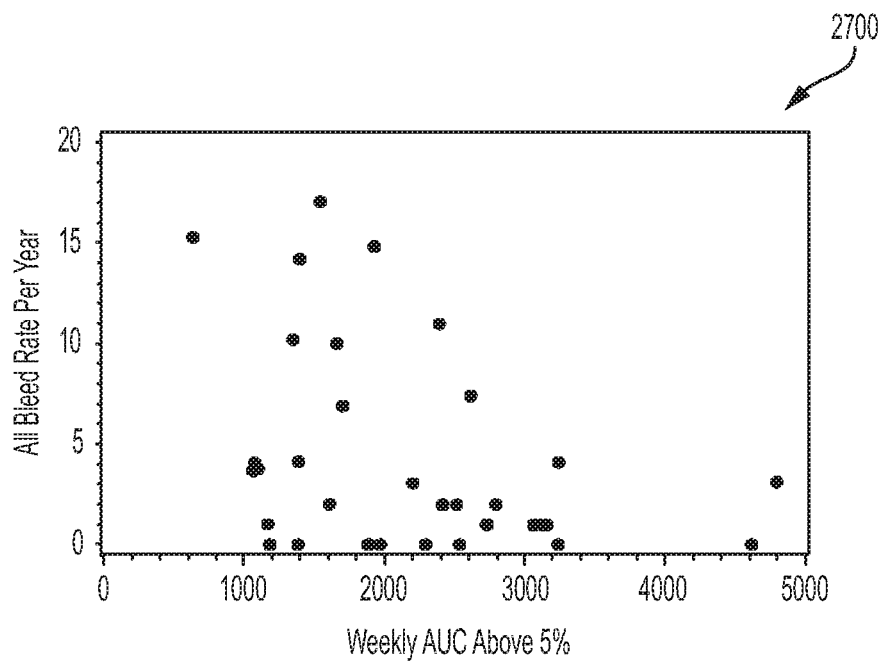


FIG. 27

2800

	All Bleeding		Joint Bleeding	
	Coeff.	P-value	Coeff.	P-value
$\text{Log}_{10}(\text{Weekly AUC} > 5\%)$	-3.1307	S	-3.0986	S
$\text{Log}_{10}(\text{Weekly AUC} > 10\%)$	-2.6189	S	-2.5856	S
$\text{Log}_{10}(\text{Weekly AUC} > 20\%)$	-1.9820	S	-1.9474	S

FIG. 28

2900

Variable	Time>05% Time>10% Time>20%	AUC>05% AUC>10% AUC>20%
Average Cmax	0.15 (ns) 0.43 (ns) 0.65 (s)	0.82 (s) 0.82 (s) 0.87 (s)
Time>05% Time>10% Time>20%		0.60 (s) 0.81 (s) 0.90 (s)

