

Supplementary Table 1. The demographic information and biochemical parameters. AST: aspartate aminotransferase; LDH: lactic dehydrogenase; CK: Creatine Kinase; α -HBDH: α -Hydroxybutyrate dehydrogenase; HCY: Homocysteine; FA: folic acid; B12: Serum Vitamin B12; MCV: mean corpuscular volume.

	Baseline: Mean (SD) (n=430)	Normal range
Age	23.90(2.765)	/
Years of abuse	2.23 (0.635)	/
Dosage(ml/day)	1796.88(453.915)	/
Craving	54.02(19.930)	/
Sleep	12.99(4.637)	/
Anxiety	45.49(10.750)	/
Depression	27.76(12.538)	/
AST(U/L)	44.04(17.337)	15~40U/L
LDH(U/L)	246.77(79.96)	109~245U/L
CK(U/L)	237.45(172.005)	18~198U/L
α -HBDH(U/L)	165.15(62.512)	72~182U/L
HCY(μ mol/L)	21.26(6.242)	<6 μ mol/L
FA (nmol/L)	24.88(8.907)	4.5-34.0nmol/L
B12(pg/ml)	801.69(223.407)	>150pg/ml
MCV (fl)	102.30(3.169)	86~100fl

Supplementary Table 2. Difference between relapsed individuals and non-relapsed individuals at baseline, according to their relapse status at the third month. Bonferroni corrected. AST: aspartate aminotransferase; LDH: lactic dehydrogenase; CK: Creatine Kinase; α -HBDH: α -Hydroxybutyrate dehydrogenase; HCY: Homocysteine; FA: folic acid; B12: Serum Vitamin B12; MCV: mean corpuscular volume

	Relapsed individuals (N=237)	Non-relapsed individuals (N=193)	Statistics		
	Mean (SD)	Mean (SD)	t	df	p
Age	23.93 (2.85)	23.90 (2.64)	0.115	428	.908
Years of abuse	2.23 (0.65)	2.26(.64)	-0.465	428	.642
Onset Age	19.61(2.52)	19.779 (2.69)	-0.613	428	.540
Number of siblings	0.119 (0.38)	0.093(0.31)	0.793	428	.428
Daily dosage (ml)	1817.72(454.39)	1775.75(466.14)	0.942	428	.347
	N	N	X ²	df	p
Education					
Middle	5	4	0.386	3	0.943
High	46	41			
College	154	120			
Master	32	28			
Daily usage period					
Morning	176	227			
Noon	7	10	1.665	2	0.435
Night	10	10			
Situation of use					
Home	14	15			
Rental	81	58			
Car	33	25	8.976	5	0.110
Hotel	49	52			
Internet café	38	17			
bar	22	26			
Nationality					
Chinese Korean	3	0			
Han	221	190			
Hui	4	1	7.312	4	0.120
Manchu	8	2			
Mongol	1	0			

Residence						
South of China	71	58	0.816	2	0.665	
North of China	165	135				
Family guardian						
Father	26	31				
Mother	61	49	2.477	2	0.290	
Grandparents	150	113				
Parental marriage						
Normal	26	14				
Separate	67	50				
Divorce	101	80	4.774	4	0.311	
Remarry	39	46				
Other	4	3				
Income(monthly)						
<10000	38	28	1.313	3	0.726	
10000-20000	70	56				
20000-30000	102	80				
>30000	27	29				
Sources of first use						
Friends	114	95				
Schoolmate	66	55	0.187	2	0.911	
bar	57	43				
Number of Friends use N₂O						
0	16	21				
1	40	30				
2	60	56	3.696	3	0.296	
>3	121	86				
Smoking	203	173	1.537	1	0.215	
Alcohol use	80	64	0.017	1	0.897	
Alcohol use frequency						
1-3	20	15				
3-5	19	24				
5-10	22	12	3.639	4	0.457	
>10	19	13				
Baseline	Mean (SD)	Mean (SD)	t	df	p	
AST	43.99(17.76)	44.31(16.88)	-0.189	428	0.850	
LDH	251.51(90.06)	248.90(69.29)	0.331	428	0.741	
CK	242.65(201.98)	235.24(118.28)	0.451	428	0.652	
α -HBDH	165.42(68.84)	168.25(51.82)	-0.486	425.48	0.627	
HCY	21.17(6.39)	21.36(6.39)	-0.310	428	0.757	
FA	24.04(9.41)	26.89 (7.49)	-3.410	428	0.001	
B12	792.68(216.86)	813.92 (230.54)	-0.982	428	0.327	
MCV	102.61(2.97)	101.95(3.42)	2.117	428	0.035	
Anxiety score	46.96 (11.40)	42.76(9.51)	4.091	428	<.0001	
Depression score	26.89(11.47)	29.56 (13.77)	-2.194	428	0.029	
Sleep score	12.99(4.40)	12.88(4.73)	0.263	428	0.793	
Craving score	55.55 (20.64)	52.67(18.71)	1.498	428	0.135	