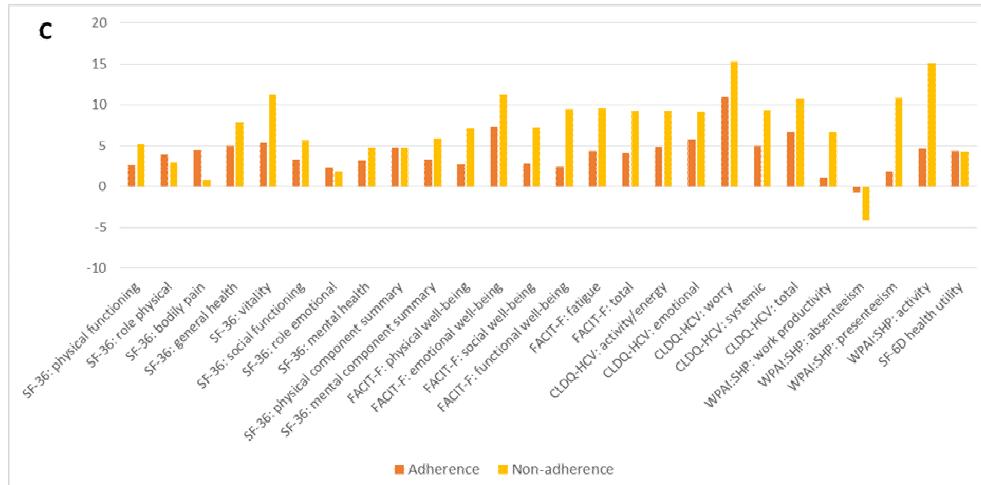
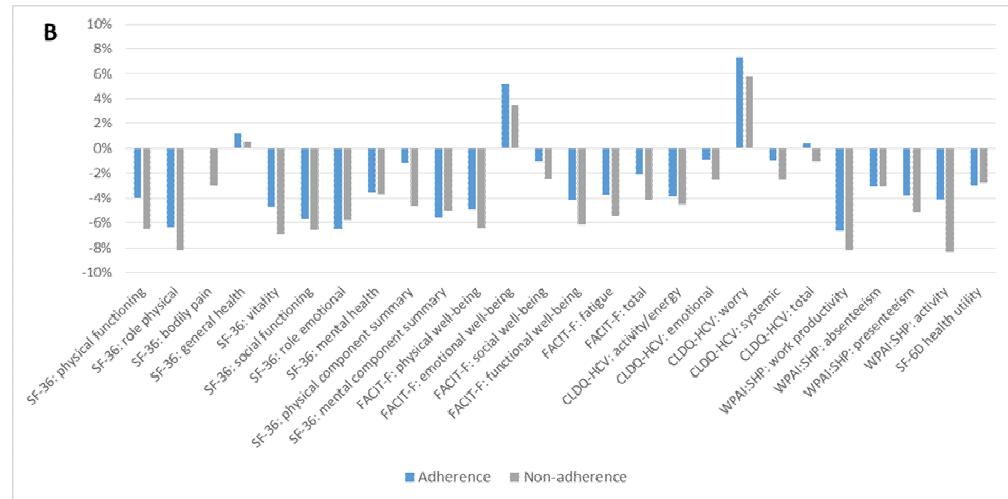
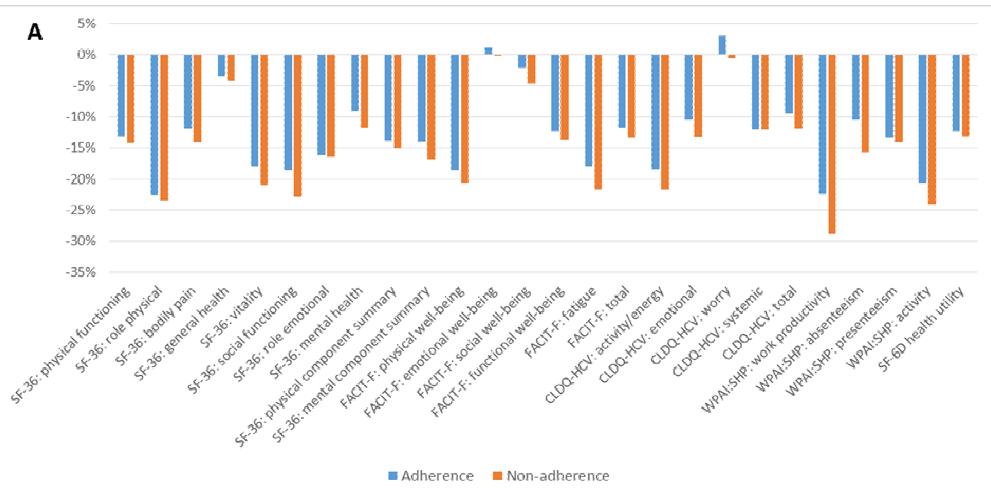