FIG. 29

3000

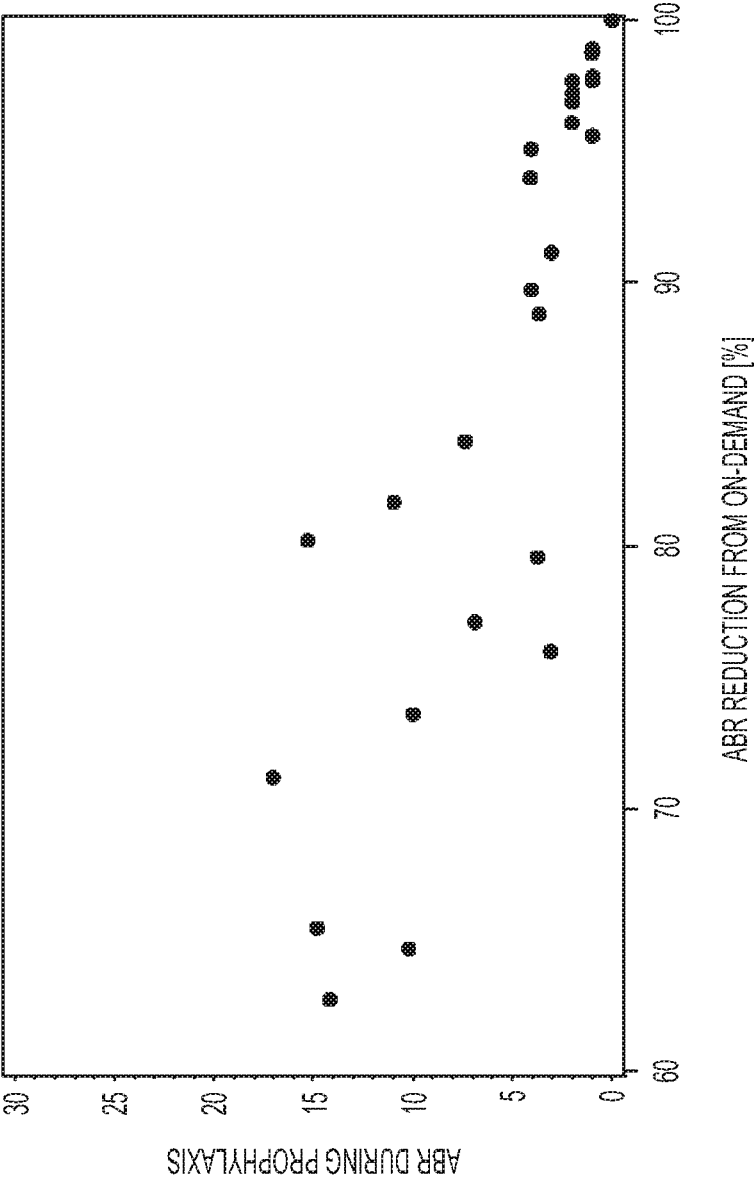


FIG. 30

