

**Supplemental Table 1**

**Individual Subject Population Sequencing at Day 1, Day 4, and Day 14 of the NS5A Coding Region**

**1 mg GS-5885**

ID	GT	D1 Change From Reference 1a H77	D4 Change from D1	D14 Change from D1
AE	1a	T64A I74V R78K K107T R123Q S131T T135A I144V E171D E181D M226V A245T	No Change From D1	No Change From D1
AF	1a	I52I/V K107K/R R123Q S131T I144V M226V N246N/T	T87T/S N/T246T	T87T/S
AG	1a	T64T/A R81K K107T R123Q S131T T135A I144V L158L/I E181D M226V A245I/T E264A I280V D292G V296I S306A/T R308K A310T R311P V326L	L31L/M H58H/P I/T245T R348Q L368V	Q30Q/R L31L/M I/T245T A/T306S K308R T310A P311R L326V
AH	1a	G2D A25T R44K V46A R81K R123Q S131T K139R I144V L158I M226V V296I A310T R311P V315I	I296V T310A P311R I315V	V326I R348Q L368V
AI	1a	A25T S85N K107A S131T I144V E171D/E S174T E181D G215K M226E A245T	No Change From D1	No Change From D1
AJ	1a	R78K R81K/R I121V R123Q S131T I144V R176I/M/R E181A M226V L239F	Unable To Genotype	R262K T270P/T E273D/E E275E/K I280I/V D285D/N E290E/K V296I E301E/K R308K/R A310A/S R311P V326I/M E327E/K K331E/K E335*/E R348K L368V N392D T395A/E V410A
AK	1a	R44K S85N K107E/K I121V R123Q S131T I144V M226L K240K/R	L226M K/R240K	Q30Q/R
AL	1a	H58P E62D K68R V75T R78K R123Q S131T I144V L153V F161Y E171D M226E P237I K240R E256D	V296I A310T R311P V326L T328S K357R E365D T370S E374Q T377A	V296I A310T R311P V326L T328S K357R E365D T370S E374Q T377A
AM	1a	M28V T71S T79S K107E I121V S131T I144V R176S E181A A197T L199V M226L N246D	Q30Q/R I280I/V V296I S306C/S R311Q V315I/V V326L T328P K331R D333G L368V	Q30H/Q/R
AN	1a	S1A I34I/V R44K I52I/V R81K/R K107E S131T I144V E171D E181D A213T M226V A245A/I/T/V	A25A/T E107E/K	No Change From D1

**3 mg GS-5885**

ID	GT	D1 Change From Reference 1a H77	D4 Change from D1	D14 Change from D1
A	1a	K107K/T R123Q S131T I144V M226V V296I R311Q V315I R348Q L368V I388V N392D T394A T395A C404H V410A G439E D441G	K24E/K Q107T I296V Q311R I315V Q348R V368L V388I D392N A394T A395T H404C A410V E439G G441D	M28M/T Q30Q/R L31L/M V388I D392N A394T A395T H404C A410V E439G G441D
B	1a	I12I/V R44K K107T R123Q S131T N137D/N I144V H180N E181A/T P183L/P A245V N246S I280I/V V296M R304L R308K R311P V315I V326I P346P/S R348Q S349P/S T367F/V L368I S369F/S P401S V410A G439E D441G	M28M/V Q30Q/R V37V/L Y93Y/C N137N/D P183P/L	I/V12I M28M/V V37L/V Y93C I121I/V D/N137N V181T S183L I/V280I M296V L304R K308R P311R I315V I326V P/S346P Q348R P/S349S F/V367T I368L F/S369S S401P A410V E439G G441D
C	1a	R44K R78K T99S K107T S131T T135A I144V R176S M226V V296I T328P L368V V410A G439E D441G	W4*/W M28M/T Q30Q/R	Q30Q/R I296V P328T V368L A410V E439G G441D
D	1a	R78K I121I/V R123Q S131T I144V E181D M226V	Q30Q/R I280V V296I A310T R311P/Q V326L T328K K331E P343P/S P347P/Q R348K/Q L368V G390S T394A/T V410A G439E/G A440A/D D441D/G	Q30H/Q/R L31L/M
E	1a	R78K/R K107T R123Q S131T I144V A213T G215K M226E A245A/T I280V V296I A310A/T R311P V315I/V V326L S349T L368V G390S T395A P401S V410A G439E A440T D441G	Q30Q/R I/V315V	Q30Q/R I/V315V

