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Acute alcohol effects on set-shifting and its moderation by baseline individual differences: a latent variable analysis

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ABSTRACT

Aims To compare the acute effects of alcohol on set-shifting task performance (relative to sober baseline performance) during ascending and descending limb breath alcohol concentration (BrAC), as well as possible moderation of these effects by baseline individual differences. Design Shifting performance was tested during an initial baseline and a subsequent drinking session, during which participants were assigned randomly to one of three beverage conditions (alcohol, placebo or control) and one of two BrAC limb conditions [ascending and descending (A/D) or descending-only (D-only)]. Setting A human experimental laboratory on the University of Missouri campus in Columbia, MO, USA. Participants A total of 222 moderate-drinking adults (ages 21–30 years) recruited from Columbia, MO and tested between 2010 and 2013. **Measurements** The outcome measure was performance on set-shifting tasks under the different beverage and limb conditions. Shifting performance assessed at baseline was a key moderator. Findings Although performance improved across sessions, this improvement was reduced in the alcohol compared with no-alcohol groups (post-drink latent mean comparison across groups, all $Ps \le 0.05$), and this effect was more pronounced in individuals with lower pre-drink performance (comparison of pre- to post-drink path coefficients across groups, all $Ps \le 0.05$). In the alcohol group, performance was better on descending compared with ascending limb ($P \le 0.001$), but descending limb performance did not differ across the A/D and D-only groups. Conclusions Practising tasks before drinking moderates the acute effects of alcohol on the ability to switch between tasks. Greater impairment in shifting ability on descending compared with ascending breath alcohol concentration is not related to task practice.

Keywords Acute alcohol, baseline EF performance, executive functioning, latent-variable approach, limbs of the BrAC, set-shifting.

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INTRODUCTION

A large literature describes an important role for higherorder cognitive abilities in addiction, such as planning, problem-solving and flexibility [1,2]. This broad class of cognitive abilities is often referred to as 'executive functioning' (EF). Many EF-related abilities are known to be impaired under acute alcohol intoxication, both in the laboratory [3–5] and in naturalistic settings [6–8]. This is a major concern, given that impairment in the regulatory processes represented by these EF abilities can promote protracted alcohol and drug use despite their health risks and contribute to a range of acute alcohol-related problems. To date, most studies investigating acute effects of alcohol on EF have focused on abilities related to response inhibition or working memory; relatively few investigations have tested alcohol's effects on the ability to switch between tasks [9]. This ability, often termed 'shifting' [10], represents the ability to perform a new operation in the face of proactive interference (see [11]). Along with inhibition and working memory updating, shifting is considered one of the core EF abilities supporting self-regulatory control [10], and although these three aspects of EF share some common features, they are also theoretically and empirically distinct [10]. Addressing this imbalance in the literature on alcohol and EF is important, because although alcohol's effects on EF are extensive, they are far

from uniform [9]. The purpose of the current study was to provide the most thorough test to date of alcohol's effects on shifting.

Set-shifting paradigms involve a sequence of trials that require either transitioning to a new task set or maintaining the current one on successive trials. Outcome measures in set-shifting paradigms include the switch cost—the additional time [represented in response time (RT)] required to reconfigure successfully a new task set relative to maintaining the current one—and perseverative errors—responding to a stimulus according to a prior task set configuration that is inappropriate in the current one. Earlier studies showed that alcohol increases perseverative errors both in laboratory [4,12,13] and in naturalistic (bar) settings [7], but no previous studies have tested acute alcohol effects on switch costs.

Extant studies in this literature have suffered from several shortcomings, leading to uncertainty over their conclusions. First, in most studies alcohol's effects on shifting have been modeled using only a single behavioral task [9]. Moreover, the tasks used in most of those studies (e.g. Wisconsin Card Sort; Tower of London; Trails-making Test part B) are cognitively complex, meaning that they tap multiple EF abilities as well as other, non-EF processes (language; visuospatial), introducing variability in performance attributable to task stimuli but unrelated to shifting ability [10,14]. Previous work [15] demonstrates how alcohol's effects can vary according to such stimulus-driven factors. The current study overcame this problem by adopting a latent variable approach, in which three exemplar tasks selected to have different non-executive requirements but to tap the same underlying shifting ability were administered. The resulting latent variable represents an estimate of the latent variable of shifting ability with no measurement error [16].

Two additional shortcomings of the extant literature were addressed in this study. First, previous studies of alcohol's effects on shifting, indeed on cognitive functions more generally, have failed to investigate potential differences associated with ascending and descending breath alcohol concentration (BrAC). This issue is important, in that reduced impairment during descending relative to ascending BrAC has been reported for some EF abilities [17-22]; for reviews, see [23,24]. However, this apparent recovery on the descending limb of the BrAC curve has been observed mainly in within-subjects comparisons, where the same individuals are tested during both limbs [17-22]. This design makes it difficult to infer whether improvements during descending BrAC merely reflect practice effects [7,25], but also see [26]. Disentangling practice from acute tolerance effects [23] requires a design in which some participants complete EF tasks under both ascending and descending BrAC (A/D group), whereas others complete the tasks only under descending BrAC (D-only group). Better descendinglimb (DL) performance in the A/D group relative to the D-only group would provide evidence of a practice effect; equivalent DL performance in the two groups would suggest that acute tolerance occurs.

Finally, very few studies have considered the moderating influence of individual differences in sober (baseline) performance on the magnitude of alcohol's effects on EE Low EF ability represents a dual hazard for harmful drinking outcomes because not only does poor performance on EF tasks predict escalation of alcohol involvement [27–29] and risk-taking behaviors in youth [30], but individuals with poor EF ability often experience greater impairment from alcohol than their higher-EF peers [31,32]. The current study included a sober baseline testing session to permit modeling effects of individual differences in shifting ability.