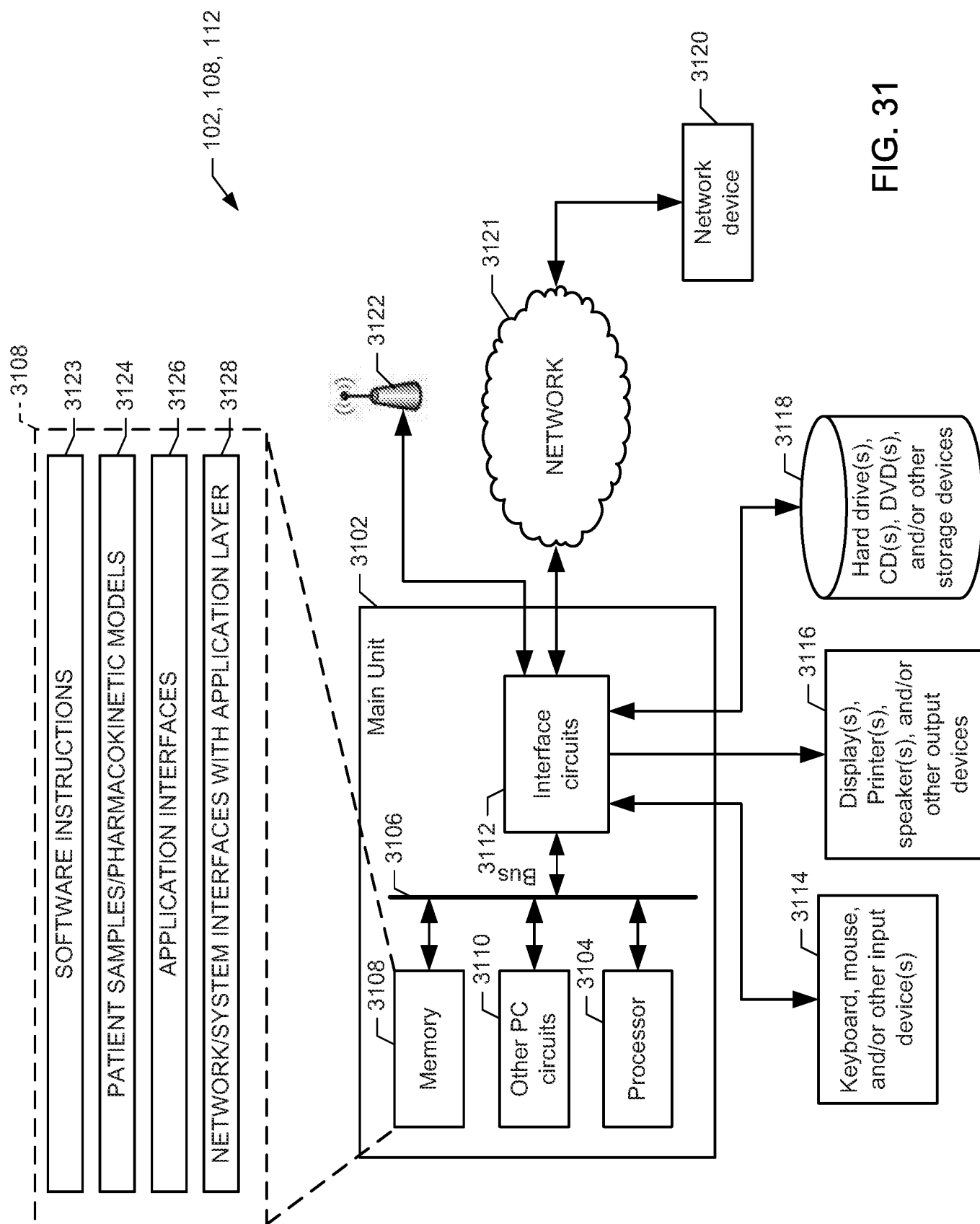
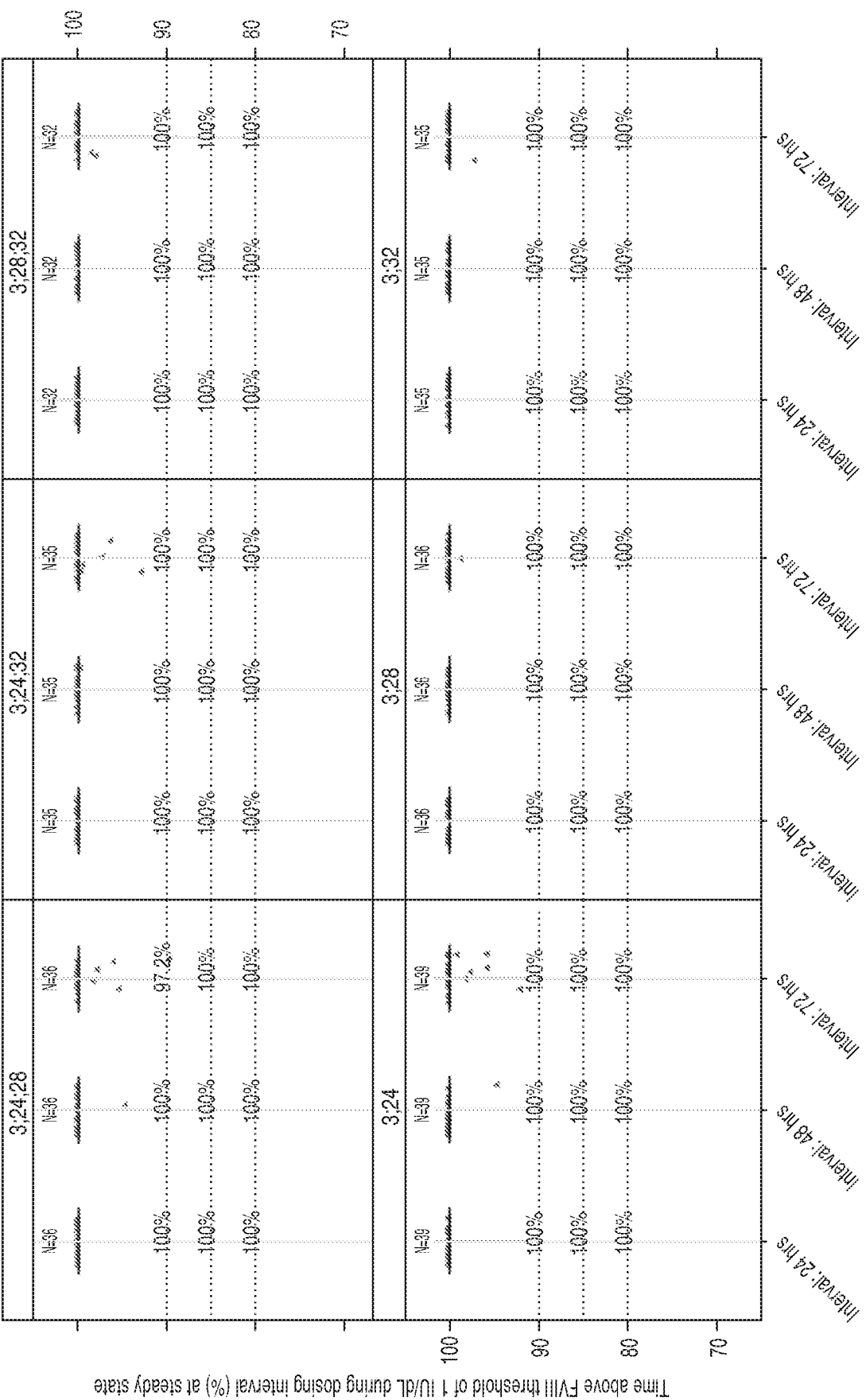