F	1a	R123Q F127S S131T N137D I144V S174N E181A M226L K240R V296I P299C A310T R311Q S349P V361I L368V	Q30Q/R E398G V410A T442A	M28M/T E398G V410A T442A
G	1a	R123Q S131T I144V M226L/V A245T D251D/E E256E/Q T270L V296I/V R308K R311Q/R V326M/V R348Q K357K/R E365D/E T367N E374D/E T389A G390A N392D/N T393S S396P A400A/S C404C/R P405S V410A D430E G439E D441G	L31L/M V37M/V S38F/S E/Q226L I/V251E I/V296V Q/R311R M/V326V K/R357R D/E365E A373A/V D/E374D D/N392N P400S	M28M/T Q30Q/R T204N/T L/V226L/M/V Q/R311R M/V326V Q348R K/R357K D/E365E N367T D/E374E A389T A390G D/N392N S393T P396S A/S400A C/R404C S405P A410V E430D E439G G441D
H	1a	R78K R123Q S131T I144V E171D M226V N246K E256Q N276D V296I A310T R311Q V326I R348K L368I/L S369P N392D T395A P405H V410A	K24K/N Q30H/Q/R D368I	Q30Q/R L31L/M D368I
I	1a	L16V C98C/S K107T R123Q/R S131T I144V E181D A213T G215K M226V I280V V296I R304L A310T R311P V315I/V V326L R348Q E365D L368V G390S T395A S397P A400G P401P/S V410D G439E A440T D441G	M28M/T I/V315V	M28M/T L31L/M I/V315V
J	1a	G2D/G H54H/Y R78K S85N/S I121V R123Q S131T T135A I144V E181D G215A M226L/M A245T I280V K305R R308K A310G R311P V315I V326L K331K/R S349P/S T367S L368V	D/G2G Q30H/Q R81K/R A92A/V R348Q/R G390S D391D/G T394G T395A/T G403V V410A	Q30Q/R L31L/M P/S349S

#### 10 mg GS-5885

ID	GT	D1 Change From Reference 1a H77	D4 Change from D1	D14 Change from D1
K	1a	R78K K107T S131T I144V M226E K240R A245T I280V V296I R311Q V326I R348K/Q L368V	R6G/R L31L/M Y93C/Y F127F/S E171D/E T328M/T	Q30Q/R L31L/M R123Q/R I296V Q311R I326V K/Q348R V368L
L	1a	R44K R78K S85T K107T E117D R123Q S131T I144V M226V G267S V296I R311Q V326I	M28M/T Q30Q/R L31L/M P343A R348Q L368V	M28M/T Q30Q/R L31L/M I296V Q311R I326V
M	1a	R44K K107T R123Q S131T I144V M226L V296I	L31L/M I296V	L31L/M I296V
N	1a	V37M S85A/G/S/T K107N R123Q S131T I144V M226V	Q30Q/R V296I R308K/R R311Q V315I/V V326I R348Q T360M S369P T377A/T T389S V410A L428Y G439E/G T442A/T	Q30Q/R N85A V296I
O	1a	V46A T64A K107T R123Q S131T T135A I144V L158I E181D M226L I280V V296I R311P V315I V326L R348Q E365D L368V N392D/N P401S G403G/S D407D/N V410A G439E D441G	Q30H/Q/R V280I I296V P311R I315V L326V Q348R D365E V368L D/N392N S401P G/S403G D/N407D A410V E439G G441D	Q30H/Q/R V280I I296V P311R I315V L326V Q348R D365E V368L D/N392N S401P G/S403G D/N407D A410V E439G G441D
P	1a	R44K/R S85N K107T R123Q S131T I144V A213S M226E E293E/G V296I A310T R311P V315I V326L R348Q L368V	Q30Q/R K/R44R Y93C/Y	Q30Q/R K/R44R V368L
Q	1a	V37M/V T64S/T R78K/R R81K/R N105D/N R123Q S131T I144V F161Y M226E	M/V37V S/T64T M/V78K K/R81R S85N Y93C D/N105N V296I R308K R311P V326L	M/V37V S/T64T M/V78K K/R81R S85N Y93C D/N105N
R	1a	S1C R44K/R R78K K107K/T F127S S131V T135A I144V E171D S174T R176S E181D A213T M226E S228G G267S S306F R308K R311P V315I V326L K331E/G/K/R R348Q L368V N392D G403V Y413C	Unable To Genotype	No Data
S	1a	G2D R44K K68R R81W K107T R123Q S131T I144V S146A/S E171D	Unable To Genotype	Q30Q/R S249F I255V R262W A289T V296I

