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The age of onset of substance use is related to the coping strategies to deal with treatment in men with substance use disorder

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ABSTRACT

Background. The age of onset of substance use (OSU) as well as the coping strategies (CS) influence both the development and the course of Substance Use Disorders (SUD). We aim to examine the differences in the CS applied to deal with treatment in men with SUD, considering whether the age of OSU had begun at age 16 or earlier (OSU \leq 16) or at 17 years or later (OSU \geq 17), as well as the associations of the CS with clinical variables were studied.

Methods. A total of 122 patients with at least three months of abstinence, 60 with $OSU \le 16$ and 62 with $OSU \ge 17$, were evaluated through the Coping Strategies Inventory and clinical assessment tools.

Results. The OSU \leq 16 patients were younger and presented a worse clinical state. Compared to the norms, the SUD patients were less likely to use adaptive CS, although this was more remarkable for the OSU \leq 16 group. Furthermore, the OSU \leq 16 patients presented a CS pattern of higher Disengagement, with lesser use of Social Support and higher Problem Avoidance and Social Withdrawal. In the whole SUD sample, the severity of addiction, number of relapses and age of OSU (as a continuous variable) were related to maladaptive coping. Nevertheless, the cut-off age of OSU modulated these results.

Conclusions. The OSU \leq 16 was a risk factor for presenting greater clinical severity and a more dysfunctional CS profile to deal with treatment. Thus, the cut-off age considered has allowed us to differentiate SUD patients with more vulnerability to present worse clinical prognosis who may require specific prevention and rehabilitation strategies discussed throughout this work.

Subjects Psychiatry and Psychology, Public Health

Keywords Coping strategies, Addiction severity, Onset substance use, Treatment coping, Substance use disorders

INTRODUCTION

Substance Use Disorders (SUD) are considered a public health issue since they have severe personal and community consequences, as well as a high worldwide prevalence (United Nations Office on Drugs and Crime (*UNODC*) 2015). Despite welfare programs, patients have a high variability of response to interventions (*Kampman et al.*, 2007) and high rates

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of relapse (*Suijkerbuijk et al., 2015; Witkiewitz & Marlatt, 2004*). Moreover, the age of onset of substance use (OSU) usually occurs at very early ages and it has been established as a strong predictor to future SUD (*Woodcock, Lundahl & Stoltman, 2015*) and linked both to worse clinical course and cognitive functioning (*Capella, Benaiges & Adan, 2015; Eddie, Epstein & Cohn, 2015; Hammond, Mayes & Potenza, 2014; Kendler et al., 2013*) and greater brain alterations (*Elofson, Gongvatana & Carey, 2013*). Thus, it is necessary to further study the risk factors and course severity in SUD, such as age of OSU, with the aim of improving prevention and treatment strategies according to their characteristics, to make them more effective.

In this line, we have attempted to clarify the reasons that lead people to consume substances. While the age of OSU is often linked to social goals and to its positive reinforcement, once SUD consumption has developed it is maintained in order to reduce the stress-based negative affect (*Blevins et al., 2014*; *Dermody, Cheong & Manuck, 2013*). Although there are multiple factors that influence the clinical prognosis of SUD patients, certain psychological characteristics are essential. Among these, we can mention the coping strategies (CS) used to face adversity since it has been shown that they influence both the development and course of SUD and its treatment outcome (*Marquez-Arrico, Benaiges & Adan, 2015*; *Walker & Stephens, 2014*).

Lazarus & Folkman (1984) define CS as cognitive and behavioral responses aimed at managing internal or external demands. They have established two major ways of coping: engagement, aimed at dealing with the stressors or their related emotions, and generally considerate adaptive; and disengagement, targeted to avoid the stressful situations or their related emotions, and mainly regarded as maladaptive (*Carver & Connor-Smith*, 2010; *Skinner et al.*, 2003; *Tobin et al.*, 1989). People are likely to show a relatively stable disposition towards the habitual use of certain CS to diverse stressors, which to some extent would vary depending on situation-specific variables in a concrete coping episode or stressor (*Bauer et al.*, 2016; *Bouchard*, *Guillemette & Landry-Léger*, 2004; *Lazarus & Folkman*, 1984) and age (*Mauro et al.*, 2015; *Woodhead et al.*, 2014).

Substance consumption is considered as a type of coping behavior to avoid stress, focused on emotions and directed to temporarily alleviate the negative affect that certain stressors generate, although in the long term it will trigger more severe problems, such as the need to consume again (*Bavojdan, Towhidi & Rahmati, 2011; Buckner et al., 2015; Hruska et al., 2011*). The avoidance-based coping style has been established as a risk factor with a poor prognosis for the initiation and maintenance of SUD. In contrast, problem-focused coping is considered a protective factor against consumption (*Blevins et al., 2014; Coriale et al., 2012; Dermody, Cheong & Manuck, 2013; Marquez-Arrico, Benaiges & Adan, 2015; Walker & Stephens, 2014; Woodhead et al., 2014*) and is associated to better mental health (*Bavojdan, Towhidi & Rahmati, 2011; Nyamathi et al., 2010*). Similarly, social support is a CS with a protective effect against stress (*Hyman et al., 2009*) and the development of depression (*Aarts et al., 2015*), having been observed that it correlates negatively with the relapse rate of SUD patients (*Chauchard, Septfons & Chabrol, 2013; Dolan et al., 2013; Hägele et al., 2014*). However, recent results are controversial (*Nyamathi et al., 2010*) and need to be contrasted.

Adolescence is a risk stage for the onset of several psychiatric disorders, such as those related with substance use (*Hägele et al., 2014*; *Kirst et al., 2014*). Since early OSU is associated to future SUD development (*Woodcock, Lundahl & Stoltman, 2015*) and to more severe characteristics (*Capella, Benaiges & Adan, 2015*; *Eddie, Epstein & Cohn, 2015*; *Hammond, Mayes & Potenza, 2014*; *Kendler et al., 2013*), studying the CS of addicts and their clinical implications, considering the age of OSU, is a research area of undoubted clinical interest.

Based on characteristics of brain ontogeny (*Kunert, Derichs & Irle, 1996; Lambe, Krimer & Goldman-Rakic, 2000; Shaw et al., 2006; Sundram, 2006*), previous studies have found that patients who begin consumption at age 16 or earlier, when compared to those with onset at age 17 or later, have a lower premorbid intelligence quotient (*Capella, Benaiges & Adan, 2015; Pope et al., 2003*), worse neuropsychological performance (*Ehrenreic et al., 1999; Jockers-Scherübl et al., 2007*) and less cerebral and gray matter volume (*Wilson et al., 2000*). However, considering this cut-off age, no previous studies have provided data about the possible differences in CS patterns and their relationship with clinical variables.