The primary aim of this study was to provide the most comprehensive test to date of alcohol's acute effects on shifting, using a latent variable approach to isolate alcohol's effects on shifting from non-EF processes (see [14]). Based on the findings of earlier acute alcohol studies on EF, we focused on the following questions: (a) is post-drink performance in the alcohol group worse than post-drink performance in the no-alcohol group; (b) is performance on the DL significantly less impaired than performance on the ascending limb (AL) in the alcohol group, which might suggest acute tolerance or practice effects; and (c) does post-drink performance vary according to both pre-drink performance levels and beverage group, such that individuals with poorer baseline shifting ability experience greater impairment from alcohol than those with better baseline ability? We also wished to test (d) whether any ostensible performance recovery on the DL appears due to practice; a difference in DL performance between the A/D and D-only groups would suggest a practice effect.

METHOD

Design

The study consisted of a baseline session and a drinking session; during the drinking session participants consumed alcohol, a placebo beverage or a control beverage. This design permitted comparison of shifting performance (measured as a latent variable derived from performance in three set-shifting tasks) as a function of alcohol pharmacology (alcohol versus placebo and control) and alcohol expectancy (alcohol and placebo versus control) on the ascending and descending limb of the breath alcohol curve, as well as moderation of these effects by baseline individual differences. Prior to data collection, the statistical power analysis of the structural models was assessed using the Monte Carlo component of Mplus, and for each combination of three effect size magnitudes (low,

medium, high) for both alcohol and expectancy effects based on previously published estimates for the updating construct and assuming a sample size of 216 individuals. The power to detect small, medium and large effects for either construct were 0.19–0.20, 0.89–0.90 and 0.99, respectively.

their eligibility (see Supporting information for exclusion criteria). Participants were paid \$35 for completion of the first session and \$14/hour for the second session (and a \$10 bonus for completing both sessions). Demographic characteristics of the sample are given in Table 1.

Sample

Two-hundred and fifty-eight participants between the ages of 21 and 30 years were recruited from the Columbia, MO community for a study examining effects of alcohol on cognition. Interested individuals were screened for

Procedure

The baseline session started at 9:00 a.m. and took approximately 3-4 hours; drinking sessions, which took place 1-3 weeks (mean = 19.1 days) later, started between 12:00 and 1:00 p.m. and lasted approximately 4 hours.

Table 1 Demographic characteristics of study sample.

	Experimental group $(n = 222)$									
	Control		Placebo		Alcohol					
	D-only $n = 40$	A/D n = 40	$D-only \ n=35$	A/D n = 37	$D-only \ n=37$	A/D n = 33				
% Male	57.5	42.5	57.14	56.75	43.24	39.39				
Age (mean, SD)	22.57 (2.57)	22.3 (2.44)	22.71 (3.1)	22.73 (2.39)	22.97 (2.75)	23.24 (3.74)				
% Caucasian	97.44	87.18	94.12	94.59	97.3	96.77				
Smoking ^a	19/16/2/3	14/18/4/3 ^b	17/16/2/0	19/15/2/1	10/18/3/6	15/13/0/5				
(never/occasional/e	ex-/current smoker)									
Drinks per week (mean, SD)	6.54 (5.24)	9.52 (8.65)	7.16 (8.61)	5.79 (4.95)	7.97 (5.46)	6.19 (4.99)				

D-only: participants tested only on the descending limb of the breath alcohol concentration (BrAC) curve; A/D: participants tested on both the ascending and the descending limbs of the BrAC curve. a Units of measurement: total number of subjects; b in the control A/D limb group, one subject's smoking data were missing. SD = standard deviation.

Table 2 Means and standard deviations of switch costs in the alcohol, placebo and control groups.

	Experimental Group $(n = 222)$							
	Control		Placebo		Alcohol			
	D-only	A/D	D-only	A/D	D-only	A/D		
Baseline	n = 40	n = 40	n =35	n = 37	n = 37	n = 33		
Number-Letter	328.34 (193.92)	316.45 (170.73)	347.01 (182.37)	311.02 (186.41)	282.84 (182.81)	313.21 (159.34)		
Color-Shape	199.28 (192.03)	220.61 (175.86)	228.79 (116.00)	183.19 (140.52)	213.64 (177.60)	185.31 (137.62)		
Category-	160.17 (125.98)	132.33 (85.05)	129.55 (119.52)	149.26 (132.96)	152.49 (125.06)	194.22 (112.87)		
switch								
Ascending								
Number-Letter	_	226.67 (124.77)	_	221.29 (144.95)	_	303.05 (184.76)		
Color-Shape	_	201.16 (150.71)	_	142.92 (133.63)	_	254.46 (165.47)		
Category-	_	117.32 (96.14)	_	159.85 (139.29)	_	189.71 (127.62)		
Switch								
Descending								
Number-Letter	262.02 (158.50)	158.96 (95.41)	255.25 (155.58)	185.81 (126.28)	250.29 (187.57)	249.04 (143.73)		
Color-Shape	199.61 (178.40)	106.66 (83.17)	156.45 (125.65)	121.75 (133.62)	170.93 (120.53)	181.55 (146.25)		
Category- Switch	148.39 (101.77)	118.88 (96.52)	144.93 (107.59)	110.97 (94.11)	120.17 (105.31)	117.17 (107.22)		

A/D = completed tasks on the ascending and descending limb; D-only = completed tasks on the descending limb only.

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At the beginning of the baseline session, participants completed demographics and other self-report measures, followed by completion of the three set-shifting tasks as well as nine additional tasks assessing other EF abilities (not reported here). When participants returned to the laboratory for the drinking session they were assigned randomly to one of three beverage conditions by a research assistant using a computerized randomizer algorithm: a no-alcohol control beverage (n = 80), an active placebo beverage (n = 72; 0.04 g/kg ethanol) or an alcohol beverage (n = 70; 0.80 g/kg ethanol for men, 0.72 g/kg ethanol)for women) (for alcohol administration procedure, see Supporting information). Participants in the control condition were told that their drink contained no alcohol; those in the placebo and alcohol conditions were told that their drink contained 'a moderate amount of alcohol'.