T	1a	G2D R44K R78K K107T I121I/V R123Q S131T I144V M226E/V A245T A252F/I/V E253D/E E256K G267G/S E291D V296I R311Q V326L R348Q L368V	Q30Q/R H58D/H I/V121I D/E252F/V G/S256E/K N276N/S K331K/N K357K/R P401S D407E V410A D441S	Q30Q/R L31L/M F/I/V252A D/E253E K256E G/S267G D291E I296V Q311R L326V Q348R V368L
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**10 mg GS-5885**

ID	GT	D1 Change From Reference 1b Con 1	D4 Change from D1	D14 Change from D1
AO	1b	T17S/T F37L Q54H R78K T122R F127Y T135A L183V T245A R246N E285D Q288R V296I R305K K308R R311S A347T S364T T377V K378Q	Unable To Genotype	Y93H
AP	1b	F37L K44R Q54N P58P/S R78K V130I V138I C165S L176Q K215G I280V Q288R A347T S364T	P/S58P Y93H	H85H/Y Y93H
AQ	1b	T99S K166G L176Q R246H Q288R R305K	Unable to Genotype	L31L/V Y93H/Y T99S K166G L176Q R246H Q288R R305K S364T
AR	1b	F37L Q54N R78K V138L E171D L176Q L183V T213A E285D Q288R R305K R311S I315V L326I R359K	F36F/L Y93H/Y	L37F/L K78K/R Y93H L138I/L A213A/T S311A/S A347A/T K359K/R
AS	1b	T14E T17S/T L31M F37F/I K44R L176Q T213S/T R246H/R Q288R R305K A347T S364T E365D	E14E/V S/T17T F/I37F S81C/S Y93H S/T213S H/R246R R311P/R I315I/V A349A/T I352I/V	E14E/V S/T17T F/I37F S81C Y93H S/T213S H/R246R R311P I315V A349T I352V
AT	1b	R30Q Q62K P77S R78K T174V L176Q E285D Q288H R305K A347T I352V R359K S364T	Y93H	Y93H
AU	1b	K44R V138I/V A164A/V L176Q A197A/T Q288R R305K I315V A347T S364T	H85H/Y Y93H	H85H/Y Y93H
AV	1b	F37I Q54H R78K E116A V138I L176Q A197T A219D T245A R246H E285D Q288R R305K K308S R311L M313L I315V L326I V335T A347T V362I S364T	Unable to Genotype	K78K/R Y93H
AW	1b	F37V K44R R78K S101C E171D L176Q T213S I280V Q288R R305K I315V A347I A349T V362I S364T E365D	Unable to Genotype	Y93H V315I/V
AX	1b	T17S/T K20K/R Q54H T56I Q62E/Q T79A V130I/M/V E171D/E L176Q A197A/T E285D/E Q288R R305K R311P/R M313L/M I315I/V L326I/L P346S A347D/N R357K S364T	S/T17S K/R20R E/Q62E Y93H I/M/V130I D/E171D A/T197A D/E285D P/R311P L/M313L I/V315V I/L326I D/N347S	I56I/T Y93H D/E285D P/R311P I/L326I S346P/S D/N347N/T