Our paper has two aims. The first is to assess the differences in the CS profile to deal with the treatment of men diagnosed with SUD, depending on whether they initiated substance use at age 16 or earlier ($OSU \le 16$) or at age 17 or later ($OSU \ge 17$), as well as in relation to normative data. The second is to explore the relationships among CS- and SUD-related clinical variables.

METHOD

Study design and participants

We enrolled 122 patients under SUD treatment in different healthcare resources (ambulatory drug use treating or residential in therapeutic community) in a cross-sectional study design. All were male, given the high prevalence of this gender in SUD (*UNODC*, 2015) and to avoid biasing the results due to sex differences (*Woodhead et al.*, 2014). Participants were derived from treatment centers after selection according our inclusion/exclusion criteria. In a first evaluation session we confirmed the diagnosis by a researcher (trained clinical psychology postgraduate) responsible of clinical assessment. In a second session, the coping strategies assessment tool along with other tests not presented in this manuscript were administered. After collecting data, they were assigned to two groups according to age of OSU: one for OSU at age 16 or earlier (OSU ≤ 16 ; n = 60), and one for OSU at age 17 or later (OSU ≥ 17 ; n = 62). The consideration of this age cut-off was based on the neurodevelopmental characteristics (*Kunert, Derichs & Irle, 1996*; *Lambe, Krimer & Goldman-Rakic, 2000*; *Shaw et al., 2006*; *Sundram, 2006*), as well as the differences in cognitive performance found in previous studies (*Capella, Benaiges & Adan, 2015*; *Ehrenreic et al., 1999*; *Jockers-Scherübl et al., 2007*; *Pope et al., 2003*).

The inclusion criteria were: (1) current or past diagnosis of SUD confirmed by a diagnostic interview according to the criteria in the Diagnostic and Statistical Manual of Mental Disorders, Fourth Edition Text Revised (DSM-IV-TR; *American Psychiatric Association, 2000*); (2) with abstinence for at least three months at the time of the study

(excluding caffeine or nicotine consumption), confirmed by urinalysis, to ensure the overcoming of withdrawal symptoms and minimum adherence to treatment; (3) age between 18–55 years. The exclusion criteria were: (1) presence of mental retardation or pervasive developmental disorder, history of traumatic brain injury, neurological injury or any other medical problem which could interfere in the assessment; (2) presence of a comorbid axis I mental disorder, such mood or affective disorders, confirmed by a diagnostic interview according to DSM-IV-TR criteria.

All patients provided written informed consent and were not compensated for their participation. The ethic committee of the University of Barcelona approved this study (IRB00003099), which meets the ethical principles of the declaration of Helsinki (*World Medical Association, 2013*). This study was part of a larger project on clinical characteristics, neuropsychological functioning, personality traits and circadian rhythmicity in SUD and Dual Diagnosis patients.

Clinical and sociodemographic measures

Through a structured interview designed specifically for our study and the Structural Clinical Interview for the DSM-IV Axis I Disorders (SCID-I; *First et al.*, 1999), we collected sociodemographic (age, marital, educational and economic status) and clinical data (presence of psychiatric pathology and substance use family history, suicidal attempts, past treatment for SUD, consumption pattern, type of drugs used, age of OSU, duration of drug use, residential or ambulatory treatment, medication, abstinence periods and relapses). This information was confirmed with the medical history of the centers' databases and with the patients' treating psychiatrist.

The Clinical Global Impression questionnaire (CGI; *Guy*, *1976*) was administered as a subjective measure of clinical severity. Furthermore, severity of SUD was assessed using the Drug Abuse Screening Test (DAST-20; *Skinner*, *1982*) through its Spanish version (*Gálvez* & *Fernández*, *2010*), which provides a total score from 0 to 20 (0 no addiction, 1–5 low, 6–10 intermediate, 11–15 substantial, and 16–20 severe).

Coping strategies assessment

CS were assessed by means of the Spanish version (*Cano-García, Rodríguez-Franco & Martínez, 2007*) of the Coping Strategies Inventory (CSI; *Tobin et al., 1989*). Patients assessed the frequency with which they had used the strategies described to deal with their SUD treatment. The CSI is composed by 41 items with 5-point Likert scale answers, of which 40 configure the primary scales with one additional item on self-perceived coping ability. The CSI has a hierarchical structure composed by eight primary, four secondary and two tertiary scales. The primary scales are: Problem Solving, Cognitive Restructuring, Social Support, Express Emotions, Problem Avoidance, Wishful Thinking, Social Withdrawal, and Self-Criticism. The secondary scales are: Problem Focused Engagement (composed by Problem Solving and Cognitive Restructuring), Emotion Focused Engagement (Social Support and Express Emotions), Problem Focused Disengagement (Problem Avoidance and Wishful Thinking) and Emotion Focused Disengagement (Social Withdrawal and Self Criticism). The tertiary scales of the CSI are Engagement (Problem and Emotion Focused Engagement) and Disengagement (Problem and Emotion Focused Disengagement).

Statistical analyses

Descriptive statistics and frequencies were calculated to describe the total study sample. Differences in sociodemographic and clinical variables between groups were explored with the Mann–Whitney U test (U) or with the Chi Square test (χ^2) for categorical variables. The Student's *t*-test (*t*) was used when the quantitative data fulfilled the necessary conditions, and the U test was used instead when those conditions were not met. Factorial analysis of eight factors for CSI was performed with a normalized varimax rotation employed to achieve factor simplicity. Furthermore, internal consistency for the primary scales was calculated with the Cronbach's alpha coefficients. Three multivariate analyses of covariance (MANCOVA) were performed considering the primary, secondary and tertiary CSI scales. Age was considered as covariate, since it could be a confounding factor (*Mauro et al., 2015*; Woodhead et al., 2014). An additional analysis was carried out considering the treatment regimen (residential or ambulatory), to assess whether this was an indicator of differences in the CS profile to deal with treatment related to the recruitment of patients and not related to the age of OSU. The Bonferroni test was applied in all analyses to reduce the occurrence of a type I error. The effect size was calculated with the partial Eta squared (η_p^2) , assuming a value of 0.01 as low, of 0.04 as moderate and of 0.1 as high (*Huberty*, 2002). The data were compared to the Spanish norms, only available for the primary scales (Cano-García, Rodríguez-Franco & Martínez, 2007), using percentiles.