To permit separate assessment of practice effects from acute alcohol tolerance on the DL, a missing-by-design method was employed [33,34] in which participants were assigned randomly to one of two task completion conditions. Participants in the A/D group completed the shifting tasks on both the AL and DL, whereas those in the D-only group completed the tasks only on the DL. All participants completed the set-shifting tasks in a predetermined order (category-switch, color-shape, number-letter), followed by two unrelated tasks. Following previous research [35], D-only participants watched episodes of a popular television sitcom (The Office) during ascending BrAC (in the alcohol condition, or until an equivalent amount of time had elapsed post-drinking in the placebo and control conditions). The order of the tasks was reversed on the DL so that each task would be completed at equivalent breath alcohol concentrations on the AL and DL.

Measures

Subjective effects of alcohol

At baseline and each post-drinking BrAC measurement, participants completed measures of self-reported stimulation, sedation and subjective intoxication. Stimulation and sedation were assessed using the Biphasic Alcohol Effects Scale (BAES) [36], and subjective intoxication ('How intoxicated do you feel right now?') was assessed using a 10-point scale (1 = not at all intoxicated, 10 = more intoxicated than I've ever been).

Set-shifting paradigms

Participants performed three cued set-shifting tasks (number-letter, color-shape and category-switch) used previously in the study by Friedman and colleagues [14]. For brevity, here we provide brief descriptions; complete details

are provided in the Supporting information. In all tasks, each trial was preceded by an informative cue indicating which one of the two subtasks should be performed on that trial. Subjects were required to switch between the two task-sets if the cue presented in the current trial differed from the cue presented in the previous trial. In the number-letter task [11], participants were presented with a number-letter or letter-number pair (e.g. 7G) and were expected to make an 'even/odd' or 'consonant/vowel' judgment depending on the pre-trial cue. In the color-shape task [37], participants were presented with a circle or a triangle in either red or green. They were expected to make a 'red/green' or 'circle/triangle' judgment depending on the cue. In the category switch task [38], participants were presented with a word and were asked either to make a 'living/non-living' or 'smaller/bigger than a soccer ball' judgment based on a symbol presented above the word. The dependent measure in each task was the switch cost, calculated as the difference between the average RTs of the trials that required a task switch and the average RTs of the trials in which no switch occurred (see Fig. 1). Each

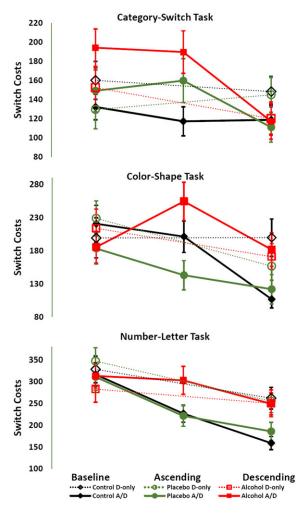


Figure I Mean switch costs in the alcohol, placebo and control groups. Vertical bars indicate standard errors

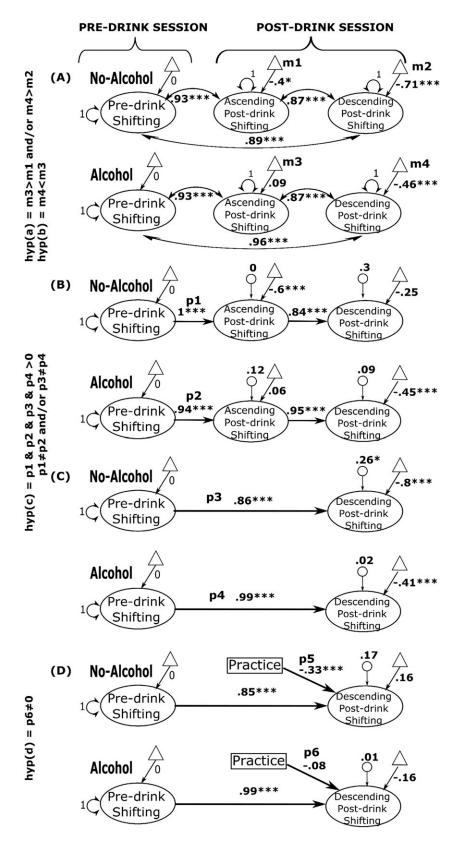


Figure 2 Standardized parameter estimates for the multi-group covariance, longitudinal factor and MIMIC models for alcohol and no-alcohol groups. For the parameter estimates: ****P < 0.001; *P < 0.05 (one-sample t-tests). Pre- and post-drink latent variables (ovals) are constructed from scores on three set-shifting tasks (category—switch, color—shape and number—letter). For simplicity, individual indicators are not presented. Triangles represent latent means (m), double-headed arrows on latent variables represent variances, double-headed arrows between latent variables represent covariances, single-headed arrows between latent variables represent path coefficients (p) and small circles with single-headed arrows represent disturbances; m: mean, p: path coefficient. Larger factor scores represent greater switch costs (i.e. worse performance)

task consisted of an equal number of switch and no-switch trials. The cue and the target were displayed on the screen until the participant responded, followed by a 350 ms response-to-cue interval. An auditory feedback ('beep') was presented if subjects responded incorrectly. Each target type and cue—target combination appeared equally often in each block. No more than four switch trials occurred in a row.

Statistical methods

For the analysis, a set of structural equation models (SEM) were applied because (i) within these models, latent factors (i.e. shifting ability) can be formed from multiple indicators (three set-shifting paradigms): in this way, measurement error can be reduced, resulting in increased reliability; and (ii) they allow us to estimate how these latent variables are affected by other (latent) variable indices across multiple groups.