30 mg GS-5885

ID	GT	D1 Change From Reference 1a H77	D4 Change from D1	D14 Change from D1
U	1a	R44K S85N T99S R123Q S131T I144V S174T V296I R311P	Q30Q/R L31L/M R348Q S369P	Q30Q/R L31L/M
V	1a	F36L V37M/V K107T R123Q S131T I144V A197T L199V M226L K240R V296I	Q30Q/R S306H R307P A310T V315I T328A R348K L368V	Q30Q/R
W	1a	S3C/S F36F/L V37I/M/V R44K H58H/P E62D/E K107S I121I/V R123Q S131T M133M/V I144V E152E/V L158I/L F161F/Y E171D/E S174S/T E181A/E/K/T	L31M F/L36L I/M/V37V H/P58P D/E62D I/V121V E/V152V I/L158I F/Y161Y D/E171D S/T174S A/E/K/T181E	C/S3C L31M F/L36L I/M/V37V H/P58P D/E62D I/V121V M/V133V E/V152V I/L158I F/Y161Y D/E171D S/T174S R176S A/E/K/T181T G215R M226E P237L A245I V296I A310T R311P V315I V326I R348Q L368V
X	1a	K24K/R R44K/R R48Q/R V75A/V K107M/T I121I/V R123Q/R S131T T135A/T I144V M226I/M/V A245T E253D	K/R24R L31L/M K/R44K Q/R48R M/T107T I/V121I Q/R123Q A/T135T I/M/V226M T245A D253E	Q30Q/R L31L/M M/T107T I/V121I I/M/V226M T245A D253E
Y	1a	R78K K107T R123Q S131T T135A I144V E171D E181D G215K M226L A245T	Q30Q/R L31L/M Y93C/Y	Q30Q/R Y93C/Y
Z	1a	R78K K107T R123Q S131T M133V I144V R176T M226L A245T E256Q I280V V296I	Unable To Genotype	Q30Q/R H58D/H I296V
AA	1a	R78K S85N S131T N137D I144V E181A M226V D248E	Q30E/G/Q/R L31L/M	L31M D137N V144I A181E V226M E248D
AB	1a	R78K I121V R123Q S131T K139R I144V L158I G215R M226V G267S	Y93H/Y V296I R311Q V326L W329*/W R348Q T377V	Q30Q/R Y93C/Y V296I R311Q V326L R348Q T377V
AC	1a	Q30E/Q R123Q S131T I144V E171D G215K M226V	E/Q30E N246D D248N	E/Q30E N246D D248N I280V V296I R308K A310T R311P V326L R348Q T367S L368V
AD	1a	Unable to Genotype	Unable to Genotype	Unable to Genotype

90 mg GS-5885

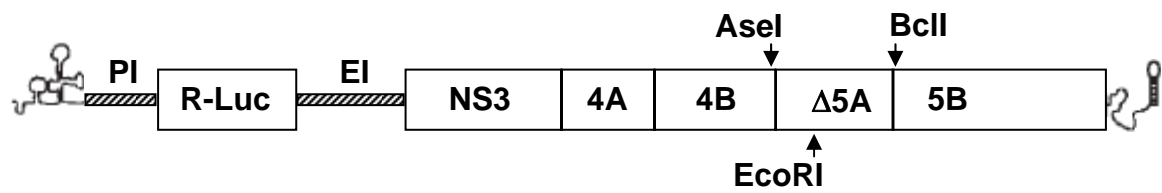
ID	GT	D1 Change From Reference 1a H77	D4 Change from D1	D14 Change from D1
BI	1a	V75A/V S85N/S K107T R123Q S131T I144V I209V G215R M226E A245T V296I/V R308K/R A310T R311Q V315I/V V326I/L R348Q L368V	K24K/R Q30Q/R T64A/T	Q30Q/R H58D/H T64A/T
BJ	1a	I12V V75A R123Q S131T I144V E181A A213A/T M226V V296I R311P	Q30Q/R Y93C/Y V326L T328S T364S L368V T370A	Q30Q/R L31L/M A181E A/T213A V226M I296V P311R
BK	1a	M83T C98S K107T R123Q S131T I144V E181D G215K V288A V296I R311Q V326I	Q30K/Q/R Y93C/H/R/Y V288A/V K331E R348Q L368V	Q30R K331E R348Q L368V
BL	1a	R44K R78K R81K K107T I121V R123Q S131T T135A/T I144V E171D L199V G215R M226E A245T E256A	Q30Q/R Y93C/Y A/T135A I280V V296I S306A R311P V315I V326L	Q30Q/R L31L/M T135A E256A
BM	1a	L31M R78K/R R81K R123Q S131T I144V A213A/T M226V A245A/T	No Change From D1	No Change From D1
BN	1a	R81K S85D R122K/R R123Q S131T I144V S174T A197T L199V G215R M226E A245T	Unable to sequence	Q30Q/R
BO	1a	R44K I52I/V T71S K107T S131T I144V L153L/V M226V I280V V296I S306Y R311P V315I/V V326L	L31M I/V52I L/V153L I/V315V L368V	L31M I/V52I L/V153L I/V315V L368V
BP	1a	R78K/R K107T R123Q S131T I144V G215K M226E A245T	L31L/M K/R78R Y93H/Y V296I R311Q V326I L368V	L31L/M
BQ	1a	V75A/V R81K/R K107T R123Q S131T T135A I144V M226V V296I R311T V326I/L/M R348Q K358K/R	Q30Q/R K358R	Q30Q/R S85N/S Y93C/Y K358R S369P/S
BR	1a	I52V T64A R78K R123Q S131T M133I T135A I144V L158I H180N G215A M226V A245T	Q30Q/R I280V V296I A310T R311P V326L D333G R348Q E365D L368V	Q30Q/R Y93C/Y