The relationships between CS and SUD clinical variables were studied in two steps, both for the total sample and for each group. First, we carried out correlational analyses between CS and clinical data; then, the significant results were introduced in a multiple stepwise regression analyses with CS as dependent variables.

Data were analyzed using the Statistical Package for the Social Sciences (SPSS; version 15.0), considering bilateral statistical significance with an established type I error at 5% (p < .05).

RESULTS

Differences in sociodemographic and clinical data

The total sample was aged 20 to 55 (M = 35.97, SD = 8.31) and most of the patients had completed the Spanish compulsory education (from 6 to 16 years). Regarding sociodemographic variables, the only observed difference between groups was the lower mean age of the OSU \leq 16 group (p < .001), while both groups did not differ in years of education, marital and economic status. The analyses of the clinical variables provided no significant differences between groups regarding relatives with other psychiatric disorders than SUD and number of suicidal attempts. Instead, in the OSU \leq 16 group it was more frequent to have relatives with SUD (p = .025). See Table 1.

With respect to SUD data, the OSU \leq 16 group had higher rates of polyconsumption (p = .030), and patients in residential rather than ambulatory treatment (p = .013), lower age of OSU (p < .001) and longer duration of drug use (p = .016). Furthermore, the groups showed differences in the type of substances used. In the OSU \leq 16 group, there were higher rates of cannabis (p < .001), and hallucinogens consumption (p = .042), while

	Total sample (<i>N</i> = 122)	$OSU \le 16$ $(N = 60)$	$OSU \ge 17$ $(N = 62)$	Statistical contrasts
Sociodemographic data				
Age (yr)	35.97 (0.75)	33.60 (1.09)	38.25 (0.96)	$U = 1,159.50^{***}$
Years of education	10.36 (0.23)	10.25 (0.33)	10.48 (0.32)	$t_{(120)} = -0.476$
Marital status				$\chi^2_{(1)} = 5.015$
Single	51.6%	51.7%	51.6%	
Separate/Divorced	22.1%	21.6%	22.5%	
Married	14.8%	13.3%	16.1%	
Stable partner	11.5%	13.3%	9.7%	
Economic status				$\chi^2_{(1)} = 4.305$
Unemployed	29.5%	30%	29%	
Active	23.8%	16.7%	30.6%	
No income	19.7%	23.3%	16.1%	
Disability pension	16.4%	20%	12.9%	
Sick leave	10.7%	10%	11.3%	
Clinical data				
Relatives with SUD	26.2%	25%	9.7%	$\chi^{2}_{(1)} = 5.024^{*}$
Relatives with others psychiatric disorder	26.1%	23.3%	29%	$\chi^2_{(1)} = 0.512$
Number of suicidal attempts	0.23 (0.66)	0.30 (0.10)	0.16 (0.06)	$t_{(120)} = 1.144$

Table 1 Descriptive statistics (frequencies or mean and standard error) of the sociodemographic and clinical data, for the total sample and groups, and the statistical contrasts carried out.

Notes.

 $OSU \le 16$, Onset of substance use at age 16 or earlier; $OSU \ge 17$, Onset of substance use at age 17 or later; yr, years; SUD, Substance Use Disorder.

*p < .05.

* *p* < .001.

in the OSU \geq 17 group there are higher rates of cocaine consumption (p = .046). In the overall sample, as well as in both groups, the substances more frequently used were cocaine, alcohol and cannabis. No differences between groups were found in the other SUD clinical characteristics studied (see Table 2).

Coping Strategies Inventory comparisons

The eight factors obtained from factorial analysis of the CSI explained the 61.95% of variance (*F*1 = 9.73, *F*2 = 9.55, *F*3 = 8.61; *F*4 = 8.35; *F*5 = 7.93; *F*6 = 7.16, *F*7 = 5.34 and F8 = 5.29). The set of items of the Problem Solving and Wishful Thinking scales are the most clearly associated with one single factor, with loadings superior to .43. Regarding the scales of Self-Criticism, Express Emotions and Cognitive Restructuring scales, only four of the five items selected converge in the factors, with values higher to .44. Finally, Social Support, Problem Avoidance and Social Withdrawal are the scales that correspond less to the items of the originals.

Cronbach's alpha coefficients of internal consistency for the primary scales were all adequate for the total sample studied: Problem Solving (0.79), Cognitive Restructuring (0.71), Social Support (0.72), Express Emotions (0.76), Problem Avoidance (0.70), Wishful Thinking (0.77), Social Withdrawal (0.74) and Self-Criticism (0.77).

SUD clinical characteristics	Total sample (<i>N</i> = 122)	$OSU \le 16$ $(N = 60)$	$OSU \ge 17$ $(N = 62)$	Statistical contrasts
Consumption pattern				
One substance	23.8%	18.3%	29%	<i>U</i> = 1,661
Two substances	34.4%	30%	38.7%	U = 1,698
Polydrug use	41.8%	51.7%	32.3%	$U = 1,499^*$
Substances used ^a				
Cocaine	87.7%	81.7%	93.5%	$\chi^2_{(1)} = 3.992^*$
Alcohol	72.1%	76.7%	67.7%	$\chi^2_{(1)} = 1.208$
Cannabis	43.4%	60%	27.4%	$\chi^2_{(1)} = 13.173^{***}$
Hallucinogens	16.4%	23.3%	9.7%	$\chi^2_{(1)} = 4.149^*$
Opioids	14.8%	16.7%	12.9%	$\chi^2_{(1)} = 0.343$
Sedatives	4.1%	5%	3.2%	$\chi^2_{(1)} = 0.244$
Age of OSU (yr)	19.16 (0.59)	14.98 (0.15)	23.20 (0.88)	$U = 17.50^{***}$
Duration of drug use (yr)	16.13 (0.77)	17.98 (1.07)	14.33 (1.06)	$t_{(120)} = 2.432^*$
Typology of treatment regimen				$\chi^2_{(1)} = 6.124^*$
Residential	62.3%	73.3%	51.6%	
Ambulatory	37.7%	26.7%	48.4%	
Daily number of medication	0.51 (0.08)	0.65 (0.14)	0.36 (0.08)	$t_{(120)} = 1.813$
Months of abstinence	8.14 (0.47)	7.60 (0.59)	8.66 (0.72)	$t_{(120)} = -1.127$
Past treatment for SUD				$\chi^2_{(1)} = 0.085$
Yes	50.5%	51.9%	48.9%	
Number of relapses				
None	57%	56.7%	57.4%	U = 1,856
One	16.5%	13.3%	19.7%	U = 1,718
Two	10.7%	15%	6.6%	U = 1,701
Three or more	15.7%	15%	16.4%	U = 1,825.50
Drug Abuse Screening Test (DAST-20)	12.15 (0.47)	12.33 (0.57)	11.94 (0.78)	$t_{(1)} = 0.418$
Clinical Global Impression (CGI)	2.45 (0.11)	2.58 (0.17)	2.33 (0.15)	$t_{(1)} = 1.069$

Table 2 Descriptive statistics (frequencies or mean and standard error) of the data related to SUD, for the total sample and groups, and the statistical contrasts carried out.