Data preparation was performed as in previous research [14]. After exclusion of outliers/dropouts, model A-B and model C-D (for models, see Fig. 2) were estimated with 218 and 222 participants' data, respectively (see Supporting information for data preparation and exclusion/outliers criteria). SEMs were estimated with Mplus version 7.2 [39] by using multi-group covariance, longitudinal factor [40] and multiple-indicator-multiplecause (MIMIC) models [41]. In baseline models, all parameters were constrained to be equal across groups (strict invariance models). Next, parameters of interest were freed across groups and the difference likelihood ratio test $(\Delta \chi^2)$ was used to examine if nested models were significantly better than the strict invariance models as described in [42]. Models with incomplete data on the ascending limb were tested by using missing data analysis with full information maximum likelihood (FIML) estimation procedures

[43]. All other models were tested with maximum likelihood estimation procedures. In addition, root mean square error of approximation (RMSEA), comparative fit index (CFI), and Tucker–Lewis index (TLI) were used to judge the fit of individual models. All strict invariance models fitted the data relatively well (for a list of models estimated and statistics, see Tables 3 and 5).

The following specifications were the same in all SEMs presented here (Fig. 3 presents measurement model for model A2 as an example): the RT switch costs derived from the three set-shifting tasks were used to construct the unobserved latent variable, shifting, separately at pre-drink (baseline) and post-drink AL and DL. A dummy-coded alcohol variable, comparing the control and placebo groups (coded 0) with the alcohol group (coded 1), was created to test for pharmacological effects. For model identification purposes, on the pre-drink shifting factor the mean and variance were set to zero and 1, respectively, in both groups. Note that freeing the pre-drink shifting latent mean in the alcohol group resulted in a mean of zero for the latent variable in the alcohol group, confirming equal latent means in alcohol and no-alcohol groups at the predrink baseline. The covariances between the three indicators at pre- and post-drink were allowed, but constrained to be equal across alcohol and no-alcohol groups. The three factor loadings, intercepts and residual variances for the three indicators were estimated freely, but were constrained to be equal for pre- and post-drink shifting factors and across alcohol and no-alcohol groups. Also note that freeing factor loadings for the three indicators across time and groups worsened the fit significantly (model A3 in Table 3), corroborating that the same latent construct was measured before and after beverage administration. This provided a sufficient condition of equal units of measurement and origin of scale to test latent means across groups.

Table 3 Results from the models with baseline, ascending and descending limb data together (n = 218).

Models	χ^2	d.f.	RMSEA (95% CI)	CFI	TLI	$\Delta \chi^2$	P
Multi-group (alcohol/no-alcohol) covariance models							
A1: Strict invariance	125.320***	76	0.077 (0.052-0.101)	0.917	0.921		
A2: Asc. & desc. post-drink factor mean	114.351**	74	0.071 (0.044-0.095)	0.932	0.934	10.969	< 0.01
A3: Factor loadings ^a	198.863***	66	0.136 (114-158)	0.776	0.755		
Multi-group (alcohol/no-alcohol) longitudinal models	S						
B1: Strict invariance	130.153***	78	0.078 (0.054-0.101)	0.912	0.919		
B2: Asc. & desc. post-drink factor mean	110.406**	76	0.064 (0.035-0.090)	0.942	0.945	43.476	< 0.001
B3: Asc. & desc. post-drink factor var.	109.898**	75	0.065 (036-0.090)	0.941	0.943	23.793	< 0.001
B4: Base-to-asc. & ascto-desc. path coefficients	102.497*	73	0.061 (029-0.087)	0.95	0.951	33.841	< 0.001

Under the Models column, parameters are listed which are estimated freely across groups. Note that the free parameters in each successive model also include the parameters freed in previous steps. For example, in model B3, ascending and descending post-drink factor variance means that, in addition to the parameters freed earlier (i.e. factor means in model B2), the ascending and descending limb factor variances were also estimated freely across groups. Likelihood ratio difference tests $(\Delta\chi^2)$ were reported in comparison to the strict invariance model. Base = baseline; Asc. = ascending limb; Desc. = descending limb; var. = variance. * $^{*}P < 0.05$; * $^{*}P > 0.01$; * $^{*}P > 0.01$. * $^{*}P > 0.01$; * $^{*}P >$

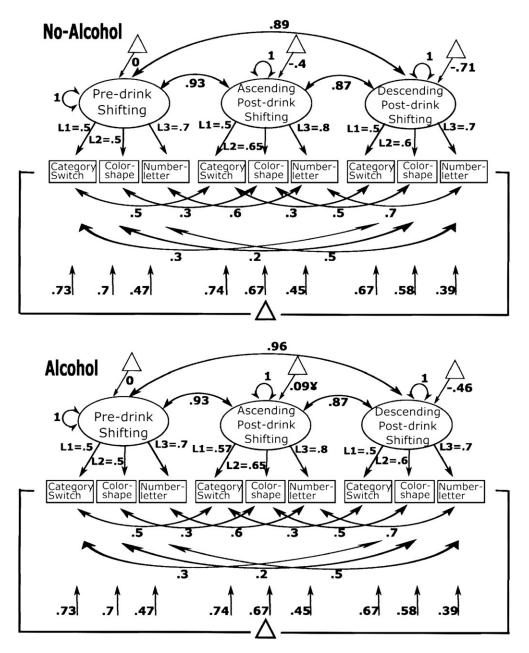


Figure 3 Standardized parameter estimates for the multi-group covariance model (model A2), presented as an example of measurement models that are simplified in Fig. 2. Values under vertical arrows pointing to measurement variables (category—switch, color—shape, etc.) represent residual variances, variance not explained by the latent variable. L1, L2, and L3 represent loadings constrained to be equal across time and groups. Triangles represent latent means, double-headed arrows on latent variables represent variances, double-headed arrows between latent variables represent covariances. Larger factor scores represent greater switch costs (i.e. worse performance). All parameters were significant at P < 0.05, except one (indicated by Y)

Note that we also tested whether performance on the DL was affected by the expectancy that alcohol was consumed, by using expectancy (0: control group, 1: placebo and alcohol groups) instead of alcohol dummy coding for grouping. Nested models testing expectancy effect were not significantly better than the strict invariance models (see Supporting information), suggesting that expectancy effects did not appear to influence performance and are therefore not discussed in the remainder of this paper.