Placebo

ID	GT	D1 Change From Reference 1a H77* or 1b Con 1^	D4 Change from D1	D14 Change from D1
AY*	1a	K107T S131T N137D I144V E181A A245A/V	No Change From D1	No Change From D1
AZ*	1a	L31L/M R44K/R R78K S85N A103A/S I121I/V R123Q S131T I144V L153L/V E181D A245T	No Change From D1	A/S103A L/V153L T245A
BA*	1a	R44K A92P K107T R123Q S131T I144V H180N A213S M226L A245T	V296I R311Q V315I/V	V296I
BB*	1a	S1A R44K K107T R123Q S131T I144V E181D G215R A219E M226V A245V E264A I280V V296I R308K A310T R311Q V315I/V T367S L368V S384S/T G390S T395A S397P P401S S402P V410A G439E	Q30H/Q/R	Q30Q/R L31L/M I/V315V S367T V368L S/T384S S390G A395T P397S S401P P402S A410V E439G
BC*	1a	R44K G49E I52V K107T R123Q/R S131T I144V L158L/V F161Y A197T L199V G215G/R D251D/N	No Change From D1	D/N251D
BD*	1a	T64S/T R78K R123Q S131T I144V L158I M226V V296I	L31L/M	No Change From D1
BE*	1a	S85N T99N S114A R123Q S131T N137D I144V E181A A213T M226V A252V I255L D285E V296I	A289A/G E293E/K R308K/R	E285D I296V
BF*	1a	I8I/V R44K R78K R123Q S131T I144V E171D M226E/V D248E	D171D/E R311P V326L K331E/K	D171D/E A310A/T R311P V326L K331E/K D333D/G T360M/T L368V
BG^	1b	G2D T17S K44R T79A E171D L176Q R246H E285D Q288R R305K R311A I315V A349T S364T	No Change From D1	No Change From D1

BH^	1b	T17S L31M F37L Q54H R78K V138I E171D L176Q L179C E285D Q288R R305K S306T R311P L326I D331D/N A347T	No Change From D1	No Change From D1
BS*	1a	V37L R78K K107T I121I/V R123Q S131T I144V E181D M226V I280V V296I S306Q A310T R311Q V326L T328P K331Q E335K L368V	No Change From D1	No Change From D1
BT*	1a	R44K R78K K107T R123Q S131V T135A I144V E181D A213T M226V A245T	No Change From D1	E290G

