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# Alcohol and drug use among adolescents - and the cooccurrence of mental health problems. Ung@hordaland, a population-based study.

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## ABSTRACT

**Objectives:** The use of alcohol and drugs is prevalent among adolescents, but little is known about the association between debut of alcohol and drug use and concurrent mental health. The aim of the study was to investigate the cross-sectional association between debut of any alcohol or drug use and alcohol- and drug-related problems, and mental health. We also wanted to examine potential interactions between gender and age, and alcohol- and drug-related exposures.

Design: Cross-sectional study.

Setting: Population-based sample of Norwegian adolescents.

**Participants:** Data stem from the large population-based ung@hordaland study (N=9,203), where all adolescents aged 16-18 years living in Hordaland county (Norway) were invited to participate. The main exposures were debut of alcohol and drug use, alcohol consumption, and presence of alcohol and drug problems as measured by CRAFFT.

Outcomes: The outcomes were self-reported symptoms of anxiety, depression, inattention and hyperactivity.

**Results:** Both debut of alcohol and drug use were associated with symptoms of depression, inattention and hyperactivity (crude ORs 1.69-2.38, Ps<0.001), while only debut of drug use was associated with increased symptoms of anxiety (OR=1.33, CI95%1.05-1.68, P=0.017). Alcohol- and drug-related problems as measured by CRAFFT were associated with all mental health problems (crude ORs 1.68-3.24, Ps<0.001). There was little evidence of any substantial age- or gender confounding on the estimated associations between alcohol- and drug-related measures and the mental health outcomes.

**Conclusion:** Early debut of alcohol and drug use and drug problems are consistently associated with more symptoms of mental health problems, indicating that these factors are an important general indicator of mental health in adolescence.

# STRENGTHS AND LIMITATIONS OF THIS STUDY

- This study employed a large well-defined population-based sample of adolescents.
- The data employed in this study is from a recent data collection.
- This study included several different measures of alcohol- and drug use, as well as measures of both internalizing and externalizing problems.
- The cross-sectional design of this study precluded any causal inference.
- This study only had a limited age-range, a wider range would enable a more thorough investigation of age-related effects of alcohol and drug use.

## BACKGROUND

The disease burden attributed to alcohol and illicit drug use is considerable worldwide.[1, 2] Alcohol and illicit drug use tend to begin and escalate during adolescence and early adulthood.[3, 4] Early life alcohol and drug patterns are predictive of later use and abuse,[5] and for alcohol it has been shown that a 10% time-delay in debut would lead to a 35% decrease in the subsequent expected alcohol consumption .[6]

Drug use and abuse have been related to a range of mental health problems, most consistently to externalizing problems such as those associated with conduct disorder (CD) and attention-deficit-hyperactivity disorder (ADHD), [7] while the increased risk for anxiety disorders are lower. Also of note, several previous studies in adult populations have found that there is a J- or U-shaped relationship between alcohol consumption and mental health outcomes,[8-12] where both abstainers and high-level alcohol consumers have increased risk for mental health problems. Several explanations of this curvilinear relationship have been suggested: moderate alcohol consumers may directly benefit from a protective effect on mental health from alcohol, or their drinking habits may be a mere reflection of their psychological wellbeing.[9] Also, in cultures where alcohol use is the norm, moderate consumers may be more well-adjusted than individuals at the extremes of the consumption spectrum.[13] To date, few studies have investigated the relationship between the full spectrum of alcohol consumption (from non-debut to high-level alcohol consumption) and symptoms of mental health problems in a population-based sample of adolescents

Boys have an earlier debut of both alcohol consumption and intoxication according to most studies,[3] while the gender-pattern for onset of drug use is more unclear.[14-19] The established gender differences with males using more alcohol and drugs seems, however, to be closing [20] with recent studies not finding a higher rate of drug use among males.[21] If the relation between alcohol and drug use and mental health is gender specific is less clear. A lack of gender difference has been documented in population-based studies where the use of alcohol and drugs is associated with worse mental health.[22-25] In contrast, some studies have reported a modifying effect of gender and age in relation to the association between alcohol use and mental health.[23, 26, 27] For instance, a recent study investigating the association between alcohol intoxications and mental health problems among 13-19-year-olds found that high level alcohol consumption was associated with conduct and attention problems for both genders, but only girls who reported frequent intoxications reported more symptoms

of anxiety and depression.[23] On the other hand, Verdurmen and colleagues reported a modifying effect of age on the association between alcohol use and mental health, but found no evidence of any modifying effects of gender.[26] The association between alcohol use and mental health is, however, not quite consistent, as a Dutch study did not confirm the expected relation between binge drinking and mental health problems among 16-18years-olds in a population based study.[28]

Less is known whether the associations between the drug use and mental health is gender-specific, but there are some indications that co-occurrence of other mental health problems are higher for girls than boys.[7] However, in a sample of late adolescent urban youth gender did not have a moderating effect on the relationship between substance abuse and mental health, and a Finnish study did not support a gender specific association between depression and substance use.[21] These studies are rather small, and the results may have been hampered by lack of statistical power. It is also uncertain if the co-occurrence is specifically related to alcohol or drug dependence or also evident for mere alcohol or drug use.[7] Overall, there is still a need for further investigation of the association between early alcohol and drug use and mental health in a general population of adolescents.[29]

To that end, the aim of the present study was to

- Describe the debut of alcohol and drug use, alcohol use patterns and potential drug problems among Norwegian adolescents using data from a recent population-based study.
- ii) To investigate the cross-sectional relationship between a) alcohol and illicit drug use and b) alcohol and drug problems and mental health among adolescents. We also investigated any interactions with age and gender on the associations of interest, as well as the association between the full spectrum of alcohol consumption and binge drinking frequency in relation to mental health.

# METHODS

#### **Study population**

This study employed information from the ung@hordaland-survey ("youth@hordaland") of adolescents in the county of Hordaland in western Norway. All adolescent born 1993-1995 living in Hordaland were invited to participate. Ung@hordaland is a population-based study, and carried out in close collaboration with the county council administration. The majority of participants responded to the questionnaires during school hours. The general aim of the ung@hordaland-survey was to assess mental health, lifestyle, school performance and health-

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service use in adolescents, with a special emphasis on the prevalence of mental health problems. The data was collected during January-February 2012. Adolescents in upper secondary education received information per email, and one school hour was allocated for them to complete the questionnaire at school. Those not in school received information by mail to their home addresses. The questionnaire was web-based and covered a broad range of mental health issues, daily life functioning, use of health and social services, demographic background variables, and a request for permission to obtain school data, and to link the information with national registries and parental questionnaires. Uni Health and Hordaland County Council are collaborating partners in conducting the study. The study was approved by The Regional Committee for Medical Research Ethics in Western Norway. After complete description of the study to the subjects, written informed consent was obtained.

The current study is based on the first version of data files released in May 2012, with a total of 10,220 participants. Based on list-wise deletion of those with missing information on both exposures and outcomes, n=1,017 were excluded, leaving 9,203 in the final study sample. Those with missing information on the exposures and the outcomes, were slightly younger than those with valid information on the included variables (mean difference 0.12 years, p<.001) and more likely to be males (61.6% versus 45.7% in the analysed sample, p<.001).

# Identification of alcohol problems and illicit drug use

Self-reported debut, frequency and amount of alcohol consumption and illicit drug use were included as main exposure variables. We included a binary measure of alcohol debut "Have you ever tried alcohol?" (Yes/No), and illicit drug use debut: "Have you ever tried hash, marihuana or other narcotic substances?" (Yes/No). Out of the total study sample, n=11 individuals did not answer the question on illicit drug use, and they were recoded to the "No" category. Alcohol use was measured using the self-reported units of beer, eider, wine, spirits and illegally distilled spirits usually consumed during the last 14 days. Based on this information about consumption and alcohol debut, a summed variable including the following gender-specific distribution was constructed: "Never tried", "Non-consumption" (if reported consumption was "0"), "0.1-19.9<sup>th</sup>", "20.0-39.9<sup>th</sup>", "40.0-59.9<sup>th</sup>", "60.0-79.9<sup>th</sup>", "80.0-89.9<sup>th</sup>", "90.0-100<sup>th</sup>". Excessive alcohol consumption was defined as above the 90<sup>th</sup> percentile sum score.[11, 30] Frequent binge drinking was defined as drinking so much that one was clearly intoxicated (drunk)?", with five categories ranging from "No, never" to "Yes, more than 10 times". There were n=147 missing responses on the binge drinking item, leaving a sample of N=9,056 for this measure. In

addition, potential alcohol and drug problems were identified employing the six-item inventory CRAFFT.[31] CRAFFT was specifically designed to identify alcohol and illicit drug problems among adolescents,[31] and have been shown to have acceptable sensitivity and specificity at a cut-off of >=2 in international studies.[32, 33] In relation to the ung@hordaland sample, a linear relationship between CRAFFT-score and excessive alcohol consumption, binge drinking and debut of illicit drug use has previously been reported.[33]

## Mental health problems

Depression was assessed using the short version of the Mood and Feelings Questionnaire (SMFQ).[34] The SMFQ comprises 13 items assessing depressive symptoms rated on a 3-point scale. The wordings of the response categories in the Norwegian translation equal the original categories of "Not true", "Sometimes true" and "True". High internal consistency between the items and a strong unidimensionality have been shown in population-based studies,[35] and was recently confirmed in a study based on the sample included in the present study.[36] For the purposes of the current study, depression was defined as a score above the 90<sup>th</sup> percentile of the Total SMFQ-score. It should be noted that term depression as used in the current study does not imply existence of a clinical diagnosis, such as MDD. Also, being a relatively brief self-report questionnaire, the SFMQ does not differentiate between different types of depressive disorders/conditions. The Cronbach's alpha of the SMFQ in the ung@hordaland-sample was 0.91.

Symptoms of anxiety was identified using the five-item inventory SCARED, which is short-form of the 41-item version screening inventory for anxiety disorders.[37] The short-form of the SCARED has showed similar psychometric properties to the full version.