MODEL RESULTS

To determine whether post-drink performance in the alcohol group was worse than in the no-alcohol group and whether performance on the AL in the alcohol group was significantly less impaired than performance on the DL, a covariance model was estimated in which the covariances among the three shifting latent factors were allowed, but constrained to be equal across the alcohol and no-alcohol groups. Testing hypotheses (a) and (b) requires comparison of group means

across time and groups by freeing the post-drink shifting latent means (Fig. 2A), which fitted the data significantly better than the baseline model (Table 3, Model A). Comparing the shifting latent mean across groups revealed that post-drink performance (both on the AL and DL) was worse (i.e. larger switch costs) in the alcohol group compared to the no-alcohol group (also see Table 4). Comparing the latent means across the limbs of the BrAC in the alcohol group revealed an improvement in performance from the AL to the DL in the alcohol group, suggesting an acute tolerance effect on the DL. These findings indicate support for hypotheses (a) and (b).

To address research question (c), whether post-drink performance varies according to pre-drink performance levels and beverage groups (specifically whether individuals with poorer baseline shifting ability experience greater impairment from alcohol than those with better baseline ability), a multi-group longitudinal factor model was estimated where the pre-drink component was regressed on the post-drink component. The post-drink latent means and variances, as well as the pre-drink-to-AL and AL-to-DL path coefficients were estimated freely (Fig. 2B). In the noalcohol group, pre-drink performance fully explained AL performance (path coefficient = 1), leaving no residual variance to be explained by the AL data (disturbance = 0). This is to be expected, given that no alcohol was administered to participants in this group. In the alcohol group, pre-drink performance also predicted AL performance but to a lesser extent than in the no-alcohol group; this

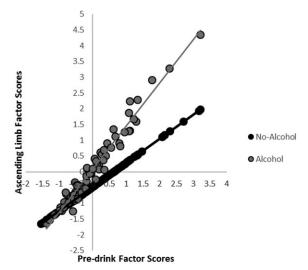
difference across groups was significant (Tables 3 and 4, model B). We then tested whether pre-drink performance predicted DL performance differently in the alcohol and no-alcohol groups. To do so, the same model was tested without the AL data (Fig. 2C; Table 5, model C). Comparing the path coefficients across groups revealed that pre-drink shifting ability explained the variance in DL performance more in the no-alcohol group than in the alcohol group (also see Fig. 4).

To address research question (d), whether performing the tasks on the AL affected performance on the DL in the alcohol group, a MIMIC model was estimated by including a practice dummy variable (1 = D-only group, 2 = A/D group) to the previous model and regressing on the DL component (Fig. 2D). Freely estimating the regression from the practice variable to the DL shifting variable resulted in a significant improvement over the baseline model (Table 5, model D). Significant negative path coefficients from practice to the DL post-drink shifting ability in the no-alcohol group (-0.33, P < 0.001) imply that the switch cost at DL post-drink was smaller (i.e. better shifting performance) in the A/D group compared to the D-only group; however, the analogous path for the alcohol group was not significant (-0.08, P = 0.23), indicating that practice did not improve DL performance in the alcohol group. Also, as shown in Table 6, these coefficients across groups were significantly different from each other. This finding is inconsistent with hypothesis (d).

Table 4 Standardized parameters from the models with baseline, ascending and descending limb data together in the no-alcohol and alcohol groups.

	Latent mean para	imeter estimates	Path coefficient estimates		
Models	No-alcohol	Alcohol	No-alcohol	Alcohol	
Multi-group (alcohol/no-alcohol) covariance factor models	Ascending				
A1: Strict invariance	-0.21	-0.21	_	_	
A2: Asc. & desc. post-drink factor mean	-0.4^{**b}	0.089^{b}	_	_	
	Descending				
A1: Strict invariance	-0.61***	-0.61***	_	_	
A2: Asc. & desc. post-drink factor mean	-0.71*** ^a	-0.46*** ^a	_	_	
A2: Asc. versus desc. limb	a	b	_	_	
Multi-group (alcohol/no-alcohol) Longitudinal factor models	Ascending		Base-to-ascend	ling	
B1: Strict invariance	-0.2	-0.2	0.98***	0.98***	
B2: Asc. & desc. post-drink factor mean	-0.48***b	0.09^{b}	1***	0.85***	
B4: Base-to-asc. & ascto-desc. path coefficient	-0.61*** ^b	0.06^{b}	1*** ^a	0.94***	
-	Descending		Ascending-to-descending		
B1: Strict invariance	-0.44***	-0.44***	0.91***	0.91***	
B2: Asc. & desc. post-drink factor mean	-0.35*	-0.51***	0.88***	0.91***	
B4: Base-to-asc. & ascto-desc. path coefficient	-0.25	-0.45***	0.84***	0.95***	

Under the Models column, parameters are listed which are estimated freely across groups. Note that the parameters listed in each successive model also include parameters that are freed in previous steps. Significance levels represent whether estimated parameters are significantly different than zero: $*P \le 0.05$; $**P \le 0.01$; $***P \le 0.001$. Significance levels for the *t*-test comparing parameter estimates for the no-alcohol versus alcohol groups and for the *t*-test comparing ascending versus descending limb factor means are indicated as, $^aP \le 0.05$; $^bP \le 0.01$. Base = baseline; Asc.= ascending limb; Desc.= descending limb.



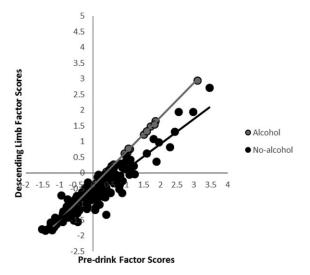


Figure 4 Post-drink shifting latent variable factor scores for the ascending limb (from model B4) and descending limb (from model C4) as a function of beverage group (alcohol versus no alcohol) and baseline task performance (pre-drink shifting latent variable factor scores). Larger factor scores represent greater switch costs (i.e. worse performance)

DISCUSSION

This experiment investigated the acute effect of alcohol on shifting. The study addressed limitations in the extant literature in several ways, including the use of a latent variable framework informed by multiple indicators of setshifting ability, examining whether individual differences in baseline shifting ability moderate alcohol effects on shifting performance and testing whether alcohol expectancy or practice (performing the tasks on both BrAC limbs) would explain ostensible acute tolerance on the DL of the BrAC curve. Several findings from this study advance understanding of alcohol's effects on shifting. First, the good fit of our model with three indicators of shifting provided evidence for a latent shifting factor tapped by the three tasks we administered. Moreover,

the SEM in which factor loadings for the three indicators were freed across time and groups worsened fit significantly (model A3 in Table 3), corroborating that the same latent construct was measured at baseline and after beverage administration.