Supplemental Figure 1

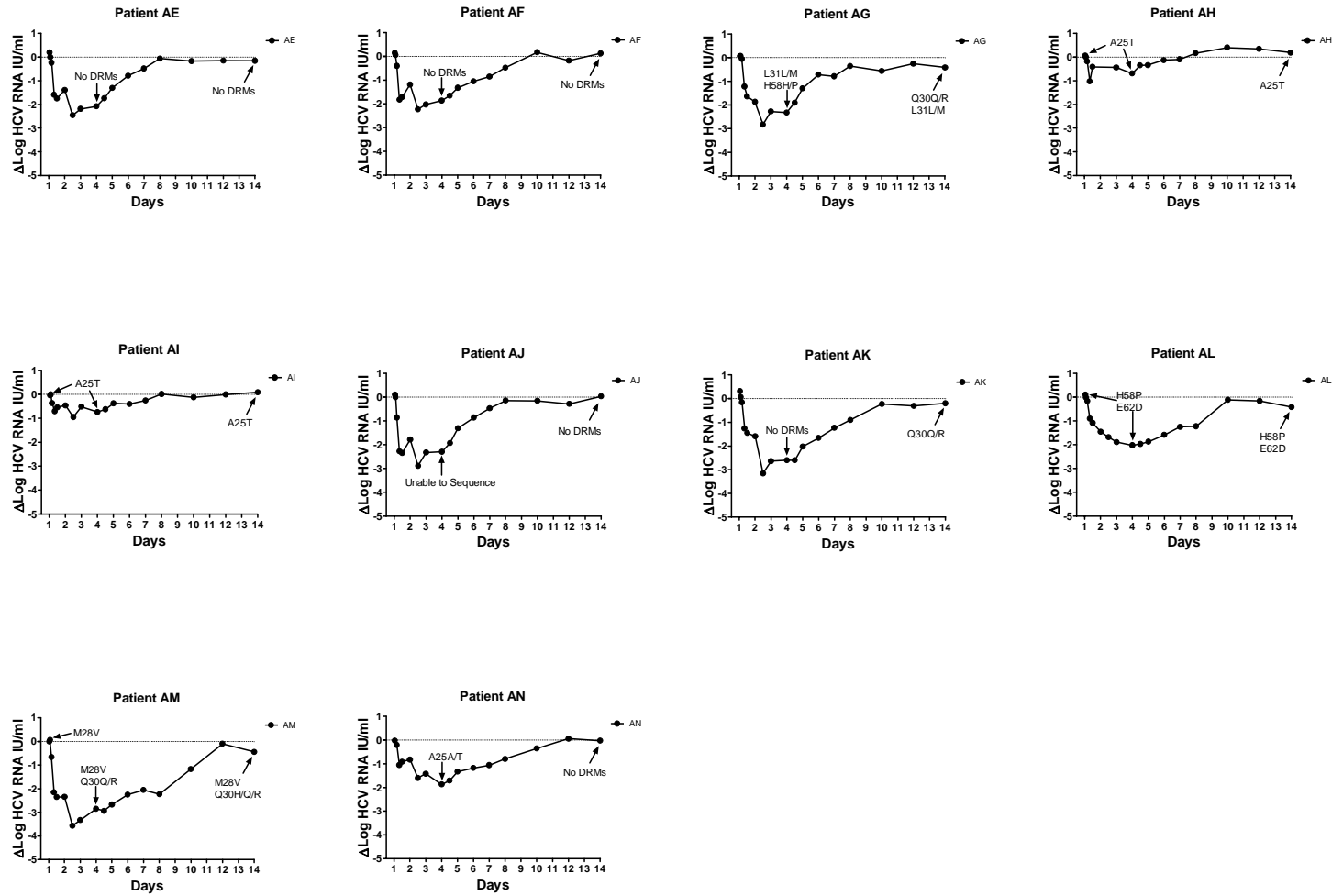


NS5A shuttle vector used for genotype 1a and genotype 1b NS5A insertion from clinical isolates. The backbone of the shuttle vector is from 1b: Con-1 AJ238799. The IRES elements driving Renilla Luciferase (R-Luc) and non-structural protein transcription respectively are the polio virus IRES and encephalomyocarditis virus IRES.

Lohmann V, Korner F, Koch J, Herian U, Theilmann L, Bartenschlager R. 1999. Replication of subgenomic hepatitis C virus RNAs in a hepatoma cell line. *Science* **285**:110-113.

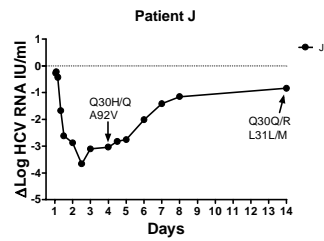
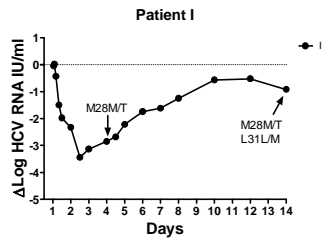
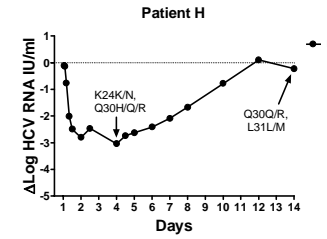
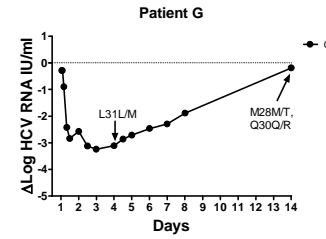
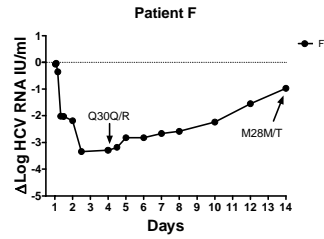
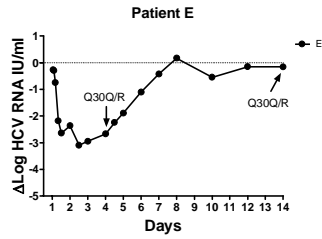
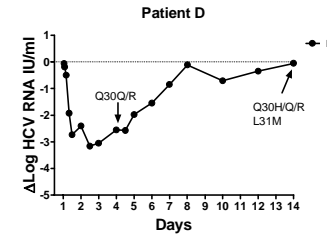
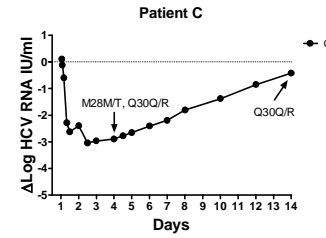
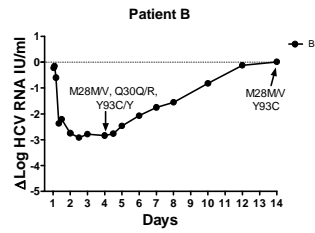
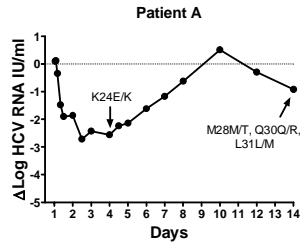
## Supplemental Figure 2