Notes.

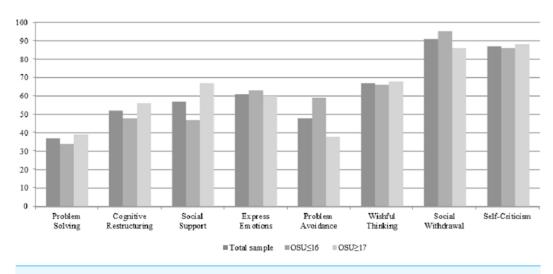
SUD, Substance Use Disorder; OSU <16, Onset of substance use at age 16 or earlier; OSU <17, Onset of substance use at age 17 or later; OSU, Onset of substance use; yr, years; DAST-20, Drug Abuse Screening Test; CGI, Clinical Global Impression.

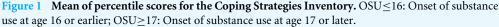
^aPercentages will not equal 100 as each participant may take more than one substance of abuse.

**p* < .05.

* p < .001.

The comparisons among percentile scores in the CSI primary scales according to the Spanish normative data (see Fig. 1) showed lower scores in the overall sample (<40 percentile) in Problem Solving, and higher scores (>60 percentile) in Express Emotions, Wishful Thinking, Social Withdrawal and Self-Criticism. When considering groups, higher scores in Wishful Thinking, Social Withdrawal and Self-Criticism were observed in both. On the other hand, the OSU \geq 17 group had higher scores in Social Support and lower scores in Problem Avoidance.





The MANCOVA analyses showed several significant differences between groups for the CSI scales (see Table 3). Results in the primary scales indicated that the OSU \leq 16 group had lower scores in Social Support (p = .019), and higher scores both in Problem Avoidance (p = .037), and in Social Withdrawal (p = .049). No differences between groups were found in the other primary and secondary scales. Regarding tertiary scales, the groups were similar in their use of Disengagement strategies, but the OSU \leq 16 group showed lower scores in Engagement strategies (p = .038).

In the additional analysis carried out considering the treatment regimen (residential or ambulatory) as a group variable, no significant differences between groups were found regarding any primary, secondary or tertiary CSI scales (p > .241; in all cases), neither considering age and age of OSU as covariates (p > .240; in all cases).

Relations among SUD, clinical characteristics and coping strategies

First, correlations between CS and clinical data, and between CS and SUD clinical characteristics, indicated that only age of OSU (considered as a continuous variable), duration of drug use, number of relapses and DAST-20 were associated with some of the CS: Wishful Thinking, Social Withdrawal, Problem Focused Disengagement, Emotion Focused Disengagement and Disengagement.

Second, the above variables were introduced in regression analyses with the different CS as dependent variables (see Table 4). In the total sample, the DAST-20 explained 11% of the variance of Problem Focused Disengagement (p = .002), 13% of Social Withdrawal (p = .001), 17% of Wishful Thinking (p < .001) and, together with age of OSU and number of relapses, 23% of the variance of Emotion Focused Disengagement (p < .001). Age of OSU and number of relapses accounted for 15% of the variance of Disengagement (p = .002).

Considering the OSU \leq 16 group, the regression analysis indicated that the model was significant in only four of the CS. The DAST-20 explained 18% of the variance of Wishful Thinking (p = .003). The DAST-20 and the number of relapses were significant for Social

Table 3Means and standard deviations for the total sample and for each group, and results of theMANCOVA analyses for both groups considering age as a covariate for the Coping Strategies Inventory(CSI).

CSI	Total sample $(N = 122)$	$OSU \le 16$ $(n = 60)$	$OSU \ge 17$ $(n = 62)$	F	Effect size
Primary subscales					
Problem solving	13.31 (4.76)	12.83 (4.69)	13.77 (4.82)	0.835	0.007
Cognitive restructuring	10.44 (5.04)	9.72 (5.05)	11.16 (4.97)	2.705	0.022
Social support	11.50 (5.03)	10.50 (5.35)	12.48 (4.52)	5.628*	0.045
Express emotions	10.37 (5.08)	10.70 (5.25) 10.05 (4.94)		0.891	0.007
Problem avoidance	5.77 (4.64)	6.85 (5.32)	4.76 (3.63)	4.471^{*}	0.036
Wishful thinking	14.69 (4.81)	14.60 (4.67)	14.77 (4.99)	0.004	0.001
Social withdrawal	9.68 (5.30)	10.77 (5.18)	8.63 (5.25)	3.959*	0.029
Self-criticism	13.57 (5.09)	13.48 (5.29)	13.66 (4.95)	0.081	0.011
Self-perceived capacity	2.70 (1.33)	2.50 (1.42)	2.90 (1.21)	1.245	0.010
Secondary subscales					
Problem focused engagement	23.78 (8.08)	22.60 (8.29)	24.91 (7.77)	2.353	0.019
Emotion focused engagement	21.79 (8.10)	21.10 (8.71)	22.45 (7.48)	0.772	0.006
Problem focused disengagement	20.53 (5.92)	21.45 (6.21)	19.62 (5.52)	2.562	0.021
Emotion focused disengagement	23.48 (8.64)	24.25 (8.28)	22.72 (8.98)	0.423	0.004
Tertiary subscales					
Engagement	45.59 (14.26)	43.70 (14.98)	47.42 (13.39)	1.944	0.016
Disengagement	43.61 (10.92)	45.87 (11.16)	41.44 (10.31)	4.417^{*}	0.036

Notes.

OSU \leq 16, Onset of substance use at age 16 or earlier; OSU \geq 17, Onset of substance use at age 17 or later. *p < .05.

Withdrawal (p = .001) and Emotion Focused Disengagement (p = .004), explaining 27% and 22% of the variance, respectively. Number of relapses explained 12% of the variance of Disengagement (p = .018).