Symptoms of inattention and hyperactivity were measured using sub-scores from the official Norwegian translation of the Adult ADHD Self-report Scale (ASRS),[38] originally constructed for use in adults, but recently validated among adolescents.[39] ASRS is an 18-item self-report scale, comprising 9 items on a hyperactivity-impulsivity subscale and 9 items on an inattention-subscale. The responses are given on a 5-point scale (ranging from "Never" to "Very often").

## Statistical analysis

Five different alcohol and drug-related binary variables were constructed as main exposures: Ever consumed alcohol (Yes/No), ever tried illicit drugs (Yes/No), those reporting 90<sup>th</sup> gender-specific percentile alcohol consumption (Yes/No), binge drinking >10 times (Yes/No) and CRAFFT-caseness according to the ≥2 cutoff

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(Yes/No). For outcome variables the mental health questionnaires were dichotomised at the 90<sup>th</sup> percentile, identifying those with high sum scores on the ASRS subscales of inattention and hyperactivity-impulsivity, the SMFQ scale of depression and the SCARED scale of anxiety.

First, the sample was described in relation to exposures stratified for age and gender. Second, odds ratios were computed using simple logistic regression models for the associations between exposures and outcomes, as well as logistic regression models adjusted for age and gender. In addition, we investigated the potential two-way interaction with age and gender for the alcohol- and drug-related exposures in relation to mental health problems. The presence of significant interactions was assessed using likelihood-ratio tests (Wald), and when significant interactions were identified the associations of interest were presented stratified for age or gender. In secondary analyses, we investigated the association between the spectrum of alcohol consumption ("Never tried alcohol", "Non-consumption", "0.1-19.9<sup>thac</sup>, "20.0-39.9<sup>th</sup>", "40.0-59.9<sup>th</sup>", "60.0-79.9<sup>th</sup>", "80.0-89.9<sup>th</sup>", "90.0-100<sup>th</sup>") and binge drinking ("Never tried alcohol", "Never", "Once", "2-3 times", 4-10 times" and ">10 times") categories separately for all mental health variables, employing age- and gender-adjusted logistic regression models with "Never tried" as reference group. All analyses were performed using Stata 12.0.[40]

# RESULTS

The final sample consisted of N=9,203 participants, where 54.3% were female and the mean age was 17.9 (SD 0.8) years. Table 1 and Figure 1 show the use of alcohol and illicit drugs stratified by age and gender. A total of 20.7% of the girls and 27.1% of the boys had *not* tried alcohol, whereas 8.8% of the girls and 12.5% of the boys *had* tried any illicit drug. Also, ~22% percent of the sample was CRAFFT-cases, indicating some alcohol- or drug-related problem. There was a significant age-trend for all exposures for both genders, where increasing age was associated with increased proportion of participants having tried alcohol or illicit drugs, as well as reporting alcohol- and drug-related problems (p-values ranging from: 0.002 to <0.001).

Table 2 depicts the crude and age- and gender-adjusted associations between having tried alcohol, illicit drugs or reporting alcohol- and drug-related problems, in relation to mental health problems. Having ever consumed alcohol was associated with increased odds of all mental health problems in both crude and age- and gender-adjusted models, except for symptoms of anxiety (crude ORs ranging from 1.13 to 2.36). Having ever tried illicit drugs was associated with increased odds of all mental health problems in both crude and age- and gender-adjusted models (crude ORs ranging from 1.33 to 2.38). In relation to alcohol- and drug-related

problems, CRAFFT-caseness was associated with increased odds of all mental health problems in both the crude and the age- and gender-adjusted model (crude ORs ranging from 1.68 to 3.24). Excessive alcohol consumption was associated with all mental health problems in both crude and age- and gender-adjusted models (crude ORs ranging from 1.61 to 2.70). Binge-drinking more than 10 times was associated with increased symptoms of depression, inattention and hyperactivity in both regression models, but not with symptoms of anxiety (ORs ranging from 0.95 to 2.08).

Table 3 shows the p-values of the two-way age- and gender-interactions across use of alcohol and illicit drugs and mental health problems. For the most part, no interactions were found; only seven out 40 (17.5%) investigated interactions were statistically significant. For age, significant interactions were found with having ever consumed alcohol and frequent binge drinking in relation to hyperactivity (p=0.002 and p=0.025, respectively). In the age-stratified analyses having ever tried alcohol was associated with increased levels of hyperactivity for 17-year-olds (OR=2.53, CI95% 1.88-3.43) and 18-year-olds (OR=2.34, CI95% 1.57-3.48), but not for 19-year-olds (OR=0.79, CI95% 0.47-1.34). Frequent binge drinking was associated with increased levels of hyperactivity for all age-groups, but more strongly among 17 (OR=2.56, CI95% 1.88-3.47) and 18-year-olds (OR=2.54, CI95% 1.93-3.34) than for 19-year-olds (OR=1.55, CI95% 1.16-2.08).

For gender, significant interactions were found with having ever tried illicit drugs and excessive alcohol consumption in relation to symptoms of anxiety (p=0.021 and p<0.001, respectively), excessive alcohol consumption and symptoms of hyperactivity (p=0.021), and with binge drinking and excessive consumption and symptoms of depression (p=0.027 and p=0.018, respectively). In the gender-stratified analyses, having ever tried illicit drugs and reporting excessive consumption were associated with increased symptoms of anxiety for boys (OR=2.44, [CI95% 1.57-3.78] and OR=4.36 [CI95% 2.52-7.57], respectively), but not for girls (OR=1.30 [CI95% 0.98-1.73] and OR=1.23 [CI95% 0.87-1.73], respectively). The same pattern was evident for depression, where frequent binge drinking was associated with increased symptoms in boys (OR=1.83, CI95%1.32-2.52) but not for girls (OR=1.19, CI95% 0.98-1.44). Excessive alcohol consumption was associated with increased symptoms of hyperactivity and depression, more strongly for boys (OR=3.78 [CI95% 2.65-5.40] and OR=3.57 [CI95% 2.32-5.51], respectively) than girls (OR=2.17 [CI95% 1.60-2.94] and OR=1.89 [CI95% 1.43-2.49], respectively).

Table 4 shows the associations between alcohol consumption and binge drinking categories with never having tried alcohol as reference. For symptoms of inattention, hyperactivity and depression there was a trend towards more symptoms of mental health problems with increasing levels of usual consumption and higher

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frequency for binge drinking (all p<0.001). This trend was not found for symptoms of anxiety where there were no associations with binge drinking categories (p-value for trend: 0.679) and alcohol consumption categories (p-value for trend: 0.058). The only significant difference was found with excessive alcohol consumption (90<sup>th</sup> percentile) compared to never having tried alcohol (OR=1.60, CI95%1.15-2.23).

## DISCUSSION

#### Main findings

In sum, most adolescents aged 17 to 19 had tried alcohol and one-fifth of the girls and one-fourth of the boys had tried some illicit drug. Also, twenty percent of the adolescents reported some alcohol- or drug-related problems, as indicated by 2 or more points on the CRAFFT-questionnaire. There was a clear positive age-trend for having tried both alcohol and illicit drugs for both boys and girls, and debut of alcohol and illicit drug use was associated with mental health problems for both genders. Having tried illicit drugs, excessive alcohol consumption and CRAFFT-caseness were associated with all mental health problems. No J- or U-shaped association was found for alcohol consumption or binge drinking in relation to any of the mental health problems included in this study. There was weak evidence of any substantial age- or gender-confounding or interaction on the estimated association between alcohol- and drug-related measures and mental health problem. Moreover, a closing gender gap on alcohol and drug use was supported.

## Strengths and limitations

The current study holds several strengths. Firstly, it is a large well-defined population-based sample of adolescents. The sample size enabled the investigation of both main effects and potential interaction effects of age and gender. Secondly, the data collection is recent which allows for an updated and contemporary insight into the current status of alcohol- and drug use, presence of potential alcohol- and drug use problems, and associations with mental health problems. Thirdly, the study included different measures of alcohol- and drug use, as well as the validated CRAFFT-questionnaire for alcohol- or drug-related problems. Fourthly, we included measures of internalizing and externalizing mental health problems, using well-validated questionnaires measuring symptoms of depression and anxiety, as well as hyperactivity and inattention symptoms that are part of the ADHD-diagnosis.

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The current study has also some limitations. Firstly, the study is cross-sectional, and this precludes any causal inference of the estimated associations. Previous findings indicate that mental health problems can be a consequence of alcohol- and drug use, be a risk factor for alcohol- and drug use or that both are influenced by a common cause.[41-43] Secondly, the age-range included in the present study is limited, and a wider range would enable a more thorough investigation of age-related effects of alcohol and drug use, and probably also increase the chances of finding any moderating effects of age.[44] Thirdly, both data on exposures and outcomes were self-reported, which could lead to common methods bias and bias in reporting, especially underreporting for the exposures. Fourthly, due to missing information, almost ten percent of the total eligible study population was not included in our analyses. The attrition from the current study could affect generalizability, and result in adolescents attending school being overrepresented. Although no data on non-responders were available in the current wave of the Bergen Child Study, previous research from the former waves of the Bergen Child Study have shown that non-participants tend to have more psychological problems than participants.[45] This could suggest that the non-responders also have a higher alcohol and drug consumption, which could bias our sample further. This non-participation bias is, however, not likely to reduce the associations of interest substantially.[46] Finally, the assessment of illicit drug does not indicate type of drug tried and frequency, thus limiting conclusions drawn according to the relation between mental health and more specific drugs or consumption patterns.

## Interpretation

The closing gender gap reported in previous studies, was partly confirmed by the present study.[20] Even though more girls in the younger age group had tried alcohol, boys showed a pattern of trying illegal drugs more often and there were more boys with problems related to alcohol and drug use, while there were no differences in patterns of problem drinking and binge drinking. The expected age pattern was confirmed with both more alcohol and drug use and binge drinking with increasing age. 18 years is the legal age when adolescents can buy alcohol in Norway, and thus an increase in alcohol use at this age is as expected. In the 19-years-olds more than a third reported binge drinking. Even though this is a lower rate than reported in a previous population-based Norwegian study ,[23] our findings suggest that this negative alcohol use pattern is highly frequent among Norwegian adolescents.