Genotype 1a Patients dosed with 1 mg GS-5885

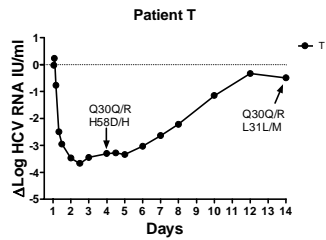
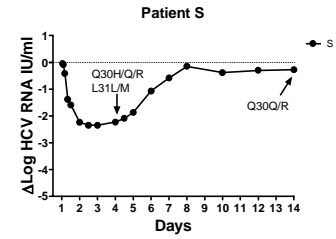
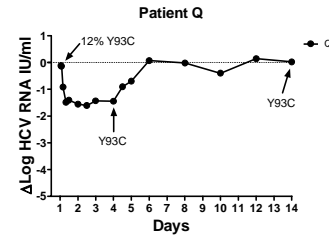
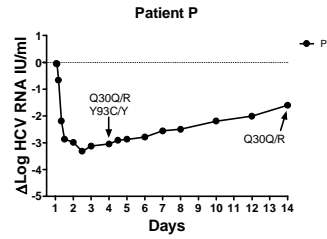
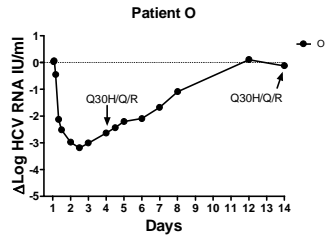
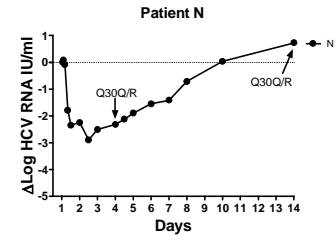
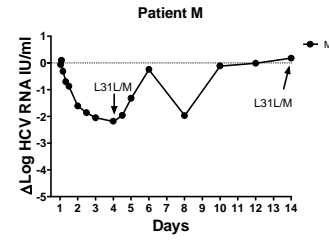
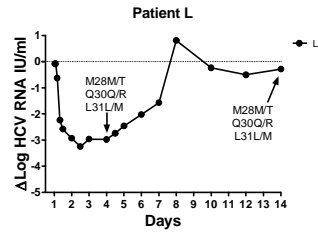
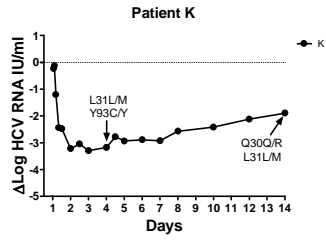