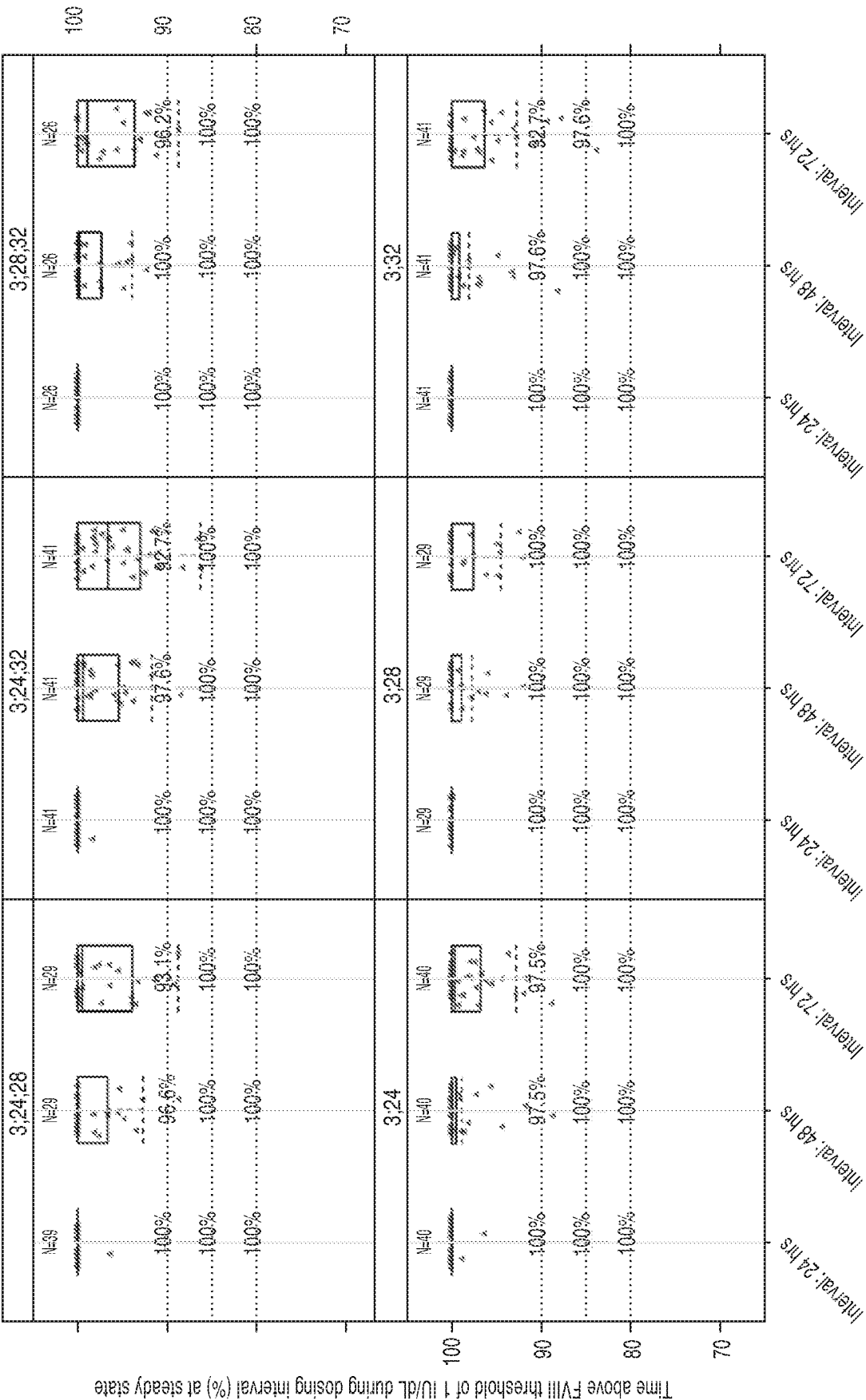
FIG. 31



Basis: Results from Bayesian algorithm on all available quantifiable samples

FIG. 32

Results based on sparse sampling profiles which passed the reliability check