In both alcohol and no-alcohol groups, performance was better at post-drink (lower switch costs and latent mean), due possibly to performing the tasks a second time (or third time for the A/D group). However, this improvement in performance was greater in the no-alcohol group than the alcohol group, suggesting that the pharmacological effects of alcohol limited the effects of practice. However, compared to the AL performance, DL performance was improved in the alcohol group, a finding similar to that seen in previous studies using within-subjects comparisons.

Additionally, baseline performance predicted post-drink performance differentially according to beverage condition: pre-drink performance explained AL performance more clearly in the no-alcohol group than in the alcohol group. Presumably, this occurred because alcohol interfered with shifting performance on the AL, reducing the extent of improvement from pre-drink levels in the alcohol group. More interestingly, the relative reduction in performance enhancement from pre- to post-drink in the alcohol condition was more pronounced in individuals with lower pre-drink performance (see Fig. 4). This could mean that individuals with low EF ability not only have relatively weak EF when sober, but they are also more susceptible to the detrimental effects of acute alcohol exposure. In the long term, these individuals might carry a greater risk for alcohol abuse and, ultimately, addiction.

The current study also investigated whether practice and/or expectancy effects influence shifting performance while BrAC is falling. There has been limited research on whether alcohol affects shifting differentially under ascending and descending BrAC. Previous studies have reported impaired EF performance on the AL and a recovery from this impairment on the DL, a finding that has been interpreted as a sign of acute tolerance. Possibly to eliminate individual variability in response to acute alcohol and increase statistical power, most studies examining such limb effects have utilized within-subjects designs in which participants were tested both under ascending and descending BrAC [17–22]. Conversely, one study compared participants tested on the AL to those tested on the DL of BrAC and reported no difference in shifting performance [7]. In the current study, we scrutinized this issue by (a) testing whether performing the tasks on the AL improves DL performance (i.e. a within-subjects comparison) and (b) testing whether DL performance differs according to whether or not the tasks were performed on the AL (i.e. a between-subjects comparison across the A/D and D-only groups). Results showed a general performance improvement on the DL compared to the AL. Moreover, performing

Table 5 Results from the models with baseline and descending limb data together (n = 222).

Models	χ^2	d.f.	RMSEA (95% CI)	CFI	TLI	$\Delta\chi^2$	P	
Multi-group (alcohol/no-alcohol) longitudinal factor models								
C1: Strict invariance	49.801*	36	0.059 (0-0.096)	0.960	0.966	_	_	
C2: Desc. post-drink factor mean	44.840	35	0.050 (0-0.090)	0.971	0.975	5.0586	< 0.025	
C3: Desc. post-drink factor var.	45.097	34	0.054 (0-0.093)	0.968	0.971	3.8219	> 0.1	
C4: Pre- to post-path coefficient	39.656	33	0.043 (0-0.085)	0.981	0.982	8.1597	< 0.05	
Multi-group (alcohol/no-alcohol) MIMIC mo	dels (with prac	tice varia	ıble)					
D1: Strict invariance	62.951*	47	0.055 (0-0.088)	0.959	0.963	_	_	
D2: Limb path coefficient	55.580	46	0.043 (0-0.080)	0.975	0.978	7.001	< 0.01	
D3: Desc. post-drink factor mean & var.	53.836	44	0.045 (0-0.082)	0.975	0.976	8.233	< 0.05	
D4: Pre- on post-path coefficient	46.635	43	0.028 (0-0.071)	0.991	0.991	14.692	< 0.01	

Names of parameters in the Models column were estimated freely across groups in each successive model are listed. Note that free parameters in each successive model also include the parameters that are freed in previous steps; e.g. in model C4. pre- to post-path coefficient means that, in addition to the parameters freed earlier (i.e. post-drink factor mean and variance in model C2 and C3), the path coefficient from pre- to post-drink latent variable was also estimated freely across groups. Likelihood ratio tests $(\Delta\chi^2)$ were reported in comparison to the strict invariance models. Desc. = descending limb; var. = variance. *P < 0.05; RMSEA = root mean square error of approximation; CI = confidence interval; CFI = comparative fit index; TLI = Tucker–Lewis index; MIMIC = multiple-indicator–multiple-cause.

Table 6 Standardized parameters from the models with baseline and descending limb data together.

	Parameter estimates								
	Latent mean		Latent variance		Pre–post path		Practice–post path		
Models	No-alcohol	Alcohol	No-alcohol	Alcohol	No-alcohol	Alcohol	No-alcohol	Alcohol	
Multi-group (alcohol/no-alcohol) los	ngitudinal facto	r models							
C1: Strict invariance	-0.660***	-0.660***	0.166	0.166	0.913***	0.913***	_	_	
C2: Des. post-drink factor mean	-0.751^{***a}	-0.476***a	0.174	0.174	0.909***	0.909***	_	_	
C3: Des. post-drink factor variance	-0.749*** ^a	-0.477*** ^a	0.176	0.166	0.908***	0.913***	_	_	
C4: Pre- to post-path coefficient	-0.808***b	-0.396*** ^b	0.260*	0.019	0.860***a	0.988*** ^a	_	_	
Multi-group (alcohol/no-alcohol) M	IMIC models (with practice v	ariable)						
D1: Strict invariance	0.029	-0.029	0.115	0.115	0.912***	0.912***	-0.230***	-0.230***	
D2: Limb path coefficient	0.009	0.009	0.121	0.126	0.902***	0.921***	-0.257***b	-0.159**b	
D3: Des. post-drink factor mean & var.	0.143	-0.265	0.097	0.181	0.901***	0.902***	-0.602*** ^a	-0.150 ^a	
D4: Pre- on post-path coefficient	0.162	-0.157	0.166	0.010	0.853***b	0.992*** ^b	-0.326***a	-0.081 ^a	