Moreover, only four significant regression models were observed in the OSU \geq 17 group. In this case, the DAST-20 were significant for Wishful Thinking (p = .022), Social Withdrawal (p = .037), Problem Focused Disengagement (p = .010) and Emotion Focused Disengagement (p = .031), accounting for 11%, 12%, 13% and 17% of the variance, respectively.

DISCUSSION

To our knowledge, this is the first study that aims to elucidate the possible existence of differences in CS related to treatment in men with SUD, depending on whether their substance use began at age 16 or earlier, or at age 17 or later. In addition, we have also assessed the influence of clinical variables related to the SUD regarding the CS pattern.

Sociodemographic and clinical differences between groups

In relation to the sociodemographic data, both groups differ only in age, the patients in the OSU \leq 16 group being the youngest. We decided to control the possible effect of this variable on the coping results, given that with the passing of years people may develop some

Table 4 Multiple linear regression for the Coping Estrategies. Multiple linear regression for the Coping Strategies Inventory (CSI) considering as independent variables the sociodemographic and clinical data that showed significant correlations, for the total sample (N = 122) and for the OSU \geq 16 (N = 60) and OSU \geq 17 (N = 62) groups.

CSI	Adjusted R ²	F	IV ^a	β Standardized	<i>p</i> values
Total sample					
Wishful thinking	0.167	15.279***	DAST-20	0.423	.0001
Social withdrawal	0.133	11.900**	DAST-20	0.381	.001
Problem focused disengagement	0.113	10.022**	DAST-20	0.166	.002
Emotion focused disengagement	0.232	8.157***	Age of OSU (yr)	-0.276	.010
			Number of relapses	0.220	.047
			DAST-20	0.302	.007
Disengagement	0.146	7.072**	Age of OSU (yr)	-0.269	.017
			Number of relapses	0.300	.008
OSU≤16					
Wishful thinking	0.187	9.732**	DAST-20	0.456	.003
Social withdrawal	0.265	7.835**	Number of relapses	0.322	.029
			DAST-20	0.387	.010
Problem focused disengagement ^b					
Emotion focused disengagement	0.218	6.307**	Number of relapses	0.315	.038
			DAST-20	0.342	.025
Disengagement	0.120	6.159*	Number of relapses	0.378	.018
OSU≥17					
Wishful thinking	0.131	5.827*	DAST-20	0.398	.022
Social withdrawal	0.105	4.740^{*}	DAST-20	0.364	.037
Problem focused disengagement	0.171	7.577*	DAST-20	0.443	.010
Emotion focused disengagement	0.115	5.138*	DAST-20	0.377	.031
Disengagement ^b					

Notes.

 $OSU \le 16$, Onset of substance use at age 16 or earlier; $OSU \ge 17$, Onset of substance use at age 17 or later; IV, Independent Variables; DAST-20, Drug Abuse Screening Test; OSU, Onset of Substance Use; yr, years.

^aOnly significant variables are presented that comprise each explicative model. In all cases, the Tolerance values were higher than 0.91 and the Variance Inflation Factor values lower than 1.09.

^bAny explicative model was significant.

*p < .05.

 $p^{**} < .01.$

 $p^{***} p < .001.$

variation in the CS they use (*Mauro et al., 2015*; *Woodhead et al., 2014*). The substances more frequently consumed in both groups were cocaine, alcohol and cannabis. However, the patients in the OSU \leq 16 group had a higher frequency of cannabis and hallucinogens consumption, whereas cocaine was the most frequently consumed drug in the OSU \geq 17 group. These differences may reflect the social preference for a certain substance at the time the patients were developing their SUD (*European Monitoring Centre for Drugs and Drug Addiction, 2016*).

When the cut-off age of OSU was considered, the clinical characteristics of both groups fall in line with the only data currently available (*Capella, Benaiges & Adan, 2015*), that is, the OSU \leq 16 patients present a more severe clinical and SUD pattern, characterized by more substance consumption, greater duration of drug use and the need for a more intensive

treatment (residential instead of ambulatory) to achieve abstinence. Other works have analyzed the age of OSU, although without establishing a cut-off point, and have obtained similar results (*Eddie, Epstein & Cohn, 2015; Hammond, Mayes & Potenza, 2014; Kendler et al., 2013*). Moreover, the higher presence of a family history of SUD in the OSU \leq 16 group supports the findings on the genetic predisposition and early environmental exposures that precede the onset of drug use in early consumers (*Hägele et al., 2014; Hammond, Mayes* & *Potenza, 2014*). These data should be further explored in future research taking into account the possible mediation of the main substance consumed, since this family factor has not yet been found in cannabis consumers (*Pope et al., 2003*). Finally, it is interesting to note that the CS profile of the patients was independent of the type of treatment regimen in agreement with a previous study (*Adan, Antúnez & Navarro, 2017*).

Coping strategies

The SUD patients show a tendency to use emotion in order to cope with the stress generated by their treatment. There is a predominance to use maladaptive CS such as avoiding any contact with those persons related to their stressful experience (Social Withdrawal), as well as self-blame for the occurrence of the stressful situation or its improper management (Self-Criticism). When considering the most adaptive pole, the predominant CS is the release of emotions that occur in the process of stress (Express Emotions). When focusing on addressing the problem, the tendency was to do it in an inappropriate way by thinking about non-stressful alternative realities (Wishful Thinking), with a low propensity to active resolution of their difficulties (Problem Solving). This is consistent with the results of previous studies that have found a tendency to maladaptive coping, with lesser use of active coping problem-solving strategies in drug consumers (*Marquez-Arrico, Benaiges & Adan,* 2015), which could be at the basis of the onset (*Walker & Stephens, 2014*; *Woodhead et al.,* 2014) and maintenance of the SUD (*Coriale et al., 2012*). The pattern described is observed in both groups, although the OSU \geq 17 group presents a more adaptive coping profile.

When compared to the OSU \geq 17, the OSU \leq 16 group is characterized by a CS pattern of greater Disengagement (lesser use of Social Support; greater Problem Avoidance and Social Withdrawal). Thus, it seems that the patients with an early OSU present fewer interpersonal skills, showing a tendency to withdraw from their social environment and to avoid contact with other people in order to express their emotions. In this sense, we find that a higher rate of substance and medication use in adults is related to a lesser early childhood social competence (*Jones, Greenberg & Crowley, 2015*). Furthermore, in SUD patients under treatment high rates of cooperation (*Andó et al., 2012*) and stronger social support (*Chauchard, Septfons & Chabrol, 2013*; *Dolan et al., 2013*) have been related to longer abstinence periods, while social withdrawal is higher when the age of OSU is younger (*Marquez-Arrico, Benaiges & Adan, 2015*). Moreover, avoidance-based coping has been consistently identified as a moderator of SUD (*Bavojdan, Towhidi & Rahmati, 2011*; *Hruska et al., 2011*), and this could be a key factor in the tension-reduced-based models, where emotional suffering is relieved through avoidance of emotional distress (*Buckner et al., 2015*).