Basis: Results from Bayesian algorithm on all available quantifiable samples

Results based on sparse sampling profiles which passed the reliability check

FIG. 33

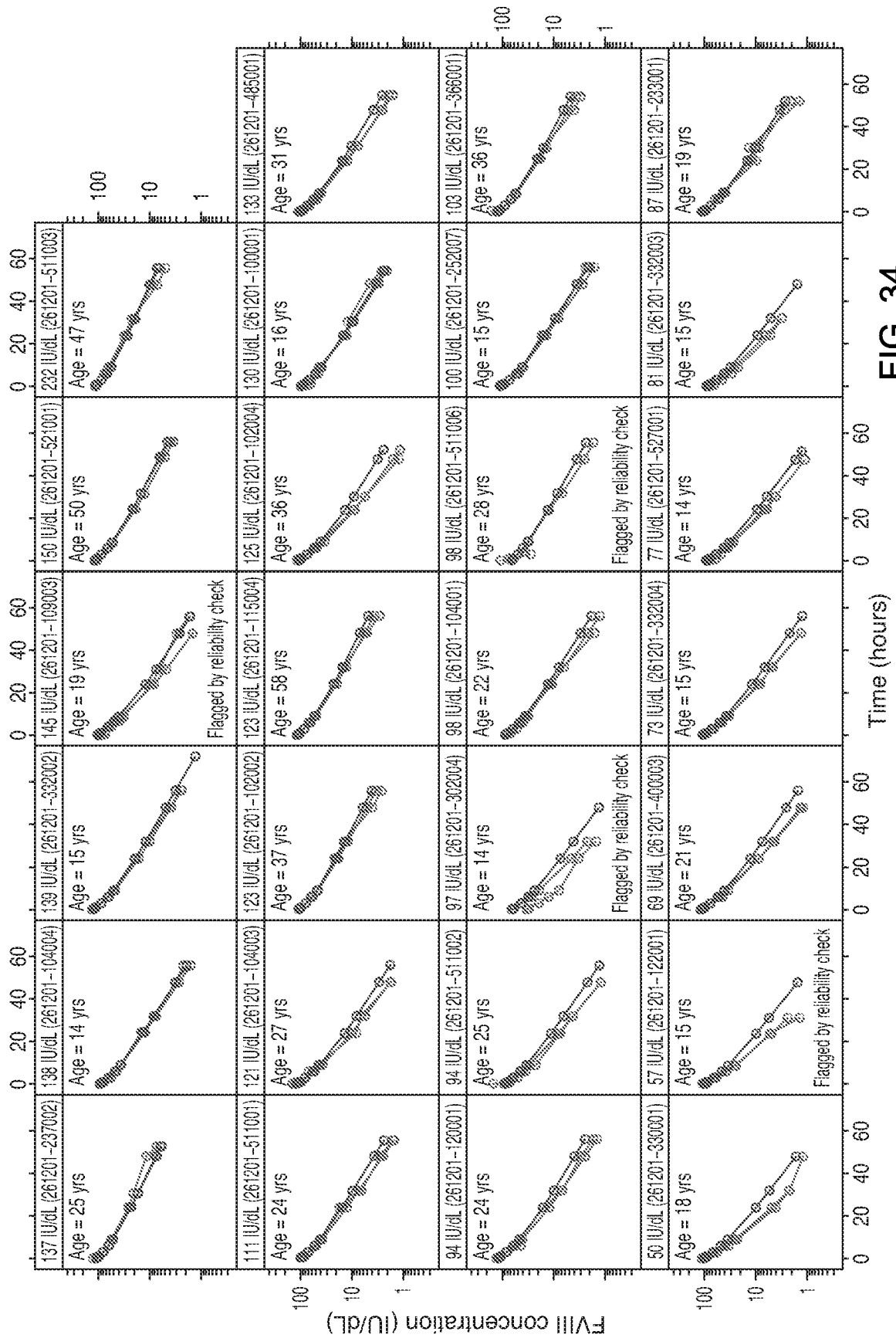