Supplementary Figure 1. End of treatment changes in PROs in patients adherent and not adherent to (A) IFN+RBV-containing regimens; (B) IFN-free RBV-containing regimens; (C) IFN-free RBV-free regimens



**Supplementary Table 1.** Comparison of adherent and non-adherent patients by treatment regimen.

	IFN-containing regimens			
	Adherence >= 80%	Adherence < 80%	p	All patients
N	510	147		657
Age, years	51.204 ± 10.775	53.558 ± 9.385	0.0068	51.731 ± 10.519
Male gender	345 (67.6%)	96 (65.3%)	0.59	441 (67.1%)
Caucasian	423 (82.9%)	114 (77.6%)	0.14	537 (81.7%)
African-American	54 (10.6%)	22 (15.0%)	0.14	76 (11.6%)
Asian	19 (3.7%)	7 (4.8%)	0.57	26 (4.0%)
Enrolled in the U.S.	397 (77.8%)	123 (83.7%)	0.13	520 (79.1%)
Employed	262 (65.5%)	63 (57.3%)	0.11	325 (63.7%)
Cirrhosis	85 (16.7%)	31 (21.4%)	0.19	116 (17.7%)
Treatment-naïve	345 (67.6%)	108 (73.5%)	0.18	453 (68.9%)
HCV genotype 1	333 (65.9%)	93 (63.7%)	0.62	426 (65.4%)
HCV genotype 2	26 (5.1%)	13 (8.9%)	0.09	39 (6.0%)
HCV genotype 3	119 (23.6%)	31 (21.2%)	0.56	150 (23.0%)
HCV genotypes 4, 5, 6	27 (5.3%)	9 (6.2%)	0.70	36 (5.5%)
coinfection with HIV	0	0		0
baseline HCV RNA > 6 log 10/mL	388 (76.1%)	107 (72.8%)	0.41	495 (75.3%)
baseline ALT > 1.5 x ULN	269 (52.7%)	75 (51.0%)	0.71	344 (52.4%)
baseline hemoglobin, g/dL	14.837 ± 1.276	14.437 ± 1.414	0.0024	14.748 ± 1.317
BMI	28.957 ± 5.558	27.671 ± 5.093	0.0094	28.670 ± 5.480
History of type 2 diabetes	57 (11.2%)	15 (10.2%)	0.74	72 (11.0%)
History of anxiety	67 (13.1%)	32 (21.8%)	0.0100	99 (15.1%)
History of depression	110 (21.6%)	32 (21.8%)	0.96	142 (21.6%)
History of fatigue	56 (11.0%)	19 (12.9%)	0.51	75 (11.4%)
History of sleep disorders	95 (18.6%)	28 (19.0%)	0.91	123 (18.7%)
Baseline PROs (range)				
SF-36: physical functioning (0-100)	81.814 ± 22.874	75.000 ± 27.507	0.0169	80.296 ± 24.128
SF-36: role physical (0-100)	80.510 ± 24.978	71.176 ± 31.468	0.0085	78.446 ± 26.805
SF-36: bodily pain (0-100)	74.543 ± 24.815	67.843 ± 29.850	0.05	73.059 ± 26.138
SF-36: general health (0-100)	66.682 ± 21.072	62.980 ± 22.850	0.10	65.855 ± 21.519
SF-36: vitality (0-100)	62.610 ± 22.533	59.574 ± 24.356	0.22	61.934 ± 22.967
SF-36: social functioning (0-100)	82.252 ± 22.669	78.723 ± 28.999	0.79	81.467 ± 24.240
SF-36: role emotional (0-100)	85.362 ± 21.689	79.492 ± 26.273	0.0283	84.056 ± 22.897
SF-36: mental health (0-100)	74.968 ± 17.934	73.661 ± 20.647	0.85	74.677 ± 18.563
SF-36: physical component summary (25-60)	50.857 ± 8.814	48.002 ± 10.971	0.0130	50.223 ± 9.402
SF-36: mental component summary (15-62)	50.752 ± 9.370	50.088 ± 11.158	0.90	50.605 ± 9.790
FACIT-F: physical well-being (0-28)	23.422 ± 4.665	21.962 ± 6.175	0.05	23.108 ± 5.057
FACIT-F: emotional well-being (0-24)	18.438 ± 4.103	18.657 ± 4.087	0.69	18.485 ± 4.097
FACIT-F: social well-being (0-28)	21.603 ± 5.948	21.656 ± 5.355	0.67	21.614 ± 5.819
FACIT-F: functional well-being (0-28)	20.964 ± 5.846	19.782 ± 6.049	0.05	20.713 ± 5.903
FACIT-F: fatigue (0-52)	40.283 ± 10.894	37.710 ± 13.484	0.18	39.725 ± 11.540