There was strong evidence for a co-occurrence of alcohol and drug problems and mental health problems among adolescents, consistent with findings in previous studies.[7, 25] While most studies have

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focused on drug abuse, we also found that even having tried alcohol or drugs in this age group are potential indicators of mental health problems for both genders. Due to the cross-sectional design of this study, we could not investigate direction of causality. But the relationship between alcohol and drug use and mental health problems in adolescence is probably complex and multidirectional.[24, 47] For instance, mental health problems like impulsivity and hyperactivity may possibly render an adolescent susceptible for using alcohol, but these problems may also be exacerbated by alcohol and drug use. There may also be common etiological factors that accounts for both an increased risk of mental health problems and alcohol consumption, such as parental alcohol use or social peer network. While both ADHD and depressive symptoms were consistently related to alcohol and drug use and abuse, the co-occurrence of these disorders may have impacted the association.

In general, we found little evidence of a substantial modifying effect of age or gender, which is contrary to some previous studies.[23, 26, 27] We did, however, find weak evidence for a modifying effect of age and gender for specific mental health domains: there was some evidence for an age-differential association between alcohol and drug exposures and symptoms of hyperactivity, where having tried alcohol or frequent binge drinking were more strongly associated with symptoms of hyperactivity for younger adolescents. This could be related to a lower level of cognitive and emotional control in early adolescence compared to at an older age,[44] increasing the susceptibility both to alcohol use, binge drinking and mental health problems.

We also found a gender-differential association between alcohol and drug exposures and symptoms of internalizing mental health problems, where boys showed more consistent association between exposures and symptoms of anxiety and depression than girls. Interestingly, we found that, among girls, excessive alcohol consumption was not associated with anxiety and frequent binge drinking was not associated with depression. This is contrary to results from a previous Finnish study which found stronger associations between heavy drinking and psychosocial dysfunction among girls than among boys.[27] One factor that may have impacted on this relation is the higher rate of depressive symptoms in girls,[36] and this may be understood in light of the "gender paradox" where the gender with the lowest prevalence rate will be more severely affected.[48] The interpretation of the findings concerning interactions should, however, be done with caution as the number of tested interactions (N=40) were high, and only 17.5% were significant. Future studies should therefore investigate the reported interactions in other study samples.

In the secondary analyses, we found no indication of any curvilinear relationship between alcohol consumption and frequency of binge drinking in relation to mental health problems. This is contrary to previous findings,[8-12] predominantly from adult populations, and may be an indication of a differential association

between alcohol habits and mental health problems in adolescent versus in an adult population. Also, it may be an indication of cohort-effects, as there is a tendency for lower levels of alcohol consumption and more abstention among adolescents in recent years.[4, 49] Thus, as the abstention group grows, the group also includes more individuals who identify with being abstinent, who have a social network and actively chooses abstinence due to an increased focus on education, career or sports.[50]

## **Clinical implications**

The high rate of alcohol and drug use in underage adolescents, 10 percent having tried illegal substances, as well as a pattern of negative alcohol and drug use characterised by binge drinking and large quantities of alcohol together represent a serious public health concern that emphasizes a need of preventive interventions. The strong co-occurrence of mental health and alcohol and drug related problems suggest that assessment of alcohol and drug use and abuse should become an integral part of mental health assessment and interventions, especially in adolescent with depression and ADHD.

In summary, it seems evident that both alcohol and drug debut, and the frequency and intensity of alcohol consumption are important and relevant indicators of mental health problems among adolescents. Investigation of the causal nature between alcohol and drug use, and mental health problems, as well as the access to health care and utilisation of health care among adolescents who use alcohol and drugs in hazardous ways should be addressed in future studies.

## **COMPETING INTERESTS**

The authors declare that they have no competing interests.

# **AUTHORS' CONTRIBUTIONS**

JCS conceived the initial concept, and the initial analyses and literature search. JCS, BS, MH interpreted the initial analyses and wrote the first draft. AJL, KMS and RJ reviewed and revised the initial analyses and first draft, and suggested further analyses, as well as revisions of the first draft. The subsequent drafts were reviewed by all authors, and all authors read and approved the final manuscript.

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# REFERENCES

1.	Rehm, J., et al., Alcohol use disorders in EU countries and Norway: An overview of
	the epidemiology. European Neuropsychopharmacology, 2005. 15: p. 377-388.
2.	Degenhardt, L. and W. Hall, Extent of illicit drug use and dependence, and their
	contribution to the global burden of disease. The Lancet, 2012. 379(9810): p. 55-70.
3.	Hellandsjø Bu, E.T., et al., Teenage alcohol and intoxication debut: The impact of
	family socialization factors, living area and participation in organized sports. Alcohol
	and Alcoholism, 2002. 37(1): p. 74-80.
4.	Hibell, B., et al., Summary: The 2007 ESPAD report among students in 35 countries.
	2009, European Monitoring Centre for Drugs and Drug Addiction: Luxembourg.
5.	Hingson, R.W., T. Heeren, and M.R. Winter, Age at drinking onset and alcohol
	dependence: Age at onset, duration, and severity. Archives of Pediatrics & Adolescent
	Medicine, 2006. 160(7): p. 739-746.
6.	Pedersen, W. and A. Skrondal, Alcohol consumption debut: Predictors and
	consequences. Journal of Studies on Alcohol, 1998. 59(1): p. 32-42.
7.	Costello, E.J., T.D. Armstrong, and A. Erkanli, Report on the developmental
	epidemiology of comorbid psychiatric and substance use disorders. 2000, National
	Institute on Drug Abuse.
8.	Rodgers, B., et al., Non-linear relationships in associations of depression and anxiety
	with alcohol use. Psychological Medicine, 2000. 30(2): p. 421-32.
9.	Rodgers, B., et al., Risk factors for depression and anxiety in abstainers, moderate
	drinkers and heavy drinkers. Addiction, 2000. 95(12): p. 1833-1845.
10.	Caldwell, T.M., et al., Patterns of association between alcohol consumption and
	symptoms of depression and anxiety in young adults. Addiction, 2002. 97: p. 583-594.
11.	Skogen, J.C., et al., Anxiety and depression among abstainers and low-level alcohol
	consumers. The Nord-Trøndelag Health Study. Addiction, 2009. 104(9): p. 1519-1529.

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2 3	12.	Pape, H. and T. Hammer, Sober adolescence - Predictor of psychosocial
4 5		maladjustment in young adulthood? Scandinavian Journal of Psychology, 1996. 37: p.
6 7 8		362-277.
9 10	13.	Upmark, M., J. Möller, and A. Romelsjö, Longitudinal, population-based study of self
11 12		reported alcohol habits, high levels of sickness absence, and disability pensions.
13 14		Journal of Epidemiology and Community Health, 1999. 53: p. 223-229.
15 16	14.	Brecht, ML., et al., Methamphetamine use behaviors and gender differences.
17 18		Addictive behaviors, 2004, <b>29</b> (1); p. 89-106.
19 20	15	Westermeyer L and A F. Boedicker Course severity and treatment of substance
21 22	15.	westernieger, 5. and A.E. Boedieker, Course, severny, und treatment of substance
23 24		abuse among women versus men. American Journal of Drug and Alcohol Abuse,
25		2000. <b>26</b> (4): p. 523-535.
26 27		
28	16.	Hser, YI., et al., <i>Gender differences in treatment outcomes over a three-year period:</i>
29		A nath model analysis, Journal of Drug Issues, 2004, 34: p. 419-440
30 31		<i>I pun mouei unuiysis.</i> Journal of Drug 13003, 2004. <b>54</b> . p. 417-440.
32	17.	Hernandez-Avila, C.A., B.J. Rounsaville, and H.R. Kranzler, Opioid-, cannabis- and
33 34		alcohol-dependent women show more rapid progression to substance abuse treatment.
35 36		Drug and Alcohol Dependence, 2004. 74(3): p. 265-272.
37		
30 39	18.	Haas, K., Relationship of gender to licit and illicit drug use among adolescents.
40		Chrestomathy: Annual Review of Undergraduate Research at the College of
41		
43		Charleston, 2004. <b>3</b> : p. 92-100.
44 45	10	Onland F.A. K.C. Winters and P.D. Stinghfield Examining and a life-
46	19.	Opland, E.A., K.C. winters, and K.D. Sunchlield, Examining gender differences in
47		<i>drug-abusing adolescents</i> . Psychology of Addictive Behaviors, 1995. <b>9</b> (3).
48		
49 50	20.	Keyes, K.M., B.F. Grant, and D.S. Hasin, Evidence for a closing gender gap in
51		alcohol use abuse and dependence in the United States population Drug Alcohol
52 53		aconoi use, ubuse, una acpenaence in ine Onnea States population. Diag Meditor
54		Depend, 2008. <b>93</b> (1-2): p. 21-9.
55		
56 57		
58		
59		

Schwinn, T.M., S.P. Schinke, and D.N. Trent, *Substance use among late adolescent urban youths: Mental health and gender influences*. Addictive behaviors, 2010. 35(1):
p. 30-34.

- Armstrong, T.D. and E.J. Costello, *Community studies on adolescent substance use, abuse, or dependence and psychiatric comorbidity.* Journal of Consulting and Clinical Psychology, 2002. 70(6): p. 1224-1239.
- 23. Strandheim, A., et al., Alcohol intoxication and mental health among adolescents a population review of 8983 young people, 13-19 years in North-Trøndelag, Norway: the Young-HUNT Study. Child and Adolescent Psychiatry and Mental Health, 2009.
  3(18).
- Trim, R.S., et al., *The relation between adolescent substance use and young adult internalizing symptoms: Findings from a high-risk longitudinal sample*. Psychology of Addictive Behaviors, 2007. 21(1): p. 97-107.
- 25. McCarty, C.A., et al., *Developmental consistency in associations between depressive symptoms and alcohol use in early adolescence*. Journal of studies on Alcohol and Drugs, 2012. **73**: p. 444-453.
- 26. Verdurmen, J., et al., *Alcohol use and mental health in adolescents: Interactions with age and gender Findings from the Dutch 2001 Health Behaviour in School-Aged Children Survey.* Journal of Studies on Alcohol and Drugs, 2005. **66**(5): p. 605-609.
- 27. Laukkanen, E.R., et al., *Heavy drinking is associated with more severe psychosocial dysfunction among girls than boys in finland*. The Journal of adolescent health :
  official publication of the Society for Adolescent Medicine, 2001. 28(4): p. 270-277.
- Theunissen, M.-J., M. Jansen, and A. van Gestel, Are mental health and binge drinking associated in Dutch adolescents? Cross-sectional public health study. BMC Research Notes, 2011. 4(1): p. 100.