Names of parameters estimated freely across groups in each successive model are listed under the Models column. These parameters are successive in that they include parameters freed in previous steps; e.g. in model C4, pre- to post-path coefficient means that, in addition to the parameters freed earlier (i.e. post-drink factor mean and variance in model C2 and C3), the path coefficient from pre- to post-drink latent variable was also estimated freely across groups. Significance levels represent whether estimated parameters are significantly different from zero: $^*P \le 0.05$; $^*P \le 0.01$; $^*P \le 0.01$. Significance levels for the t-tests comparing parameter estimates for the no-alcohol versus alcohol groups are indicated as: $^*P \le 0.05$; $^*P \le 0.01$. Desc. = descending limb; var. = variance; pre-post path = pre- to post-drink (descending limb) path coefficient: practice—post path = practice to post-drink (descending limb) path coefficient.

the tasks on the AL improved DL performance, but only in the no-alcohol group. A lack of difference between the performance of the A/D and D-only groups in the alcohol condition suggests that the acute tolerance observed on the DL cannot be explained as merely a practice effect. Regarding the expectancy factor, the findings were inconclusive as to whether or not expectancy influenced EF performance.

In summary, our results indicate that individuals low in a key aspect of EF, switching, might experience loss of executive control after initiation of a drinking episode. However, whether individuals low or high in EF would be more prone to initiate a drinking episode is a question for future research. Limitations of the current study should be noted, however. First, although this sample is large by the standards of alcohol challenge studies, it was not large enough to permit estimation of numerous parameters in our models. Future studies with larger samples could test this possibility. Secondly, the alcohol effect on shifting reported here is related to a broad, latent shifting construct and may not generalize to specific tasks.

However, our use of a latent variable approach to capture an underlying shifting ability measured during both a sober baseline and a subsequent drinking session represents a significant advance over typical designs utilizing only a single shifting task and measuring shifting at only one (drinking) session, and stands as the most comprehensive examination to date of the acute effect of alcohol on setshifting performance.

Declaration of interests

None.

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References

- Crews F. T., Boettiger C. A. Impulsivity, frontal lobes and risk for addiction. *Pharmacol Biochem Behav* 2009; 93: 237–47.
- Moselhy H., Georgiou G., Kahn A. Frontal lobe changes in alcoholism: a review of the literature. *Alcohol Alcohol* 2001; 36: 357–68.
- Dry M. J., Burns N. R., Nettelbeck T., Farquharson A. L., White J. M. Dose-related effects of alcohol on cognitive functioning. PLOS ONE 2012; 7: e50977.
- Casbon T. S., Curtin J. J., Lang A. R., Patrick C. J. Deleterious effects of alcohol intoxication: diminished cognitive control and its behavioral consequences. *J Abnorm Psychol* 2003; 112: 476–87.
- McCarthy D. M., Niculete M. E., Treloar H. R., Morris D. H., Bartholow B. D. Acute alcohol effects on impulsivity: associations with drinking and driving behavior. *Addiction* 2012; 107: 2109–14.
- Day A., Celio M., Lisman S., Johansen G., Spear L. Acute and chronic effects of alcohol on trail making test performance among underage drinkers in a field setting. *J Stud Alcohol Drugs* 2013; 74: 635–41.
- Lyvers M., Tobias-Webb J. Effects of acute alcohol consumption on executive cognitive functioning in naturalistic settings. Addict Behav 2010; 35: 1021–8.
- 8. Scholey A., Benson S., Neale C., Owen L., Tiplady B. Neurocognitive and mood effects of alcohol in a naturalistic setting. *Hum Psychopharmacol Clin Exp* 2012; 27: 514–16.
- Day A., Kahler C., Ahern D., Clark U. Executive functioning in alcohol use studies: a brief review of findings and challenges in assessment. Curr Drug Abuse Rev 2015; 8: 26–40.
- Miyake A., Friedman N. P., Emerson M. J., Witzki A. H., Howerter A., Wager T. D. The unity and diversity of executive functions and their contributions to complex 'Frontal Lobe' tasks: a latent variable analysis. *Cogn Psychol* 2000; 41: 49–100.
- 11. Rogers R., Monsell S. Costs of a predictible switch between simple cognitive tasks. *J Exp Psychol* 1995; **124**: 207–31.
- 12. Guillot C., Fanning J., Bullock J., McCloskey M., Berman M. Effects of alcohol on tests of executive functioning in men