FACIT-F: total (0-160)	$124.761 \pm 24.966$	$119.431 \pm 30.023$	0.17	$123.624 \pm 26.187$
CLDQ-HCV: activity/energy (1-7)	$5.483 \pm 1.269$	$5.160 \pm 1.537$	0.09	$5.414 \pm 1.335$
CLDQ-HCV: emotional (1-7)	$5.513 \pm 1.119$	$5.395 \pm 1.302$	0.66	$5.487 \pm 1.160$
CLDQ-HCV: worry (1-7)	$5.428 \pm 1.234$	$5.471 \pm 1.336$	0.46	$5.437 \pm 1.255$
CLDQ-HCV: systemic (1-7)	$5.258 \pm 1.213$	$4.903 \pm 1.405$	0.0309	$5.182 \pm 1.263$
CLDQ-HCV: total (1-7)	$5.420 \pm 1.054$	$5.232 \pm 1.228$	0.25	$5.380 \pm 1.095$
WPAI:SHP: work productivity (1-0)	$0.105 \pm 0.205$	$0.114 \pm 0.218$	0.69	$0.107 \pm 0.207$
WPAI:SHP: absenteeism (1-0)	$0.028 \pm 0.117$	$0.040 \pm 0.128$	0.12	$0.030 \pm 0.119$
WPAI:SHP: presenteeism (1-0)	$0.077 \pm 0.156$	$0.073 \pm 0.132$	0.91	$0.077 \pm 0.151$
WPAI:SHP: activity (1-0)	$0.143 \pm 0.227$	$0.217 \pm 0.296$	0.0273	$0.159 \pm 0.245$
SF-6D health utility (20-100)	$72.347 \pm 14.571$	$69.454 \pm 17.029$	0.12	$71.708 \pm 15.182$
Treatment-related adverse events and SVR				
Blood-related and anemia	55 (10.8%)	56 (38.1%)	<0.0001	111 (16.9%)
Fatigue	269 (52.7%)	93 (63.3%)	0.0239	362 (55.1%)
Flu-like symptoms	194 (38.0%)	61 (41.5%)	0.45	255 (38.8%)
Gastrointestinal	195 (38.2%)	82 (55.8%)	0.0001	277 (42.2%)
Musculo-skeletal	153 (30.0%)	54 (36.7%)	0.12	207 (31.5%)
Nervous	212 (41.6%)	76 (51.7%)	0.0292	288 (43.8%)
Psychiatric	227 (44.5%)	71 (48.3%)	0.42	298 (45.4%)
Skin and subcutaneous tissue	201 (39.4%)	53 (36.1%)	0.46	254 (38.7%)
Other	304 (59.6%)	123 (83.7%)	<0.0001	427 (65.0%)
No adverse events	50 (9.8%)	3 (2.0%)	0.0023	53 (8.1%)
SVR-12	438 (85.9%)	110 (74.8%)	0.0015	548 (83.4%)