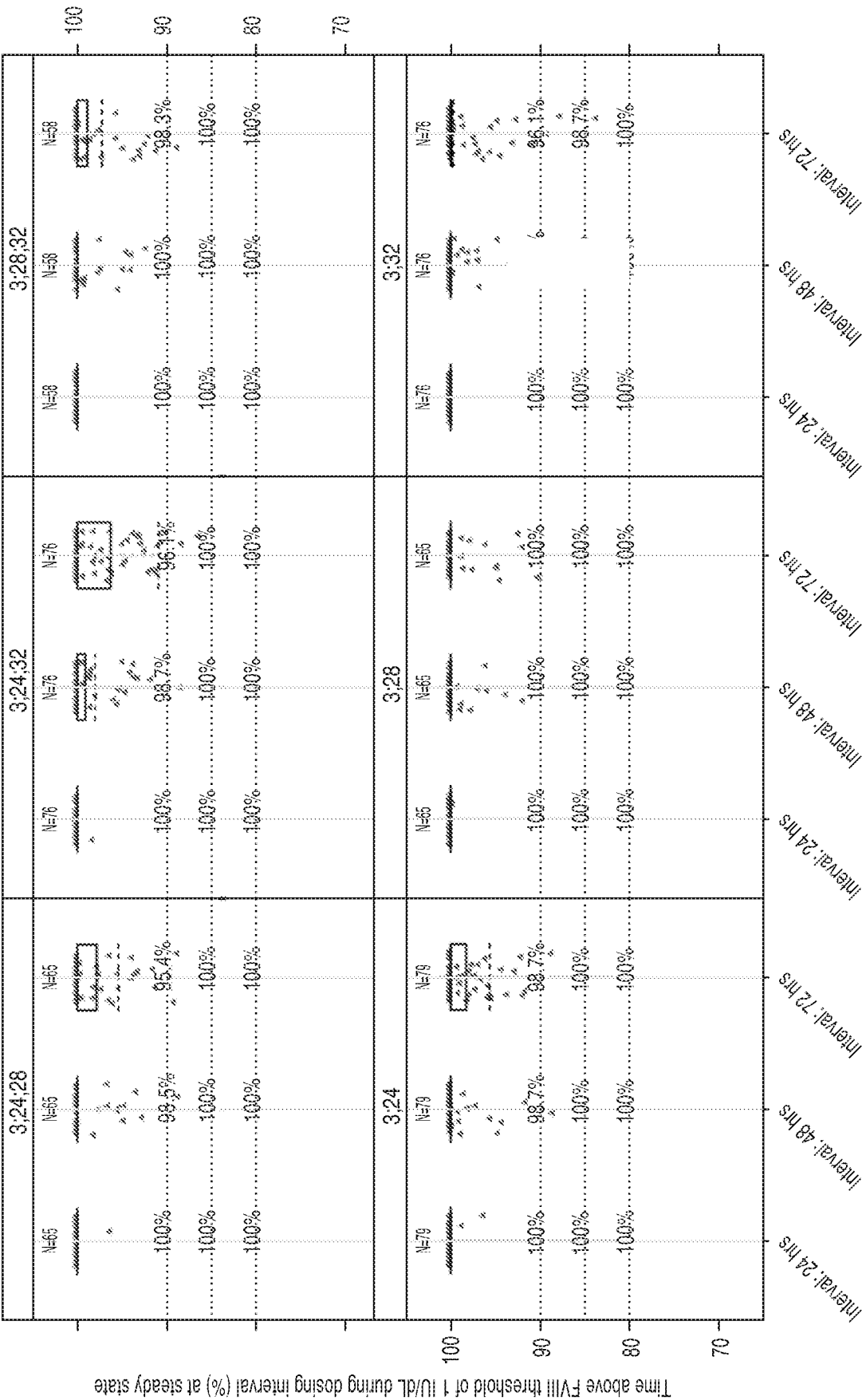


FIG. 34



Basis: Results from Bayesian algorithm on all available quantifiable samples

Results based on sparse sampling profiles which passed the reliability check

FIG. 35