Influence on coping strategies of SUD clinical characteristics

The relations observed among the clinical variables and maladaptive coping are modulated by the age of OSU and, specifically, by the cut-off age studied in this work. In SUD patients, a higher severity of addiction is related to a more dysfunctional coping pattern when they face treatment (higher scores in Wishful Thinking, Social Withdrawal, Problem Focused Disengagement and Emotion Focused Disengagement). Moreover, a higher number of relapses is related to a more frequent use of a coping style based on emotional disengagement (Emotion Focused Disengagement and Disengagement strategies), which is explained by the pattern of the OSU \leq 16 group. Thus, those patients with a more severe SUD tend to use maladaptive CS (*Blevins et al., 2014*; *Marquez-Arrico, Benaiges & Adan, 2015*; *Nyamathi et al., 2010*), and this is related to a higher number of relapses (*Chauchard, Septfons & Chabrol, 2013*; *Dolan et al., 2013*), even the more so as the age of OSU decreases.

There are some limitations in our study that should be mentioned. Almost half of the patients were polyconsumers, which was an impediment to assess separately the effect of each type of substance on CS. However, their possible effect was relatively controlled since the groups had consumed the same main substances (cocaine, alcohol and cannabis). The inclusion of only men in the sample, knowing the influence of gender in CS (Bouchard, Guillemette & Landry-Léger, 2004; Woodhead et al., 2014), limits the generalization of our results. The wide range of age in the sample may have also contributed to a type-II error. We have analyzed cross-sectional data, which does not allow establishing causal or sequential relations among variables, or if the coping strategies currently used by patients are the same as when they started the substance use or even the treatment. Moreover, the level of perceived stress and affect or mood of participants were not assessed, which could help explain their CS pattern and clinical traits (Aarts et al., 2015; Hyman et al., 2009). While the data obtained in the CSI indicated an adequate internal consistency for the primary scales, the factorial analysis with the eight factor structure model of the Spanish version (Cano-García, Rodríguez-Franco & Martínez, 2007) was only partially fulfilled, although it is widely agreed and validated. Finally, the low explanatory power obtained in the regression analyses warns us to interpret the results with caution. Future works should include larger patient samples of both men and women, differentiating the main substance consumed, with a longitudinal design that could contribute to clarify the attitudinal or situational character of coping and which variables determine it in order to obtain a better knowledge of its role in later outcomes.

Our results may have clinical interest. We find that SUD patients deal with treatment applying a dysfunctional emotion-based coping style, a style also present at the onset, maintenance and severity of the disorder (*Bavojdan, Towhidi & Rahmati, 2011*; *Blevins et al., 2014*; *Buckner et al., 2015*; *Dermody, Cheong & Manuck, 2013*; *Hruska et al., 2011*). This suggests that those interventions aimed at developing CS which may prevent consumption, such as problem-focused coping o social support, could be more effective both in addiction prevention and treatment programs. The assessment of cognitive skills seems another key factor to be considered, given that a low neuropsychological performance is related both to the development of SUD in adulthood (*Pechtel, Woodman & Lyons-Ruth, 2012*) and to the response to treatment (*Kiluk, Nich & Carroll, 2011*). In this regard, it has been shown that, in the rehabilitation of SUD patients, higher cognitive abilities are associated with greater improvement in the quality of the coping skills acquired, which in turn is indirectly associated with treatment benefits. This could be significantly relevant for patients with OSU \leq 16, since they present a worse coping style, as shown in the present study, and a lower cognitive performance, as shown in other works (*Capella, Benaiges & Adan, 2015; Ehrenreic et al., 1999; Jockers-Scherübl et al., 2007; Pope et al., 2003*). Further research is required to shed light on this issue.

CONCLUSIONS

A main priority in public health should be to target populations at risk of developing SUD or with a worse clinical prognosis, in order to design more specific intervention programs. In this sense, the cut-off age considered in our study is a contribution, since the OSU \leq 16 patients exhibited more vulnerability to present both higher clinical severity and frequency to use of a dysfunctional CS profile to cope with treatment, which were related to the severity of the addiction and relapses. Further studies are needed to explore the possible benefits of improving adaptive coping in these patients for better treatment outcomes, as well as for prevention programs of SUD.

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Competing Interests

The authors declare there are no competing interests.

Author Contributions

• Maria del Mar Capella performed the experiments, analyzed the data, contributed reagents/materials/analysis tools, wrote the paper, prepared figures and/or tables, reviewed drafts of the paper.

• Ana Adan conceived and designed the experiments, contributed reagents/materials/analysis tools, wrote the paper, prepared figures and/or tables, reviewed drafts of the paper, conceived the original idea for the study, sought funding, and wrote the protocol.

Human Ethics

The following information was supplied relating to ethical approvals (i.e., approving body and any reference numbers):

The ethics committee of the University of Barcelona granted approval to carry out the study.

Data Availability

The following information was supplied regarding data availability:

The raw data has been supplied as a Supplementary File.

Supplemental Information

Supplemental information for this article can be found online at http://dx.doi.org/10.7717/ peerj.3660#supplemental-information.

REFERENCES

- Aarts JW, Deckx L, Van Abbema DL, Tjan-Heijnen VC, Van den Akker M, Buntinx F.
 2015. The relation between depression, coping and health locus of control: differences between older and younger patients, with and without cancer. *Psychooncology* 24:950–957 DOI 10.1002/pon.3748.
- Adan A, Antúnez JM, Navarro JF. 2017. Coping strategies related to treatment in substance use disorder patients with and without comorbid depression. *Psychiatry Research* 251:325–332 DOI 10.1016/j.psychres.2017.02.035.
- **American Psychiatric Association. 2000.** *Diagnostic and statistical manual of mental disorders.* 4th edition text rev. Washington, D.C.: American Psychiatric Association.
- Andó B, Must A, Kurgyis E, Szkaliczki A, Drótos G, Rózsa S, Szikszay P, Horváth S, Janka Z, Almos PZ. 2012. Personality traits and coping compensate for disadvantageous decision-making in long-term alcohol abstinence. *Alcohol and Alcoholism* 47:18–24 DOI 10.1093/alcalc/agr144.
- Bauer MR, Harris LN, Wiley JF, Crespi CM, Krull JL, Weihs KL, Stanton AL. 2016. Dispositional and situational avoidance and approach as predictors of physical symptom bother following breast cancer diagnosis. *Annals of Behavioral Medicine* 50:370–384 DOI 10.1007/s12160-015-9763-7.
- **Bavojdan MR, Towhidi A, Rahmati A. 2011.** The relationship between mental health and general self-efficacy beliefs, coping strategies and locus of control in male drug users. *Addiction and Health* **3**:111–118.
- Blevins CE, Stephens RS, Walker DD, Roffman RA. 2014. Situational determinants of use and treatment outcomes in marijuana dependent adults. *Addictive Behaviors* 39:546–552 DOI 10.1016/j.addbeh.2013.10.031.