# **BMJ Open**

29.	McLeod, J.D., R. Uemura, and S. Rohrman, Adolescent Mental Health, Behavior
	Problems, and Academic Achievement. Journal of Health and Social Behavior, 2012.
	<b>53</b> (4): p. 482-497.
30.	Skogen, J.C., et al., Concurrent validity of the CAGE questionnaire. The Nord-
	Trøndelag Health Study. Addictive behaviors, 2011. 36(4): p. 302-307.
31.	Knight, J.R., et al., A new brief screen for adolescent substance abuse. Archives of
	Pediatrics & Adolescent Medicine, 1999. 153: p. 591-596.
32.	Knight, J.R., et al., Validity of brief alcohol screening tests among adolescents: A
	comparison of the AUDIT, POSIT, CAGE, and CRAFFT. Alcoholism: Clinical and
	Experimental Research, 2003. 27(1): p. 67-73.
33.	Skogen, J.C., et al., Psychometric properties and concurrent validity of the CRAFFT
	among Norwegian adolescents. Ung@hordaland, a population-based study. Addictive
	behaviors, 2013(Ahead of print).
34.	Thapar, A. and P. McGuffin, Validity of the shortened Mood and Feelings
	Questionnaire in a community sample of children and adolescents: a preliminary
	research note. Psychiatry Res., 1998. 81(2): p. 259-68.
35.	Sharp, C., I.M. Goodyer, and T.J. Croudace, The Short Mood and Feelings
	Questionnaire (SMFQ): a unidimensional item response theory and categorical data
	factor analysis of self-report ratings from a community sample of 7-through 11-year-
	old children. J Abnorm Child Psychol, 2006. <b>34</b> (3): p. 379-91.
36.	Lundervold, A.J., et al., Symptoms of depression as reported by Norwegian
	adolescents on the Short Mood and Feelings Questionnaire. Frontiers in Psychology,
	2013. 4.

37. Birmaher, B., et al., Psychometric properties of the screen for child anxiety related emotional disorders scale (SCARED): A replication study. Journal of the American Academy of Child and Adolescent Psychiatry, 1999. 38(10): p. 1230-1236. 38. Kessler, R.C., et al., Validity of the World Health Organization Adult ADHD Selfreport Scale (ASRS) Screener in a representative sample of health plan members. International Journal of Methods in Psychiatric Research, 2007. 16(2): p. 52-65. 39. Adler, L.A., et al., Preliminary examination of the reliability and concurrent validity of the attention-deficit/hyperactivity disorder self-report scale v1.1 symptom checklist to rate symptoms of attention-deficit/hyperactivity disorder in adolescents. Journal of Child and Adolescent Psychopharmacology, 2012. 22(3): p. 238-44. 40. StataCorp, Statistical Software: Release 12.0. 2012, Stata Corporation: College Station, TX. Tapert, S.F., L. Caldwell, and C. Burke, Alcohol and the adolescent brain: Human 41. studies. Alcohol Research & Health, 2004. 28(4): p. 205-212. 42. Chassin, L., et al., A longitudinal study of children of alcoholics: Predicting young adult substance use disorders, anxiety, and depression. Journal of Abnormal Psychology, 1999. **108**(1): p. 106-119. 43. Sher, K.J., et al., *Characteristics of children of alcoholics: Putative risk factors*, substance use and abuse, and psychopathology. Journal of Abnormal Psychology, 1991. **100**(4): p. 427-448. 44. Willoughby, T., et al., Examining the link between adolescent brain development and risk taking from a social-developmental perspective. Brain and Cognition, 2013. (3): p. 315-323.

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54 55		
56 57		
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- 45. Stormark, K.M., et al., *Predicting nonresponse bias from teacher ratings of mental health problems in primary school children*. J Abnorm Child Psychol, 2008. 36(3): p. 411-9.
- 46. Knudsen, A.K., et al., *The health status of nonparticipants in a population-based health study: The Hordaland Health Study*. American Journal of Epidemiology, 2010.
  172(11): p. 1306-14.
- 47. O'Neil, K.A., B.T. Conner, and P.C. Kendall, *Internalizing disorders and substance use disorders in youth: Comorbidity, risk, temporal order, and implications for intervention.* Clinical Psychology Review, 2011. **31**(1): p. 104-112.
- 48. Loeber, R. and K. Keenan, *Interaction between Conduct Disorder and Its Comorbid* -*Effects of Age and Gender*. Clinical Psychology Review, 1994. **14**(6): p. 497-523.
- 49. Danielsson, A.-K., et al., *Alcohol use, heavy episodic drinking and subsequent problems among adolescents in 23 European countries: does the prevention paradox apply?* Addiction, 2012. **107**(1): p. 71-80.
- 50. Hoel, S., et al., Adolescent alcohol use, psychological health, and social integration.
   Scandinavian Journal of Public Health, 2004. 32: p. 361-367.

Figure 1: Proportion of participants having ever consumed alcohol and tried illicit drugs across gender-stratified age groups (N=9,203, girls n=4,995, boys n=4,208). Error bars represent 95% confidence intervals.



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	Total sample (95% CI)	Born 1995 (95% CI)	Born 1994 (95% CI)	Born 1993 (95% CI)	OR for age trend (95% CI)	p-value
Never consumed alcohol (yes%)						
Girls	20.7% (19.6-21.8)	31.4% (29.4-33.5)	19.4% (17.6-21.4)	6.3% (5.1-7.7)	0.42 (0.38-0.47)	<.001
Boys	27.1% (25.7-28.4)	39.7% (37.4-42.1)	26.4% (24.2-28.7)	7.6% (6.1-9.4)	0.41 (0.37-0.45)	<.001
Ever tried illicit drugs (yes%)						
Girls	8.8% (8.1-9.6)	6.1% (5.1-7.3)	8.2% (7.0-9.6)	13.7% (11.9-15.6)	1.57 (1.39-1.77)	<.001
Boys	12.5% (11.5-13.5)	10.8% (8.5-11.2)	10.2% (8.8-11.9)	18.5% (16.3-21.0)	1.37 (1.22-1.54)	<.001
90 <sup>th</sup> percentile alcohol consumption <sup>a</sup> (yes%)						
Girls	7.7% (6.9-8.6)	6.0% (4.8-7.4)	7.7% (6.4-9.2)	9.7% (8.2-11.5)	1.30 (1.13-1.51)	<.001
Boys	7.4% (6.5-8.4)	5.9% (4.6-7.6)	5.9%(5.5-8.5)	9.6% (7.9-11.6)	1.31 (1.10-1.55)	=.002
Binge drinking (>10; yes%) <sup>b</sup>						
Girls	20.5% (19.4-21.6)	10.6% (9.3-12.0)	19.7% (17.9-21.7)	36.1% (33.6-38.8)	2.20 (2.01-2.41)	<.001
Boys	21.3% (20.1-22.6)	10.7% (9.3-12.3)	21.9% (19.9-24.1)	37.1% (34.2-40.1)	2.21 (2.00-2.44)	<.001
CRAFFT positive (≥2; yes%)						
Girls	23.3% (22.2-24.5)	20.0% (18.3-21.8)	22.4% (20.5-24.4)	29.5% (27.1-32.1)	1.29 (1.19-1.40)	<.001
Boys	20.3% (19.1-21.5)	15.8% (14.1-17.6)	19.3% (17.4-21.4)	29.0% (26.3-31.8)	1.47 (1.34-1.62)	<.001

<sup>b</sup>Only includes those with valid response on binge drinking (n=9,056).

Table 2: Logistic regressions of use of alcohol and illicit drugs as risk factor for mental health problems. N=9,203 (girls n=4,995, boys n=4,208).

	ASRS Inattention OR (95%CI)	ASRS Hyperactivity OR (95%CI)	SMFQ OR (95%CI)	SCARED OR (95%CI)
Ever consumed alcohol				
Crude	2.36 (1.90-2.93)	2.08 (1.68-2.58)	1.73 (1.43-2.09)	1.13 (0.94-1.37)
Adjusted for age and gender	2.21 (1.77-2.76)	2.07 (1.66-2.58)	1.64 (1.35-1.99)	1.03 (0.84-1.25)
Ever tried illicit drugs				
Crude	2.38 (1.98-2.87)	2.26 (1.86-2.74)	1.69 (1.39-2.05)	1.33 (1.05-1.68)
Adjusted for age and gender	2.49 (2.06-3.02)	2.36 (1.94-2.89)	1.98 (1.62-2.42)	1.55 (1.23-1.98)
90 <sup>th</sup> percentile alcohol consumption <sup>a</sup>				
Crude	1.97 (1.55-2.51)	2.70 (2.14-3.40)	2.19 (1.74-2.75)	1.61 (1.21-2.14)
Adjusted for age and gender	1.96 (1.54-2.50)	2.76 (2.19-3.48)	2.27 (1.79-2.88)	1.64 (1.22-2.19)
Binge >10 times <sup>b</sup>				
Crude	1.83 (1.56-2.15)	2.08 (1.77-2.45)	1.30 (1.10-1.52)	0.95 (0.78-1.16)
Adjusted for age and gender	1.80 (1.52-2.12)	2.18 (1.84-2.59)	1.36 (1.15-1.61)	0.97 (0.79-1.20)
CRAFFT-caseness (≥2)				
Crude	3.15 (2.71-3.66)	3.24 (2.78-3.78)	2.84 (2.46-3.28)	1.68 (1.42-1.99)
Adjusted for age and gender	3.06 (2.63-3.56)	3.22 (2.76-3.77)	2.84 (2.45-3.29)	1.62 (1.38-1.94)

95% confidence intervals in parentheses

 <sup>a</sup>Only includes those who have tried alcohol (n=7,031).