#### IFN-free RBV-containing regimens

N	2,131	397	2,528	
Age, years	$51.395 \pm 9.811$	$50.914 \pm 9.976$	0.23	$51.319 \pm 9.837$
Male gender	1,385 (65.0%)	257 (64.7%)	0.92	1,642 (65.0%)
Caucasian	1,862 (87.4%)	332 (83.6%)	0.0428	2,194 (86.8%)
African-American	169 (7.9%)	40 (10.1%)	0.15	209 (8.3%)
Asian	52 (2.4%)	12 (3.0%)	0.50	64 (2.5%)
Enrolled in the U.S.	1,258 (59.0%)	207 (52.1%)	0.0106	1,465 (58.0%)
Employed	1,062 (61.7%)	171 (51.2%)	0.0003	1,233 (60.0%)
Cirrhosis	397 (18.6%)	71 (17.9%)	0.74	468 (18.5%)
Treatment-naïve	1,311 (61.5%)	291 (73.3%)	<0.0001	1,602 (63.4%)
HCV genotype 1	920 (44.5%)	182 (47.2%)	0.34	1,102 (44.9%)
HCV genotype 2	353 (17.1%)	46 (11.9%)	0.0117	399 (16.3%)
HCV genotype 3	789 (38.2%)	158 (40.9%)	0.31	947 (38.6%)
HCV genotypes 4, 5, 6	5 (0.2%)	0 (0.0%)	0.33	5 (0.2%)
coinfection with HIV	353 (16.6%)	142 (35.8%)	<0.0001	495 (19.6%)
baseline HCV RNA > 6 log 10/mL	1,612 (75.6%)	290 (73.0%)	0.27	1,902 (75.2%)
baseline ALT > 1.5 x ULN	1,198 (56.2%)	204 (51.4%)	0.08	1,402 (55.5%)
baseline hemoglobin, g/dL	$14.727 \pm 1.293$	$14.376 \pm 1.484$	<0.0001	$14.673 \pm 1.330$
BMI	$27.070 \pm 5.077$	$26.628 \pm 4.950$	0.13	$27.001 \pm 5.059$
History of type 2 diabetes	198 (9.3%)	36 (9.1%)	0.89	234 (9.3%)
History of anxiety	329 (15.4%)	62 (15.6%)	0.93	391 (15.5%)
History of depression	555 (26.0%)	134 (33.8%)	0.0015	689 (27.3%)
History of fatigue	257 (12.1%)	37 (9.3%)	0.12	294 (11.6%)

History of sleep disorders	401 (18.8%)	74 (18.6%)	0.93	475 (18.8%)
Baseline PROs (range)				
SF-36: physical functioning (0-100)	82.378 ± 21.751	78.487 ± 24.988	0.0163	81.758 ± 22.338
SF-36: role physical (0-100)	77.078 ± 26.279	72.166 ± 28.281	0.0020	76.298 ± 26.662
SF-36: bodily pain (0-100)	72.617 ± 25.487	68.717 ± 27.279	0.0156	71.999 ± 25.813
SF-36: general health (0-100)	63.551 ± 22.135	60.932 ± 22.119	0.0376	63.138 ± 22.149
SF-36: vitality (0-100)	60.772 ± 22.896	58.128 ± 22.576	0.0285	60.358 ± 22.861
SF-36: social functioning (0-100)	79.872 ± 24.887	73.669 ± 26.429	<0.0001	78.886 ± 25.235
SF-36: role emotional (0-100)	81.682 ± 23.954	76.356 ± 26.532	0.0002	80.839 ± 24.452
SF-36: mental health (0-100)	72.862 ± 18.653	68.533 ± 20.357	0.0004	72.187 ± 18.990