- **Bouchard G, Guillemette A, Landry-Léger N. 2004.** Situational and dispositional coping: an examination of their relation to personality, cognitive appraisals and psychological distress. *European Journal of Personality* **18**:221–238 DOI 10.1002/per.512.
- Buckner JD, Zvolensky MJ, Crosby RC, Wonderlich SA, Ecker AH, Richter A. 2015. Antecedents and consequences of cannabis use among racially diverse cannabis users: an analysis from Ecological Momentary Assessment. *Drug and Alcohol Dependence* 147:20–25 DOI 10.1016/j.drugalcdep.2014.12.022.
- **Cano-García FJ, Rodríguez-Franco L, Martínez JG. 2007.** Adaptación española del Inventario de Estrategias de afrontamiento. *Actas Españolas de Psiquiatría* **35**:29–39.
- Capella MM, Benaiges I, Adan A. 2015. Neuropsychological performance in polyconsumer men under treatment. Influence of age of onset of substance use. *Scientific Reports* 5:12038 DOI 10.1038/srep12038.
- Carver CS, Connor-Smith J. 2010. Personality and coping. *Annual Review of Psychology* 61:679–704 DOI 10.1146/annurev.psych.093008.100352.
- **Chauchard E, Septfons A, Chabrol H. 2013.** Motivations for cannabis cessation, coping and adaptation strategies, and perceived benefits: impact on cannabis use relapse and abstinence. *Encephale* **39**:385–392 DOI 10.1016/j.encep.2013.03.008.
- Coriale G, Bilotta E, Leone L, Cosimi F, Porrari R, De Rosa F, Ceccanti M. 2012. Avoidance coping strategies, alexithymia and alcohol abuse: a mediation analysis. *Addictive Behaviors* 37:1124–1129 DOI 10.1016/j.addbeh.2012.05.018.
- **Dermody SS, Cheong J, Manuck S. 2013.** An evaluation of the stress-negative affect model in explaining alcohol use: the role of components of negative affect and coping style. *Substance Use and Misuse* **48**:297–308 DOI 10.3109/10826084.2012.761713.
- **Dolan SL, Rohsenow DJ, Martin RA, Monti PM. 2013.** Urge-specific and lifestyle coping strategies of alcoholics: relationships of specific strategies to treatment outcome. *Drug and Alcohol Dependence* **128**:8–14 DOI 10.1016/j.drugalcdep.2012.07.010.
- Eddie D, Epstein EE, Cohn AM. 2015. Pathways to vulnerability for alcohol problem severity in a treatment-seeking sample. *Addictive Disorders and Their Treatment* 14:82–94 DOI 10.1097/ADT.0000000000045.
- Ehrenreic H, Rinn T, Kunert HJ, Moeller MR, Poser W, Schilling L, Gigerenzer G, Hoehe MR. 1999. Specific attentional dysfunction in adults following early start of cannabis use. *Psychopharmacology* 142:295–301 DOI 10.1007/s002130050892.
- Elofson J, Gongvatana W, Carey KB. 2013. Alcohol use and cerebral white matter compromise in adolescence. *Addictive Behaviors* **38**:2295–2305 DOI 10.1016/j.addbeh.2013.03.001.
- **European Monitoring Centre for Drugs and Drug Addiction. 2016.** European drug report: trends and development. *Available at http://www.emcdda.europa.eu/system/files/publications/2637/TDAT16001ENN.pdf/* (accessed on 15 January 2017).
- **First MB, Spitzer RL, Gibbon M, Williams JBW. 1999.** *Entrevista Clínica Estructurada para los Trastornos del Eje I del DSM-IV, versión clínica (SCID-I).* Barcelona: Masson.
- **Gálvez BP, Fernández LG. 2010.** Validación Española del Drug Abuse Screening Test (DAST-20 y DAST-10). *Health and Addictions* **10**:35–50.

- **Guy W. 1976.** *Early Clinical Drug Evaluation (ECDEU) assessment manual.* Rockville: National Institute of Mental Health.
- Hägele C, Friedel E, Kienast T, Kiefer F. 2014. How do we 'learn' addiction? Risk factors and mechanisms getting addicted to alcohol. *Neuropsychobiology* 70:67–76 DOI 10.1159/000364825.
- Hammond CJ, Mayes LC, Potenza MN. 2014. Neurobiology of adolescent substance use and addictive behaviors: prevention and treatment implications. *Adolescent Medicine: State of the Art Reviews* 25:15–32.
- Hruska B, Fallon W, Spoonster E, Sledjeski EM, Delahanty DL. 2011. Alcohol use disorder history moderates the relationship between avoidance coping and posttraumatic stress symptoms. *Psychology of Addictive Behaviors* 25:405–414 DOI 10.1037/a0022439.
- Huberty CJ. 2002. A history of effect sizes indices. *Educational and Psychological Measure*ment 62:227–240 DOI 10.1177/0013164402062002002.
- Hyman SM, Hong KA, Chaplin TM, Dabre Z, Comegys AD, Kimmerling A, Sinha R.
 2009. A stress-coping profile of opioid dependent individuals entering naltrexone treatment: a comparison with healthy controls. *Psychology of Addictive Behaviors* 23:613–619 DOI 10.1037/a0017324.
- Jockers-Scherübl MC, Wolf T, Radzei N, Schlattmann P, Rentzsch J, Gómez-Carrillo A, Kühl KP. 2007. Cannabis induces different cognitive changes in schizophrenic patients and in healthy controls. *Progress in Neuro-Psychopharmacology and Biological Psychiatry* 31:1054–1063 DOI 10.1016/j.pnpbp.2007.03.006.
- Jones DE, Greenberg M, Crowley M. 2015. Early social-emotional functioning and public health: the relationship between kindergarten social competence and future wellness. *American Journal of Public Health* 105:2283–2290 DOI 10.2105/AJPH.2015.302630.
- Kampman KM, Pettinati HM, Lynch KG, Whittingham T, Macfadden W, Dackis C, Tirado C, Oslin DW, Sparkman T, O'Brien CP. 2007. A double-blind, placebocontrolled pilot trial of quetiapine for the treatment of Type A and Type B alcoholism. *Journal of Clinical Psychopharmacology* 27:344–351 DOI 10.1097/JCP.0b013e3180ca86e5.
- Kendler KS, Ohlsson H, Sundquist K, Sundquist J. 2013. A latent class analysis of drug abuse in a national Swedish sample. *Psychological Medicine* **43**:2169–2178 DOI 10.1017/S0033291713000081.
- Kiluk B, Nich C, Carroll K. 2011. Relationship of cognitive function and the acquisition of coping skills in computer assisted treatment for substance use disorders. *Drug and Alcohol Dependence* 114:169–176 DOI 10.1016/j.drugalcdep.2010.09.019.
- Kirst M, Mecredy G, Borland T, Chaiton M. 2014. Predictors of substance use among young adults transitioning away from high school: a narrative review. *Substance Use and Misuse* **49**:1795–1807 DOI 10.3109/10826084.2014.933240.
- Kunert HJ, Derichs G, Irle E. 1996. Entwicklung von Aufmerksamkeitsfunktionen im Kindesalter: ergebnisse einer vorläufigen Normierung der computergestützten