<sup>b</sup>Only includes those with valid response on binge drinking (n=9,056).

Table 3: Summary of level of statistical significance for age- and gender-interactions across exposures and outcomes. N=9,203 (girls n=4,995, boys n=4,208).

	ASRS in	ASRS inattention		ASRS Hyperactivity		SMFQ		SCARED	
	Gender	Age	Gender	Age	Gender	Age	Gender	Age	
Ever consumed alcohol	p=0.187	p=0.595	p=0.466	p=0.002	p=0.257	p=0.255	p=0.052	p=0.182	
Ever tried illicit drugs	p=0.666	p=0.630	p=0.542	p=0.384	p=0.054	p=0.978	p=0.021	p=0.425	
90 <sup>th</sup> percentile alcohol consumption	p=0.752	p=0.359	p=0.021	p=0.937	p=0.018	p=0.639	p<0.001	p=0.198	
Binge >10	p=0.083	p=0.427	p=0.514	p=0.025	p=0.027	p=0.889	p=0.245	p=0.452	
CRAFFT-caseness (≥2)	p=0.413	p=0.263	p=0.293	p=0.190	p=0.100	p=0.840	p=0.991	p=0.464	

\*Significant interactions in bold. Interactions assessed using Wald likelihood-ratio tests.

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Table 4: Logistic regression models across alcohol consumption and binge drinking categories, "Never tried alcohol" as reference. Adjusted for age and gender (N=9,203 for alcohol consumption and N=9,056 for binge drinking).

	ASRS Inattention	ASRS Hyperactivity	SMFQ	SCARED		
	OR (95%CI)	OR (95%CI)	OR (95%CI)	OR (95%CI)		
Categories		Alcohol consumption ca	ategories as exposure			
Never consumed alcohol	1.00 (Reference)	1.00 (Reference)	1.00 (Reference)	1.00 (Reference)		
Non-consumption	1.90 <sup>***</sup>	1.46 <sup></sup>	1.43 <sup>**</sup>	1.01		
	(1.45-2.48)	(1.10-1.93)	(1.12-1.82)	(0.78-1.30)		
0.1-19.9 <sup>th</sup> percentile	1.61 <sup>**</sup>	1.69 <sup>**</sup>	1.44 <sup>**</sup>	1.00		
	(1.19-2.17)	(1.25-2.27)	(1.11-1.88)	(0.75-1.32)		
20.0-39.9 <sup>th</sup> percentile	1.98	1.91	1.53	0.90		
	(1.46-2.69)	(1.40-2.60)	(1.16-2.03)	(0.66-1.23)		
40.0-59.9 <sup>th</sup> percentile	1.92 (1.42-2.59)	1.55 <sup>°°</sup> (1.13-2.13)	1.16 (0.87-1.56)	0.97 (0.72-1.30)		
60.0-79.9 <sup>th</sup> percentile	2.56	2.55	1.85	0.84		
	(1.92-3.40)	(1.92-3.40)	(1.42-2.41)	(0.62-1.14)		
80.0-89.9 <sup>th</sup> percentile	3.77	3.56	2.21	1.39		
	(2.75-5.17)	(2.59-4.91)	(1.62-3.02)	(0.99-1 <u>.</u> 96)		
90.0-100 <sup>th</sup> percentile	4.16	5.28	3.47	1.60		
	(3.05-5.68)	(3.91-7.12)	(2.60-4.62)	(1.15-2.23)		
o-value for trend	p<0.001	p<0.001	p<0.001	p=0.058		
Categories	Binge drinking categories as exposure					
Never consumed alcohol	1.00 (Reference)	1.00 (Reference)	1.00 (Reference)	1.00 (Reference)		
Never binge	1.65 <sup></sup> (1.24-2.20)	1.35 <sup>*</sup> (1.00-1.81)	1.30 (1.00-1.67)	1.10 (0.85-1.43)		
Dnce	1.71 <sup>**</sup>	2.06 <sup>***</sup>	1.55 <sup>**</sup>	1.02		
	(1.23-2.38)	(1.50-2.82)	(1.16-2.08)	(0.74-1.39)		
2-3 times	2.00 <sup>m</sup>	1.62 <sup>**</sup>	1.61 <sup></sup>	0.88		
	(1.51-2.63)	(1.21-2.15)	(1.26-2.07)	(0.67-1.16)		
-10 times	2.49	2.05	1.85	1.05		
	(1.90-3.26)	(1.55-2.72)	(1.44-2.36)	(0.81-1.38)		
•10 times	3.14	3.36	1.95	0.98		
	(2.43-4.05)	(2.61-4.33)	(1.54-2.47)	(0.76-1.27)		
o-value for trend	p<0.001	p<0.001	p<0.001	p=0.679		

p < 0.05, p < 0.01, p < 0.001 in comparison to "Never tried alcohol"-category

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# **Figure legends**

 Figure 1: Proportion of participants having ever consumed alcohol and tried illicit drugs across gender-stratified age groups (N=9,203, girls n=4,995, boys n=4,208). Error bars represent 95% confidence intervals.

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# **BMJ Open**

STROBE Statement—checklist of items that should be included in reports of observational studies **"X" indicates checked** 

	Item No	Recommendation
Title and abstract	1 X	(a) Indicate the study's design with a commonly used term in the title or the abstract
The and about act	1 24	(b) Provide in the abstract an informative and balanced summary of what was done
		and what was found
Inter la dian		
	2 V	Fundain the activities have been developed and strate the device of the basic second of
Background/rationale	2 X	Explain the scientific background and rationale for the investigation being reported
Objectives	3 X	State specific objectives, including any prespecified hypotheses
Methods		
Study design	4 X	Present key elements of study design early in the paper
Setting	5 X	Describe the setting, locations, and relevant dates, including periods of recruitment,
		exposure, follow-up, and data collection
Participants	6 X	(a) Cohort study—Give the eligibility criteria, and the sources and methods of
		selection of participants. Describe methods of follow-up
		<i>Case-control study</i> —Give the eligibility criteria, and the sources and methods of
		case ascertainment and control selection. Give the rationale for the choice of cases
		and controls
		Cross-sectional study—Give the eligibility criteria, and the sources and methods of
		selection of participants
		(b) Cohort study—For matched studies, give matching criteria and number of
		exposed and unexposed
		Case-control study-For matched studies, give matching criteria and the number of
		controls per case
Variables	7 X	Clearly define all outcomes, exposures, predictors, potential confounders, and effect
		modifiers. Give diagnostic criteria, if applicable
Data sources/	8* X	For each variable of interest, give sources of data and details of methods of
measurement		assessment (measurement). Describe comparability of assessment methods if there
		is more than one group
Bias	9 X	Describe any efforts to address potential sources of bias
Study size	10 X	Explain how the study size was arrived at
Quantitative variables	11 X	Explain how quantitative variables were handled in the analyses. If applicable,
		describe which groupings were chosen and why
Statistical methods	12 X	(a) Describe all statistical methods, including those used to control for confounding
		(b) Describe any methods used to examine subgroups and interactions
		(c) Explain how missing data were addressed
		(d) Cohort study—If applicable, explain how loss to follow-up was addressed
		Case-control study-If applicable, explain how matching of cases and controls was
		addressed
		Cross-sectional study-If applicable, describe analytical methods taking account of
		sampling strategy
		( <u>e</u> ) Describe any sensitivity analyses

Continued on next page

Results		
Participants	13* X	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed
		(b) Give reasons for non-participation at each stage
		(c) Consider use of a flow diagram
Descriptive	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and
data	Х	information on exposures and potential confounders
		(b) Indicate number of participants with missing data for each variable of interest
		(c) Cohort study—Summarise follow-up time (eg, average and total amount)
Outcome data	15	Cohort study—Report numbers of outcome events or summary measures over time
	X*	<i>Case-control study</i> —Report numbers in each exposure category, or summary measures of exposure
		Cross-sectional study—Report numbers of outcome events or summary measures
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their
	Х	precision (eg, 95% confidence interval). Make clear which confounders were adjusted for and
		why they were included
		(b) Report category boundaries when continuous variables were categorized
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a
		meaningful time period
Other analyses	17	Report other analyses done-eg analyses of subgroups and interactions, and sensitivity
	Х	analyses
Discussion		
Key results	18	Summarise key results with reference to study objectives
	Х	
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision.
	Х	Discuss both direction and magnitude of any potential bias
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations,
	Х	multiplicity of analyses, results from similar studies, and other relevant evidence
Generalisability	21	Discuss the generalisability (external validity) of the study results
	Х	
Other information	on	
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable,
	Х	for the original study on which the present article is based

\*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

**Note:** An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.

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# Alcohol and drug use among adolescents - and the cooccurrence of mental health problems. Ung@hordaland, a population-based study.

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SCHOLARONE<sup>™</sup> Manuscripts

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## ABSTRACT

**Objectives:** The use of alcohol and drugs is prevalent among adolescents, but too little is known about the association between debut of alcohol and drug use, problematic use and concurrent mental health. The aim of the study was to investigate the cross-sectional association between debut of any alcohol or drug use and alcohol- and drug-related problems, and mental health. We also wanted to examine potential interactions between gender and age, and alcohol- and drug-related variables.

Design: Cross-sectional study.

Setting: Population-based sample of Norwegian adolescents.

**Participants:** Data stem from the large population-based ung@hordaland study (N=9,203), where all adolescents aged 16-18 years living in Hordaland county (Norway) were invited to participate. The main independent variables were debut of alcohol and drug use, alcohol consumption, and presence of alcohol and drug problems as measured by CRAFFT.

**Outcomes:** The dependent variables were self-reported symptoms of anxiety, depression, inattention and hyperactivity. Statistical analyses included logistic regression models.

**Results:** Both debut of alcohol and drug use were associated with symptoms of depression, inattention and hyperactivity (crude ORs 1.69-2.38, Ps<0.001), while only debut of drug use was associated with increased symptoms of anxiety (OR=1.33, CI95%1.05-1.68, P=0.017). Alcohol- and drug-related problems as measured by CRAFFT were associated with all mental health problems (crude ORs 1.68-3.24, Ps<0.001). There was little evidence of any substantial age- or gender confounding on the estimated associations between alcohol- and drug-related measures and the mental health problems.