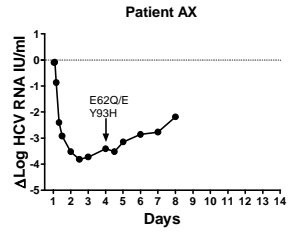
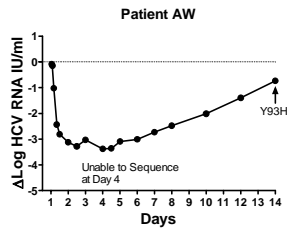
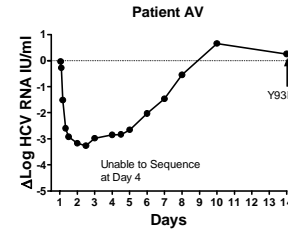
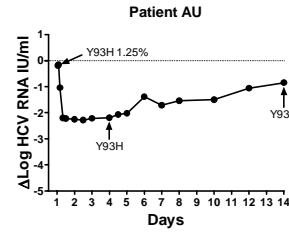
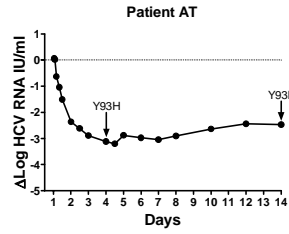
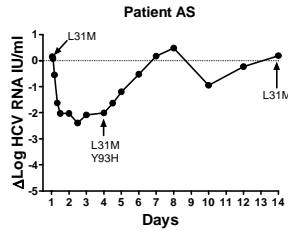
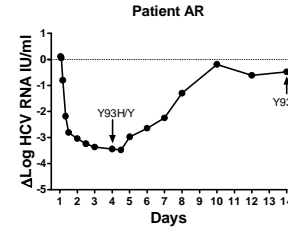
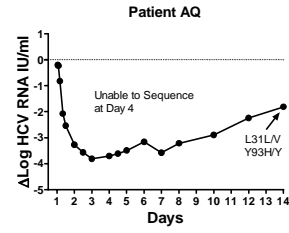
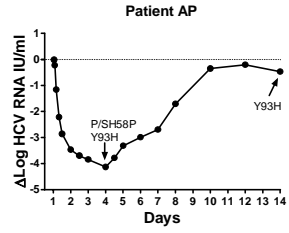
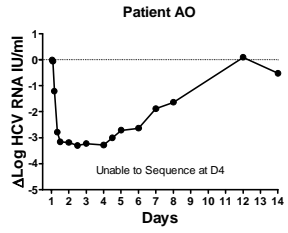
Genotype 1a Patients dosed with 3 mg GS-5885



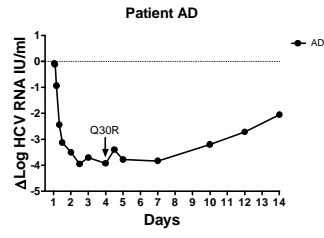
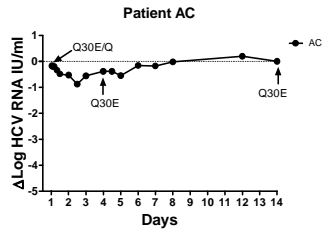
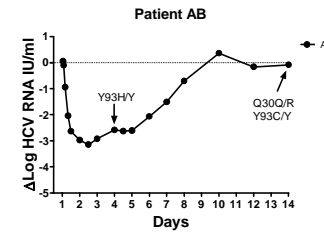
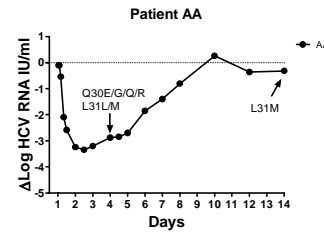
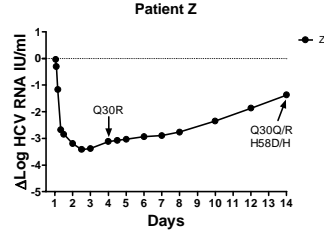
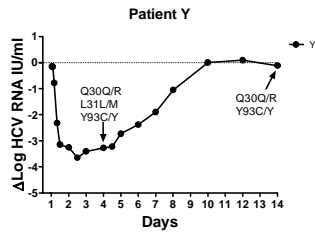
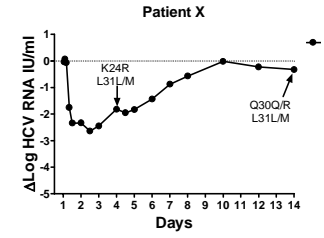
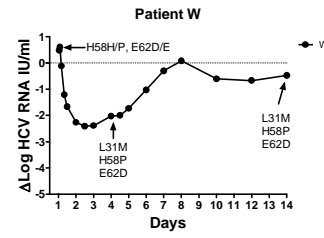
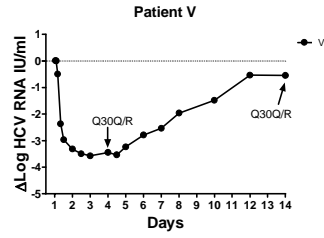
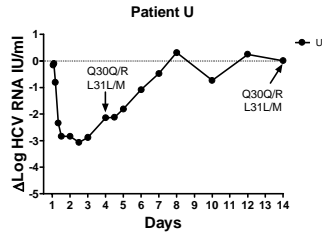
Genotype 1a Patients dosed with 10 mg GS-5885



Genotype 1b Patients dosed with 10 mg GS-5885



Genotype 1a Patients dosed with 30 mg GS-5885



Genotype 1a Patients dosed with 90 mg GS-5885