Testbatterie zur Aufmerksamkeitsprüfung (TAP) an 9- bis 12jährigen Kindern. *Zeitschrift Neuropsychology* **7**:92–113.

- Lambe EK, Krimer LS, Goldman-Rakic PS. 2000. Differential postnatal development of catecholamine and serotonin inputs to identified neurons in prefrontal cortex of rhesus monkey. *Journal of Neuroscience* 20:8780–8787.
- Lazarus RH, Folkman S. 1984. *Stress, appraisal and coping.* New York: Springer Publishing Company.
- Marquez-Arrico JE, Benaiges I, Adan A. 2015. Strategies to cope with treatment in substance use disorder male patients with and without schizophrenia. *Psychiatry Research* 228:752–759 DOI 10.1016/j.psychres.2015.05.028.
- Mauro PM, Canham SL, Martins SS, Spira AP. 2015. Substance-use coping and selfrated health among US middle-aged and older adults. *Addictive Behavior* 42:96–100 DOI 10.1016/j.addbeh.2014.10.031.
- Nyamathi A, Hudson A, Greengold B, Slagle A, Marfisee M, Khalilifard F, Patton R. 2010. Predictors of substance use severity among homeless youth. *Journal of Child and Adolescent Psychiatric Nursing* 23:214–222 DOI 10.1016/j.addbeh.2009.04.002.
- **Pechtel P, Woodman A, Lyons-Ruth K. 2012.** Early maternal withdrawal and nonverbal childhood IQ as precursors for substance use disorder in young adulthood: results of a 20-year prospective study. *International Journal of Cognitive Therapy* **5**:316–329 DOI 10.1521/ijct.2012.5.3.316.
- **Pope HG, Gruber AJ, Hudson JI, Cohane G, Huestis MA, Yurgelun-Todd D. 2003.** Early-onset cannabis use and cognitive deficits: what is the nature of the association? *Drug and Alcohol Dependence* **69**:303–310 DOI 10.1016/S0376-8716(02)00334-4.
- Shaw P, Greenstein D, Lerch J, Clasen L, Lenroot R, Gogtay N, Evans A, Rapoport J, Giedd J. 2006. Intellectual ability and cortical development in children and adolescents. *Nature* **440**:676–679 DOI 10.1038/nature04513.
- Skinner EA. 1982. The drug abuse screening test. *Addictive Behaviors* 7:363–371 DOI 10.1016/0306-4603(82)90005-3.
- Skinner EA, Edge K, Altman J, Sherwood H. 2003. Searching for the structure of coping: a review and critique of category systems for classifying ways of coping. *Psychological Bulletin* 129:216–269 DOI 10.1037/0033-2909.129.2.216.
- Suijkerbuijk AWM, Van Gils PF, Greeven PGJ, De Wit GA. 2015. De kosteneffectiviteit van interventies gericht op verslaving aan alcohol en drugs. *Tijdschr Voor Psychiatrie* 57:498–507.
- Sundram S. 2006. Cannabis and neurodevelopment: implications for psychiatric disorders. *Human Psychopharmacology* 21:245–254 DOI 10.1002/hup.762.
- **Tobin DL, Holroyd KA, Reynolds RV, Wigal JK. 1989.** The hierarchical factor structure of the coping strategies inventory. *Cognitive Therapy and Research* **13**:343–361 DOI 10.1007/BF01173478.
- United Nations Office on Drugs and Crime. 2015. World drug report. Available at https://www.unodc.org/documents/wdr2015/World_Drug_Report_2015pdf/ (accessed on 15 January 2017).

- Walker R, Stephens R. 2014. Protective behavioral strategies mediate problem focused coping and alcohol use in college students. *Addictive Behaviors* **39**:1003–1037 DOI 10.1016/j.addbeh.2014.02.006.
- Wilson W, Mathew R, Turkington T, Hawk T, Coleman RE, Provenzale J. 2000. Brain morphological changes and early marijuana use: A magnetic resonance and positron emission tomography study. *Journal of Addictive Diseases* 19:1–22 DOI 10.1300/J069v19n01_01.
- Witkiewitz K, Marlatt GA. 2004. Relapse prevention for alcohol and drug problems: that was Zen, this is Tao. *American Psychologist* **59**:224–235 DOI 10.1037/0003-066X.59.4.224.
- Woodcock EA, Lundahl LH, Stoltman JJK. 2015. Greenwald M. K. Progression to regular heroin use: examination of patterns, predictors, and consequences. *Addictive Behaviors* 42:96–100 DOI 10.1016/j.addbeh.2015.02.014.
- Woodhead E, Cronkite RC, Moos RH, Timko C. 2014. Coping strategies predictive of adverse outcomes among community adults. *Journal of Clinical Psychology* 70:1183–1195 DOI 10.1002/jclp.21924.
- World Medical Association. 2013. WMA declaration of Helsinki: ethical principles for medical research involving human subjects. *Available at https://www.wma.net/policies-post/wma-declaration-of-helsinki-ethical-principles-for-medical-research-involving-human-subjects/* (accessed on 15 January 2017).