- and women: a dose response examination. *Exp Clin Psychopharmacol* 2010; **18**: 409–17.
- Lyvers M., Maltzman I. Selective effects of alcohol on Wisconsin card sorting test performance. Br J Addict 1991; 86: 399–407.
- Friedman N., Miyake A., Young S., Defries J., Corley R., Hewitt J. Individual differences in executive functions are almost entirely genetic in origin. J Exp Psychol Gen 2008; 137: 201–25.
- Hernández O. H., Vogel-Sprott M., Huchín-Ramirez T. C., Aké-Estrada F. Acute dose of alcohol affects cognitive components of reaction time to an omitted stimulus: differences among sensory systems. *Psychopharmacology (Berl)* 2006; 184: 75–81.
- Loehlin J. C. Latent Variable Models: An Introduction to Factor, Path, and Structural Equation Analysis, 4th edn. Mahwah, NJ: Erlbaum; 2011.
- 17. Gilbertson R., Ceballos N. A., Prather R., Nixon S. J. Effects of acute alcohol consumption in older and younger adults: perceived impairment versus psychomotor performance. *J Stud Alcohol Drugs* 2009; 70: 242–52.
- Söderlund H., Parker E. S., Schwartz B. L., Tulving E. Memory encoding and retrieval on the ascending and descending limbs of the blood alcohol concentration curve. *Psychophar-macology (Berl)* 2005; 182: 305–17.
- Schweizer T. A., Vogel-Sprott M., Danckert J., Roy E. A., Skakum A., Broderick C. E. Neuropsychological profile of acute alcohol intoxication during ascending and descending blood alcohol concentrations. *Neuropsychopharmacology* 2006; 31: 1301–9.
- Schreckenberger M., Amberg R., Scheurich A., Lochmann M., Tichy W., Klega A. et al. Acute alcohol effects on neuronal and attentional processing: striatal reward system and inhibitory sensory interactions under acute ethanol challenge. Neuropsychopharmacology 2004; 29: 1527–37.
- Hiltunen A. Acute alcohol tolerance in cognitive and psychomotor performance: influence of the alcohol dose and prior alcohol experience. *Alcohol* 1997; 14: 125–30.
- Fillmore M. T., Marczinski C. A., Bowman A. M. Acute tolerance to alcohol effects on inhibitory and activational mechanisms of behavioral control. *J Stud Alcohol* 2005; 66: 663–72.
- 23. Vogel-Sprott M. Is behavioral tolerance learned? *Alcohol Health Res World* 1997; 21: 161–8.
- Schweizer T. A., Vogel-Sprott M. Alcohol-impaired speed and accuracy of cognitive functions: a review of acute tolerance and recovery of cognitive performance. *Exp Clin Psychopharmacol* 2008; 16: 240–50.
- Pihl R. O., Paylan S. S., Gentes-Hawn A., Hoaken P. N. S. Alcohol affects executive cognitive functioning differentially on the ascending versus descending limb of the blood alcohol concentration curve. *Alcohol Clin Exp Res* 2003; 27: 773–9.
- Fillmore M. T., Ostling E. W., Martin C. A., Kelly T. H. Acute effects of alcohol on inhibitory control and information processing in high and low sensation-seekers. *Drug Alcohol Depend* 2009; 100: 91–9.
- Peeters M., Janssen T., Monshouwer K., Boendemaker W., Pronk T., Wiers R., et al. Weaknesses in executive functioning predict the initiating of adolescents' alcohol use. *Dev Cogn Neurosci* 2015; 16: 139–46.
- Nigg J. T., Wong M. M., Martel M. M., Jester J. M., Puttler L. I., Glass J. M., et al. Poor response inhibition as a predictor of problem drinking and illicit drug use in adolescents at risk for alcoholism and other substance use disorders. J Am Acad Child Adolesc Psychiatry 2006; 45: 468–75.

- Khurana A., Romer D., Betancourt L., Brodsky N., Giannetta J., Hurt H. Working memory ability predicts trajectories of early alcohol use in adolescents: the mediational role of impulsivity. Addiction 2013; 108: 506–15.
- Pharo H., Sim C., Graham M., Gross J., Hayne H. Risky business: executive function, personality, and reckless behavior during adolescence and emerging adulthood. *Behav Neurosci* 2011; 125: 970–8.
- Curtin J. J., Fairchild B. A. Alcohol and cognitive control: implications for regulation of behavior during response conflict. *J Abnorm Psychol* 2003; 112: 424–36.
- Finn P. R., Justus A., Mazas C., Steinmetz J. E. Working memory, executive processes and the effects of alcohol on Go/ No-Go learning: testing a model of behavioral regulation and impulsivity. *Psychopharmacology (Berl)* 1999; 146: 465–72.
- Rhemtulla M., Little T. D. Tools of the trade: planned missing data designs for research in cognitive development. *J Cogn Dev* 2012; 13: 425–38.
- Graham J. W., Taylor B. J., Olchowski A. E., Cumsille P. E. Planned missing data designs in psychological research. *Psychol Methods* 2006; 11: 323–43.
- Roehrich L., Goldman M. S. Implicit priming of alcohol expectancy memory processes and subsequent drinking behavior. *Exp Clin Psychopharmacol* 1995; 3: 402–10.
- Martin C. S., Earleywine M., Musty R. E., Perrine M. W., Swift R. M. Development and validation of the biphasic alcohol effects scale. *Alcohol Clin Exp Res* 1993; 17: 140–6.
- Miyake A., Emerson M. J., Padilla E., Ahn J. Inner speech as a retrieval aid for task goals: the effects of cue type and articulatory suppression in the random task cuing paradigm. *Acta Psychol (Amst)* 2004; 115: 123–42.
- Mayr U., Kliegl R. Task-set switching and long-term memory retrieval. J Exp Psychol Learn Mem Cogn 2000; 26: 1124–40.
- 39. Muthén LK, Muthén BO. Mplus Version 7.2. 2013.

- Cudeck R., MacCallum R. C. Factor Analysis at 100: Historical Developments and Future Directions. Mahwah, NJ: Lawrence Erlbaum Associates; 2003.
- Jöreskog K., Goldberger A. Estimation of a model with multiple indicators and multiple causes of a single latent variable. *J Am Stat Assoc* 1975; 70: 631–9.
- Byrne B. M. Structural Equation Modeling with Mplus: Basic concepts, applications, and programming. New York, NY: Routledge; 2012.
- Enders C. K., Bandalos D. L. The relative performance of full Information maximum likelihood estimation for missing data in structural equation models. *Struct Equ Model* 2001; 8: 430–57.

Supporting Information

Additional Supporting Information may be found in the online version of this article at the publisher's web-site:

In the online supplementary materials, Table S1 and Table S2 present model results for the Expectancy and No-expectancy groups.

Table S1 Results from the models with baseline, ascending and descending limb data together.

Table S2 Standardized parameters from the models with baseline, ascending and descending limb data together in the no-expectancy and expectancy groups.

Table S3 Means (and standard deviations) of breath alcohol concentration (BrAC), subjective intoxication ratings and Biphasic Alcohol Effects Scale (BAES) stimulation and sedation ratings throughout the experiment as a function of experimental group.