SF-36: physical component summary (25-60)	50.547 ± 8.662	49.238 ± 9.267	0.0133	50.346 ± 8.768
SF-36: mental component summary (15-62)	49.267 ± 10.309	47.190 ± 11.124	0.0016	48.948 ± 10.462
FACIT-F: physical well-being (0-28)	23.481 ± 4.993	22.557 ± 5.479	0.0022	23.334 ± 5.084
FACIT-F: emotional well-being (0-24)	18.180 ± 4.239	17.807 ± 4.332	0.17	18.120 ± 4.256
FACIT-F: social well-being (0-28)	21.067 ± 6.331	19.321 ± 7.017	<0.0001	20.789 ± 6.475
FACIT-F: functional well-being (0-28)	20.138 ± 6.078	19.047 ± 6.366	0.0039	19.963 ± 6.136
FACIT-F: fatigue (0-52)	39.381 ± 11.470	37.422 ± 12.175	0.0050	39.065 ± 11.606
FACIT-F: total (0-160)	122.312 ± 26.989	116.162 ± 28.590	0.0002	121.331 ± 27.336
CLDQ-HCV: activity/energy (1-7)	5.341 ± 1.335	5.138 ± 1.433	0.0284	5.309 ± 1.353
CLDQ-HCV: emotional (1-7)	5.419 ± 1.195	5.222 ± 1.276	0.0118	5.387 ± 1.210
CLDQ-HCV: worry (1-7)	5.519 ± 1.243	5.411 ± 1.243	0.09	5.502 ± 1.243
CLDQ-HCV: systemic (1-7)	5.093 ± 1.287	4.872 ± 1.335	0.0052	5.058 ± 1.297
CLDQ-HCV: total (1-7)	5.343 ± 1.118	5.165 ± 1.168	0.0103	5.315 ± 1.128
WPAI:SHP: work productivity (1-0)	0.115 ± 0.215	0.119 ± 0.201	0.60	0.116 ± 0.214
WPAI:SHP: absenteeism (1-0)	0.030 ± 0.129	0.024 ± 0.092	0.80	0.030 ± 0.125
WPAI:SHP: presenteeism (1-0)	0.086 ± 0.161	0.094 ± 0.155	0.40	0.087 ± 0.160
WPAI:SHP: activity (1-0)	0.175 ± 0.253	0.190 ± 0.257	0.33	0.177 ± 0.254
SF-6D health utility (20-100)	70.595 ± 14.380	67.319 ± 13.555	0.0001	70.095 ± 14.303
Treatment-related adverse events and SVR				
Blood-related and anemia	131 (6.1%)	94 (23.7%)	<0.0001	225 (8.9%)
Fatigue	770 (36.1%)	171 (43.1%)	0.0086	941 (37.2%)
Flu-like symptoms	68 (3.2%)	11 (2.8%)	0.66	79 (3.1%)
Gastrointestinal	564 (26.5%)	117 (29.5%)	0.22	681 (26.9%)
Musculo-skeletal	238 (11.2%)	44 (11.1%)	0.96	282 (11.2%)
Nervous	531 (24.9%)	106 (26.7%)	0.45	637 (25.2%)
Psychiatric	593 (27.8%)	110 (27.7%)	0.96	703 (27.8%)
Skin and subcutaneous tissue	486 (22.8%)	98 (24.7%)	0.41	584 (23.1%)
Other	583 (27.4%)	144 (36.3%)	0.0003	727 (28.8%)
No adverse events	657 (30.8%)	87 (21.9%)	0.0003	744 (29.4%)
SVR-12	1,819 (85.4%)	314 (79.1%)	0.0016	2,133 (84.4%)