**Conclusion:** Early debut of alcohol and drug use and drug problems are consistently associated with more symptoms of mental health problems, indicating that these factors are an important general indicator of mental health in adolescence.

# STRENGTHS AND LIMITATIONS OF THIS STUDY

- This study employed a large well-defined population-based sample of adolescents.
- The data employed in this study is from a recent data collection.
- This study included several different measures of alcohol- and drug use, as well as measures of both internalizing and externalizing problems.
- The cross-sectional design of this study precluded any causal inference.
- This study only had a limited age-range; a wider range would enable a more thorough investigation of age-related effects of alcohol and drug use.

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### BACKGROUND

Previous studies of adolescents have found that a considerable proportion with one mental disorder also meets the criteria for at least one other mental disorder, including substance-use disorder.[1-3] Furthermore, drug use and abuse have been related to a range of mental health problems, most consistently to externalizing problems such as those associated with conduct disorder (CD) and attention-deficit-hyperactivity disorder (ADHD),[4] while the increased risk for anxiety disorders are lower. Also of note, several previous studies in adult populations have found that there is a J- or U-shaped relationship between alcohol consumption and mental health outcomes,[5-9] where both abstainers and high-level alcohol consumers have increased risk for mental health problems. Several explanations of this curvilinear relationship have been suggested: moderate alcohol consumers may directly benefit from a protective effect on mental health from alcohol, or their drinking habits may be a mere reflection of their psychological wellbeing.[6] Also, in cultures where alcohol use is the norm, moderate consumers may be more well-adjusted than individuals at the extremes of the consumption spectrum.[10] To date, few studies have investigated the relationship between the full spectrum of alcohol consumption (from non-debut to high-level alcohol consumption) and symptoms of mental health problems in a population-based sample of adolescents

Boys have an earlier debut of both alcohol consumption and intoxication according to most studies,[11] while the gender-pattern for onset of drug use is more unclear.[12-17] The established gender differences with males using more alcohol and drugs seems, however, to be closing [18] with recent studies not finding a higher rate of drug use among males.[19] Whether the relation between alcohol and drug use and mental health is gender specific is less clear. A lack of gender difference has been documented in population-based studies where the use of alcohol and drugs is associated with worse mental health.[20-23] In contrast, some studies have reported a modifying effect of gender and age in relation to the association between alcohol use and mental health.[21, 24, 25] For instance, a study investigating the association between alcohol intoxications and mental health problems among 13-19-year-olds found that high level alcohol consumption was associated with conduct and attention problems for both genders, but only the girls who reported frequent intoxication reported a modifying effect of age on the association between alcohol use and mental health, but found no evidence of any modifying effects of gender.[24] The association between alcohol use and mental health is, however, not quite consistent, as a Dutch study did not confirm the expected relation between binge drinking and mental health problems among

16-18-years-olds in a population based study.[26]. Furthermore, most of the previous studies have focused on drug and alcohol use, whereas less is known about the relationship between mental health and debut in adolescence.

In general, the mental health problems show a consistent gender pattern in adolescence, with girls having a higher rate of depression and internalizing problems, and boys a higher rate of externalising disorders. [1] Less is known whether the associations between the drug use and mental health is gender-specific, but there are some indications that co-occurrence of other mental health problems are higher for girls than boys.[4] However, in a sample of late adolescent urban youth gender did not have a moderating effect on the relationship between substance abuse and mental health, and a Finnish study did not support a gender specific association between depression and substance use.[19, 27]

Also, it is uncertain if the co-occurrence is specifically related to alcohol or drug dependence or if it is also evident for mere alcohol or drug use.[4] Overall, there is still a need for further investigation of the association between early alcohol and drug use and mental health in a general population of adolescents.[28]

To that end, the aim of the present study was to

- Describe the debut of alcohol and drug use, alcohol use patterns and potential drug problems among Norwegian adolescents using data from a recent population-based study.
- ii) To investigate the cross-sectional relationship between a) alcohol and illicit drug use and b) alcohol and drug problems and mental health among adolescents. We also investigated any interactions with age and gender on the associations of interest, as well as the association between the full spectrum of alcohol consumption and frequency of drinking to intoxication in relation to mental health.

## **METHODS**

#### **Study population**

This study employed information from the ung@hordaland-study of adolescents in the county of Hordaland in western Norway. All adolescents born 1993-1995 living in Hordaland were invited to participate (n=19,430), and 10,220 participated, yielding a participation rate of 53%. Ung@hordaland is a population-based study, carried out in close collaboration with the county council administration. The fourth wave of the longitudinal population-based study "Bergen Child Study" is also nested within ung@hordaland. The general aim of the ung@hordaland-survey was to assess mental health, lifestyle, school performance and health-service use in

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adolescents, with a special emphasis on the prevalence of mental health problems. The data was collected during January-February 2012. Adolescents in upper secondary education received information per e-mail, and one school hour was allocated for them to complete the questionnaire at school. Those not in school received information by mail to their home addresses. In addition, mental health facilities, juvenile detention centres and other institutions were contacted and the adolescents had the opportunity to participate. The questionnaire was web-based and covered a broad range of mental health issues, daily life functioning, use of health and social services, and demographic background variables. Upon participation electronic informed consent was obtained. The study was approved by The Regional Committee for Medical Research Ethics in Western Norway. Hordaland County is generally regarded representative of Norway as a whole, comprising both the second largest city of Norway (Bergen), as well as large rural areas. Official statistics of the general population (not just adolescents) show that Hordaland County does not deviate in any substantial degree from the national average on key parameters, including both socio-demographic indicators and several health indicators.

The current study is based on the first version of data files released in May 2012, with a total of 10,220 participants. Based on list-wise deletion of those with missing information on both independent and dependent variables, n=1,017 were excluded, leaving 9,203 in the final study sample. Those with missing information on the variables of interest were slightly younger than those with valid information on the included variables (mean difference 0.12 years, p<.001) and more likely to be males (61.6% versus 45.7% in the analysed sample, p<.001).

## **Demographic information**

Information about age and gender was obtained on all participants. Furthermore, the participants indicated the perceived family socioeconomic status (SES) as i) "about the same as others" (67.9%), ii) "better than others" (25.1%) or iii) "worse than others" (7.1%), with n=172 missing responses recoded to "about the same as others".

#### Identification of alcohol problems and illicit drug use

Self-reported debut, frequency and amount of alcohol consumption and illicit drug use were included as main independent variables. We included a binary measure of alcohol debut "Have you ever tried alcohol?" (Yes/No), and illicit drug use debut: "Have you ever tried hash, marihuana or other narcotic substances?" (Yes/No). Out of the total study sample, n=11 individuals did not answer the question on illicit drug use, and they were recoded to the "No" category. Alcohol use was measured using the self-reported units of beer, cider, wine, spirits and

illegally distilled spirits usually consumed during the last 14 days. Based on this information about consumption and alcohol debut, a summed variable on gender-specific distributions was constructed: "Never tried", "Nonconsumption" (if reported consumption was "0"), "0.1-19.9<sup>that</sup>, "20.0-79.9<sup>th</sup>", "80.0-89.9<sup>th</sup>", "90.0-100<sup>th</sup>". Excessive alcohol consumption was defined as above the 90<sup>th</sup> percentile sum score.[8, 29] Frequent intoxication (binge drinking) was defined as drinking so much that one was clearly intoxicated more than 10 times, based on the question: "Have you ever consumed so much alcohol that you were clearly intoxicated (drunk)?", with five categories ranging from "No, never" to "Yes, more than 10 times". There were n=147 missing responses on the intoxication item, leaving a sample of N=9,056 for this measure. In addition, potential alcohol and drug problems were identified employing the six-item inventory CRAFFT.[30] CRAFFT was specifically designed to identify alcohol and illicit drug problems among adolescents,[30] and have been shown to have acceptable sensitivity and specificity at a cut-off of >=2 in international studies.[31, 32] In relation to the ung@hordaland sample, a linear relationship between CRAFFT-score and excessive alcohol consumption, frequency of intoxication and debut of illicit drug use has previously been reported.[32] In the current study, 28.4% of those having tried alcohol were CRAFFT-positive, while 61.1% of those having tried illicit drugs were CRAFFTpositive.

## Mental health problems

Symptoms of depression, anxiety and ADHD were included as the main dependent variables. Depression was assessed using the short version of the Mood and Feelings Questionnaire (SMFQ).[33] The SMFQ comprises 13 items assessing depressive symptoms rated on a 3-point scale. The wordings of the response categories in the Norwegian translation equal the original categories of "Not true", "Sometimes true" and "True". High internal consistency between the items and a strong unidimensionality have been shown in population-based studies,[34] and was recently confirmed in a study based on the sample included in the present study.[35] For the purposes of the current study, depression was defined as a score above the 90<sup>th</sup> percentile of the Total SMFQ-score. It should be noted that term depression as used in the current study does not imply existence of a clinical diagnosis, such as MDD. Also, being a relatively brief self-report questionnaire, the SFMQ does not differentiate between different types of depressive disorders/conditions. The Cronbach's alpha of the SMFQ in the ung@hordaland-sample was 0.91.
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Symptoms of anxiety was identified using the five-item inventory SCARED, which is short-form of the 41-item version screening inventory for anxiety disorders.[36] The short-form of the SCARED has showed similar psychometric properties to the full version.

Symptoms of inattention and hyperactivity were measured using sub-scores from the official Norwegian translation of the Adult ADHD Self-report Scale (ASRS),[37] originally constructed for use in adults, but recently validated among adolescents.[38] ASRS is an 18-item self-report scale, comprising 9 items on a hyperactivity-impulsivity subscale and 9 items on an inattention-subscale. The responses are given on a 5-point scale (ranging from "Never" to "Very often").