#### IFN-free RBV-free regimens

N	1,436	57	1,493	
Age, years	53.394 ± 9.638	51.509 ± 10.984	0.38	53.322 ± 9.695
Male gender	969 (67.5%)	35 (61.4%)	0.34	1,004 (67.2%)
Caucasian	1,107 (77.1%)	41 (71.9%)	0.36	1,148 (76.9%)
African-American	289 (20.1%)	13 (22.8%)	0.62	302 (20.2%)
Asian	21 (1.5%)	0 (0.0%)	0.36	21 (1.4%)

Enrolled in the U.S.	1,147 (79.9%)	49 (86.0%)	0.26	1,196 (80.1%)
Employed	863 (60.5%)	24 (44.4%)	0.0180	887 (59.9%)
Cirrhosis	245 (17.1%)	11 (19.3%)	0.67	256 (17.2%)
Treatment-naïve	968 (67.4%)	44 (77.2%)	0.12	1,012 (67.8%)
HCV genotype 1	1,426 (99.9%)	54 (94.7%)	<0.0001	1,480 (99.7%)
HCV genotype 2	0	0		0
HCV genotype 3	0	0		0
HCV genotypes 4, 5, 6	2 (0.1%)	3 (5.3%)	<0.0001	5 (0.3%)
coinfection with HIV	319 (22.2%)	16 (28.1%)	0.30	335 (22.4%)
baseline HCV RNA > 6 log 10/mL	1,204 (83.8%)	47 (82.5%)	0.78	1,251 (83.8%)
baseline ALT > 1.5 x ULN	692 (48.2%)	30 (52.6%)	0.51	722 (48.4%)
baseline hemoglobin, g/dL	14.740 ± 1.386	14.374 ± 1.204	0.06	14.726 ± 1.381
BMI	27.276 ± 4.980	27.970 ± 5.653	0.46	27.302 ± 5.007
History of type 2 diabetes	169 (11.8%)	8 (14.0%)	0.60	177 (11.9%)
History of anxiety	290 (20.2%)	12 (21.1%)	0.87	302 (20.2%)
History of depression	397 (27.6%)	22 (38.6%)	0.07	419 (28.1%)
History of fatigue	200 (13.9%)	3 (5.3%)	0.06	203 (13.6%)
History of sleep disorders	310 (21.6%)	13 (22.8%)	0.83	323 (21.6%)
Baseline PROs (range)				
SF-36: physical functioning (0-100)	80.672 ± 24.133	74.218 ± 27.547	0.17	80.436 ± 24.285
SF-36: role physical (0-100)	76.622 ± 27.781	75.490 ± 28.854	0.77	76.582 ± 27.809
SF-36: bodily pain (0-100)	72.531 ± 25.688	71.679 ± 27.241	0.88	72.500 ± 25.736
SF-36: general health (0-100)	65.613 ± 22.022	63.637 ± 22.105	0.45	65.542 ± 22.021
SF-36: vitality (0-100)	61.153 ± 23.241	55.896 ± 24.404	0.10	60.964 ± 23.296
SF-36: social functioning (0-100)	80.182 ± 25.063	75.472 ± 29.105	0.44	80.014 ± 25.222
SF-36: role emotional (0-100)	81.400 ± 25.516	78.922 ± 26.265	0.60	81.314 ± 25.538
SF-36: mental health (0-100)	73.655 ± 19.024	69.811 ± 20.684	0.13	73.516 ± 19.092
SF-36: physical component summary (25-60)	50.267 ± 9.085	49.655 ± 10.031	0.97	50.245 ± 9.117
SF-36: mental component summary (15-62)	49.654 ± 10.432	48.079 ± 10.791	0.19	49.599 ± 10.445
FACIT-F: physical well-being (0-28)	23.281 ± 5.264	22.059 ± 5.572	0.0451	23.238 ± 5.278
FACIT-F: emotional well-being (0-24)	18.411 ± 4.318	17.529 ± 4.388	0.11	18.381 ± 4.322
FACIT-F: social well-being (0-28)	21.153 ± 6.220	20.741 ± 6.688	0.77	21.139 ± 6.235
FACIT-F: functional well-being (0-28)	20.344 ± 6.056	18.712 ± 7.395	0.17	20.286 ± 6.112
FACIT-F: fatigue (0-52)	39.097 ± 12.035	35.798 ± 12.704	0.06	38.976 ± 12.071
FACIT-F: total (0-160)	122.386 ± 27.559	115.134 ± 30.163	0.09	122.130 ± 27.677
CLDQ-HCV: activity/energy (1-7)	5.369 ± 1.375	5.093 ± 1.516	0.21	5.359 ± 1.381
CLDQ-HCV: emotional (1-7)	5.471 ± 1.216	5.192 ± 1.305	0.09	5.461 ± 1.220
CLDQ-HCV: worry (1-7)	5.530 ± 1.278	5.306 ± 1.383	0.25	5.522 ± 1.282
CLDQ-HCV: systemic (1-7)	5.087 ± 1.279	4.777 ± 1.379	0.10	5.076 ± 1.283
CLDQ-HCV: total (1-7)	5.365 ± 1.142	5.092 ± 1.264	0.13	5.355 ± 1.147
WPAI:SHP: work productivity (1-0)	0.097 ± 0.193	0.125 ± 0.197	0.57	0.098 ± 0.193
WPAI:SHP: absenteeism (1-0)	0.022 ± 0.113	0.000 ± 0.000	0.25	0.022 ± 0.111
WPAI:SHP: presenteeism (1-0)	0.075 ± 0.144	0.125 ± 0.197	0.39	0.076 ± 0.146
WPAI:SHP: activity (1-0)	0.176 ± 0.264	0.215 ± 0.278	0.23	0.177 ± 0.265
SF-6D health utility (20-100)	71.106 ± 15.038	68.660 ± 14.070	0.23	71.019 ± 15.006
Treatment-related adverse events and SVR				
Blood-related and anemia	4 (0.3%)	0 (0.0%)	0.69	4 (0.3%)
Fatigue	274 (19.1%)	13 (22.8%)	0.48	287 (19.2%)

Flu-like symptoms	17 (1.2%)	3 (5.3%)	0.0086	20 (1.3%)
Gastrointestinal	261 (18.2%)	12 (21.1%)	0.58	273 (18.3%)
Musculo-skeletal	91 (6.3%)	6 (10.5%)	0.21	97 (6.5%)
Nervous	304 (21.2%)	9 (15.8%)	0.33	313 (21.0%)
Psychiatric	167 (11.6%)	5 (8.8%)	0.51	172 (11.5%)
Skin and subcutaneous tissue	106 (7.4%)	5 (8.8%)	0.69	111 (7.4%)
Other	158 (11.0%)	11 (19.3%)	0.05	169 (11.3%)
No adverse events	764 (53.2%)	29 (50.9%)	0.73	793 (53.1%)
SVR-12	1,395 (97.1%)	47 (82.5%)	<0.0001	1,442 (96.6%)