#### Statistical analysis

Five different alcohol and drug-related binary variables were constructed as main independent variables: ever consumed alcohol (Yes/No), ever tried illicit drugs (Yes/No), those reporting  $\geq$ 90<sup>th</sup> gender-specific percentile alcohol consumption (Yes/No), intoxication >10 times (Yes/No) and CRAFFT-positive according to the  $\geq$ 2 cutoff (Yes/No). For dependent variables the mental health questionnaires were dichotomised at the 90<sup>th</sup> percentile, identifying those with high sum scores on the ASRS subscales of inattention (raw score range for cut-point 24-36, mean 27.2) and hyperactivity-impulsivity (raw score range for cut-point 20-36, mean 22.9), the SMFQ scale of depression (raw score range for cut-point 15-26, mean 19.3) and the SCARED scale of anxiety (raw score range for cut-point 5-10, mean 6.2).

First, the sample was described in relation to the independent variables, stratified for age and gender. Second, odds ratios were computed using crude logistic regression models for the associations of interest, as well as logistic regression models adjusted for age and gender, and then adjusted for age, gender and SES, with mental health variables both analysed separately, and collectively (i.e. entered into the same logistic regression model). In addition, as previous studies have shown differential age- and gender effects in the development of both mood disorder, anxiety disorder, behaviour disorder and substance use disorders among adolescents,[1] we investigated the potential two-way interaction with age and gender for the alcohol- and drug-related variables in relation to mental health problems. The presence of significant interactions was assessed using likelihood-ratio tests (Wald). When significant interactions were identified, the associations of interest were presented stratified for age or gender. In secondary analyses, we investigated the association between the spectrum of alcohol consumption ("Never tried alcohol", "Non-consumption", "0.1-19.9<sup>th</sup>, "20.0-79.9<sup>th</sup>", "80.0-89.9<sup>th</sup>", "90.0-100<sup>th</sup>") and frequency of intoxication ("Never tried alcohol", "Never intoxicated", "Once", "2-3 times", 4-10

times" and ">10 times") categories separately for all mental health variables, employing age- and genderadjusted logistic regression models with "Never tried" as reference group. All analyses were performed using Stata 12.0.[39] Due to missing information on the frequency of intoxication question, the analyses involving this variable only included n=9,056 participants. Frequency of intoxication was included as an independent variable in order to retain as much information as possible.

## RESULTS

The final sample consisted of N=9,203 participants, where 54.3% were female and the mean age was 17.9 (SD 0.8) years. Table 1 and Figure 1 show the use of alcohol and illicit drugs stratified by age and gender. A total of 20.7% of the girls and 27.1% of the boys had *not* tried alcohol, whereas 8.8% of the girls and 12.5% of the boys *had* tried any illicit drug. Among the participants aged 17, 35.3% had *not* tried alcohol, compared to 16.0% aged 18 or more (p<0.001). Also, ~22% percent of the sample were CRAFFT-positive, indicating some alcohol- or drug-related problem. There was a significant age-trend for all independent variables for both genders, where increasing age was associated with increased proportion of participants having tried alcohol or illicit drugs, as well as reporting alcohol- and drug-related problems (p-values ranging from: 0.002 to <0.001).

Table 2 depicts the crude, age- and gender-adjusted, and age-, gender- and SES-adjusted associations between having tried alcohol, illicit drugs or reporting alcohol- and drug-related problems, in relation to mental health problems . Having ever consumed alcohol was associated with increased odds of all mental health problems in both crude and age- and gender-adjusted models, except for symptoms of anxiety (crude ORs ranging from 1.13 to 2.36). Having ever tried illicit drugs was associated with increased odds of all mental health problems in both crude and age- and gender-adjusted models (crude ORs ranging from 1.33 to 2.38). In relation to alcohol- and drug-related problems, being CRAFFT-positive was associated with increased odds of all mental health problems in both the crude and the age- and gender-adjusted model (crude ORs ranging from 1.68 to 3.24). Excessive alcohol consumption was associated with all mental health problems in both crude and age- and gender-adjusted with all mental health problems in both crude and age- and gender-adjusted with all mental health problems in both crude and age- and gender-adjusted with all mental health problems in both trude and age- and gender-adjusted with all mental health problems in both crude and age- and gender-adjusted model (crude ORs ranging from 1.68 to 3.24). Excessive alcohol consumption was associated with all mental health problems in both crude and age- and gender-adjusted models (crude ORs ranging from 1.61 to 2.70). Frequent intoxication was associated with increased symptoms of depression, inattention and hyperactivity in both regression models, but not with symptoms of anxiety (ORs ranging from 0.95 to 2.08). Further adjustment for self-reported perceived family SES only slightly altered the estimated associations.

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Table 3 shows the age- and gender-adjusted associations between having tried alcohol, illicit drugs or reporting alcohol- and drug-related problems, in relation to mental health problems analysed collectively. Entering all of the mental health variables into the same logistic regression model substantially attenuated the associations of interest, but the pattern of associations were similar to entering them separately.

For the most part, no two-way age- and gender-interactions across use of alcohol and illicit drugs and mental health problems were found; only seven out 40 (17.5%) investigated interactions were statistically significant (data not shown). For age, significant interactions were found with having ever consumed alcohol and frequent intoxication in relation to hyperactivity (p=0.002 and p=0.025, respectively). In the age-stratified analyses having ever tried alcohol was associated with increased levels of hyperactivity for 17-year-olds (OR=2.53, CI95% 1.88-3.43) and 18-year-olds (OR=2.34, CI95% 1.57-3.48), but not for 19-year-olds (OR=0.79, CI95% 0.47-1.34). Frequent intoxication was associated with increased levels of hyperactivity for all age-groups, more strongly among 17- (OR=2.56, CI95% 1.88-3.47) and 18-year-olds (OR=2.54, CI95% 1.93-3.34) than for 19-year-olds (OR=1.55, CI95% 1.16-2.08).

For gender, significant interactions were found with having ever tried illicit drugs and excessive alcohol consumption in relation to symptoms of anxiety (p=0.021 and p<0.001, respectively), excessive alcohol consumption and symptoms of hyperactivity (p=0.021), and with frequent intoxication and excessive consumption and symptoms of depression (p=0.027 and p=0.018, respectively). In the gender-stratified analyses, having ever tried illicit drugs and reporting excessive consumption were associated with increased symptoms of anxiety for boys (OR=2.44, [CI95% 1.57-3.78] and OR=4.36 [CI95% 2.52-7.57], respectively), but not for girls (OR=1.30 [CI95% 0.98-1.73] and OR=1.23 [CI95% 0.87-1.73], respectively). The same pattern was evident for depression, where frequent intoxication was associated with increased symptoms among boys (OR=1.83, CI95%1.32-2.52) but not for girls (OR=1.19, CI95% 0.98-1.44). Excessive alcohol consumption was associated with increased symptoms of hyperactivity and depression, more strongly for boys (OR=3.78 [CI95% 2.65-5.40] and OR=3.57 [CI95% 2.32-5.51], respectively) than girls (OR=2.17 [CI95% 1.60-2.94] and OR=1.89 [CI95% 1.43-2.49], respectively).

Table 4 shows the associations between alcohol consumption and frequency of intoxication categories with never having tried alcohol as reference. For symptoms of inattention, hyperactivity and depression there was a trend towards more symptoms of mental health problems with increasing levels of usual consumption and higher frequency of intoxication (all p<0.001). This trend was not found for symptoms of anxiety where there were no associations with intoxication categories (p-value for trend: 0.679) and alcohol consumption categories

(p-value for trend: 0.061). The only significant difference was found with excessive alcohol consumption (90<sup>th</sup> percentile) compared to never having tried alcohol (OR=1.60, CI95%1.15-2.23).

#### DISCUSSION

#### **Main findings**

In sum, most adolescents aged 17 to 19 had tried alcohol and one-fifth of the girls and one-fourth of the boys had tried some illicit drug. Also, twenty percent of the adolescents reported some alcohol- or drug-related problems, as indicated by 2 or more points on the CRAFFT-questionnaire. There was a clear positive age-trend for having tried both alcohol and illicit drugs for both boys and girls, and debut of alcohol and illicit drug use was associated with mental health problems for both genders. Having tried illicit drugs, excessive alcohol consumption and being CRAFFT-positive were associated with all mental health problems. No J- or U-shaped association was found for alcohol consumption or frequency of intoxication in relation to any of the mental health problems included in this study. There was no clear evidence of any substantial age-, gender- or SES- confounding or modifying effect of age and gender on the estimated association between alcohol- and drug-related measures and mental health problem. Moreover, a closing gender gap on alcohol and drug use was supported.

#### Strengths and limitations

The current study holds several strengths. Firstly, it is a large well-defined population-based sample of adolescents. The sample size enabled the investigation of both main effects and potential interaction effects of age and gender. Secondly, the data collection is recent and allows for an updated and contemporary insight into the current status of alcohol- and drug use, presence of potential alcohol- and drug use problems, and associations with mental health problems. Thirdly, the study included different measures of alcohol- and drug use, as well as the validated CRAFFT-questionnaire for alcohol- or drug-related problems. Fourthly, we included measures of internalizing and externalizing mental health problems, using well-validated questionnaires measuring symptoms of depression and anxiety, as well as hyperactivity and inattention symptoms that are part of the ADHD-diagnosis.

The current study has also some limitations. Firstly, the study is cross-sectional, and this precludes any causal inference of the estimated associations. Previous findings indicate that mental health problems can be a

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consequence of alcohol- and drug use, be a risk factor for alcohol- and drug use or that both are influenced by a common cause.[40-42] Secondly, the age-range included in the present study is limited, and a wider range would enable a more thorough investigation of age-related effects of alcohol and drug use, and probably also increase the chances of finding any moderating effects of age.[43] Thirdly, both data on the independent and dependent variables were self-reported, which could lead bias in reporting, especially underreporting for the independent variables. Fourthly, due to missing information, almost ten percent of the total eligible study population was not included in our analyses. The attrition from the current study could affect generalizability, and result in adolescents attending school being overrepresented. The problem with non-participation in survey research seems unfortunately to be on the rise.[44] Official data show that in 2012, 92% of all adolescents in Norway aged 16-18 attended high school, [45] compared to 98% in the current study. Based on previous research from the former waves of the Bergen Child Study (the nested within the current study), non-participants have also been shown to have more psychological problems than participants.[46] It is therefore likely that the prevalence of mental health problems is underestimated in the current study. This could suggest that the non-responders also have a higher alcohol and drug consumption, which could bias our sample further. This non-participation bias is, however, not likely to reduce the associations of interest substantially.[47] Finally, the assessment of illicit drug does not indicate type of drug tried and frequency, thus limiting conclusions drawn according to the relation between mental health and more specific drugs or consumption patterns. Lastly, the study is based on selfreported questionnaires assessing symptoms of both alcohol and drug problems and mental health problems, and multiple information would have been have been preferable to reduce informant bias and common method bias. Also, self-reported questionnaires do not provide information regarding the existence of a clinical diagnosis, and the lack of a clinical interview in confirming a formal diagnosis is a limitation of the present study.

#### Interpretation

The closing gender gap reported in previous studies was partly confirmed by the present study.[18] Even though more girls in the younger age group had tried alcohol, boys showed a pattern of trying illegal drugs more often and there were more boys with problems related to alcohol and drug use, while there were no differences in patterns of problem drinking and frequency of intoxication. The expected age pattern was confirmed with both more alcohol and drug use and frequent intoxication with increasing age. 18 years is the legal age when adolescents can buy alcohol in Norway, and thus an increase in alcohol use at this age is as expected. In the 19-years-olds more than a third reported drinking to intoxication. Even though this is a lower rate than reported in a

previous population-based Norwegian study,[21] our findings suggest that this negative alcohol use pattern is frequent among Norwegian adolescents.

There was strong evidence for a co-occurrence of alcohol and drug problems and mental health problems among adolescents, consistent with findings in previous studies.[4, 23] While most studies have focused on drug abuse, we also found that even having tried alcohol or drugs in this age group are potential indicators of mental health problems for both genders. Due to the cross-sectional design of this study, we could not investigate direction of causality. But the relationship between alcohol and drug use and mental health problems in adolescence is probably complex and multidirectional.[22, 48] For instance, mental health problems like impulsivity and hyperactivity may possibly render an adolescent susceptible for using alcohol, but these problems may also be exacerbated by alcohol and drug use. There may also be common etiological factors that accounts for both an increased risk of mental health problems and alcohol consumption, such as parental alcohol use or social peer network. While both ADHD and depressive symptoms were consistently related to alcohol and drug use and abuse, the co-occurrence of these disorders may have impacted the association. In line with this the strength of the association was attenuated when including all of the mental health variables, while the pattern was similar and the associations were still significant. Thus, while co-occurrence of symptoms is frequent, both depressive and ADHD symptoms seems to have an independent contribution to the association between mental health and alcohol and drug use.

In general, we found little evidence of a substantial modifying effect of age or gender, which is contrary to some previous studies.[21, 24, 25] We did, however, find weak evidence for a modifying effect of age and gender for specific mental health domains: there was some evidence for an age-differential association between alcohol and drug variables and symptoms of hyperactivity, where having tried alcohol or frequent intoxication were more strongly associated with symptoms of hyperactivity for younger adolescents. This could be related to a lower level of cognitive and emotional control in early adolescence compared to at an older age,[43] increasing the susceptibility both to alcohol use, drinking to intoxication and mental health problems.

We also found a gender-differential association between alcohol and drug variables and symptoms of internalizing mental health problems, where boys showed more consistent association between alcohol and drug variables and symptoms of anxiety and depression than girls. Interestingly, we found that, among girls, excessive alcohol consumption was not associated with anxiety and frequent intoxication was not associated with depression. This is contrary to results from a previous Finnish study which found stronger associations between heavy drinking and psychosocial dysfunction among girls than among boys.[25] One factor that may have

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impacted on this relationship is the higher rate of depressive symptoms in girls,[35] and this may be understood in light of the "gender paradox" where the gender with the lowest prevalence rate will be more severely affected.[49] The interpretation of the findings concerning interactions should, however, be done with caution as the number of tested interactions (N=40) were high, and only 17.5% were significant. Future studies should therefore investigate the reported interactions in other study samples.

In the secondary analyses, we found no indication of any curvilinear relationship between alcohol consumption and frequency of intoxication in relation to mental health problems. This is contrary to previous findings,[5-9] predominantly from adult populations, and may be an indication of a differential association between alcohol habits and mental health problems in adolescents versus adults. Also, it may be an indication of cohort-effects, as there is a tendency for lower levels of alcohol consumption and more abstention among adolescents in recent years.[50, 51] Thus, as the abstention group grows, the group also includes more individuals who identify with being abstinent, who have a social network and actively chooses abstinence due to an increased focus on education, career or sports.[52]

Overall, there may be multiple reasons for discrepancies between our findings and previous studies. The recency of our data collection in comparison to a decade or so older comparison studies might have impacted the results, especially with regards to the pattern of drug and alcohol use. Also, the assessment of mental health differs across studies, with most other studies using brief general mental health measures.

#### **Clinical implications**

The high rate of alcohol and drug use in underaged adolescents, 10 percent having tried illegal substances, as well as a pattern of negative alcohol and drug use characterised by frequent intoxication and drinking large quantities of alcohol, together represent a serious public health concern that emphasizes a need of preventive interventions. The strong co-occurrence of mental health and alcohol and drug related problems suggests that assessment of alcohol and drug use and abuse should become an integral part of mental health assessment and interventions, especially in adolescents with depression and ADHD.

In summary, it seems evident that both alcohol and drug debut, and the frequency and intensity of alcohol consumption are important and relevant indicators of mental health problems among adolescents. Investigation of the causal nature between alcohol/drug abuse and mental health problems, as well as the access to health care and utilisation of health care among adolescents who use alcohol and drugs in hazardous ways should be addressed in future studies.

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### **COMPETING INTERESTS**

The authors declare that they have no competing interests.

## **AUTHORS' CONTRIBUTIONS**

JCS conceived the initial concept, and performed the initial analyses and literature search. JCS, BS, MH interpreted the initial analyses and wrote the first draft. AJL, KMS and RJ reviewed and revised the initial analyses and first draft, and suggested further analyses, as well as revisions of the first draft. The subsequent drafts were reviewed by all authors, and all authors read and approved the final manuscript. All authors contributed to the revision of the manuscript after peer-review.

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### DATA SHARING STATEMENT

Data for research projects from the population-based ung@hordaland-study may be made available at request from Regional Centre for Child and Youth Mental Health and Child Welfare, Uni Health, Uni Research, Bergen, Norway.

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# REFERENCES

- Merikangas, K.R., et al., *Lifetime Prevalence of Mental Disorders in U.S. Adolescents: Results from the National Comorbidity Survey Replication–Adolescent Supplement* (NCS-A). Journal of the American Academy of Child and Adolescent Psychiatry, 2010. 49(10): p. 980-989.
- 2. Ormel, J., et al., *Mental health in Dutch adolescents: a TRAILS report on prevalence, severity, age of onset, continuity and co-morbidity of DSM disorders.* Psychological Medicine, 2014. **FirstView**: p. 1-16.
- 3. Teesson, M., T. Slade, and K. Mills, *Comorbidity in Australia: Findings of the 2007 National Survey of Mental Health and Wellbeing*. Australian and New Zealand Journal of Psychiatry, 2009. **43**(7): p. 606-614.
- 4. Costello, E.J., T.D. Armstrong, and A. Erkanli, *Report on the developmental epidemiology of comorbid psychiatric and substance use disorders*. 2000, National Institute on Drug Abuse.
- 5. Rodgers, B., et al., *Non-linear relationships in associations of depression and anxiety with alcohol use.* Psychological Medicine, 2000. **30**(2): p. 421-32.
- 6. Rodgers, B., et al., *Risk factors for depression and anxiety in abstainers, moderate drinkers and heavy drinkers.* Addiction, 2000. **95**(12): p. 1833-1845.
- 7. Caldwell, T.M., et al., *Patterns of association between alcohol consumption and symptoms of depression and anxiety in young adults*. Addiction, 2002. **97**: p. 583-594.
- 8. Skogen, J.C., et al., *Anxiety and depression among abstainers and low-level alcohol consumers. The Nord-Trøndelag Health Study.* Addiction, 2009. **104**(9): p. 1519-1529.
- 9. Pape, H. and T. Hammer, *Sober adolescence Predictor of psychosocial maladjustment in young adulthood?* Scandinavian Journal of Psychology, 1996. **37**: p. 362-277.
- 10. Upmark, M., J. Möller, and A. Romelsjö, *Longitudinal, population-based study of self reported alcohol habits, high levels of sickness absence, and disability pensions.* Journal of Epidemiology and Community Health, 1999. **53**: p. 223-229.
- 11. Hellandsjø Bu, E.T., et al., *Teenage alcohol and intoxication debut: The impact of family socialization factors, living area and participation in organized sports.* Alcohol and Alcoholism, 2002. **37**(1): p. 74-80.
- 12. Brecht, M.-L., et al., *Methamphetamine use behaviors and gender differences*. Addictive behaviors, 2004. **29**(1): p. 89-106.
- 13. Westermeyer, J. and A.E. Boedicker, *Course, severity, and treatment of substance abuse among women versus men.* American Journal of Drug and Alcohol Abuse, 2000. **26**(4): p. 523-535.
- 14. Hser, Y.-I., et al., *Gender differences in treatment outcomes over a three-year period: A path model analysis.* Journal of Drug Issues, 2004. **34**: p. 419-440.
- 15. Hernandez-Avila, C.A., B.J. Rounsaville, and H.R. Kranzler, *Opioid-, cannabis- and alcohol-dependent women show more rapid progression to substance abuse treatment.* Drug and Alcohol Dependence, 2004. **74**(3): p. 265-272.
- 16. Haas, K., *Relationship of gender to licit and illicit drug use among adolescents*. Chrestomathy: Annual Review of Undergraduate Research at the College of Charleston, 2004. **3**: p. 92-100.
- 17. Opland, E.A., K.C. Winters, and R.D. Stinchfield, *Examining gender differences in drug-abusing adolescents*. Psychology of Addictive Behaviors, 1995. **9**(3).
- 18. Keyes, K.M., B.F. Grant, and D.S. Hasin, *Evidence for a closing gender gap in alcohol use, abuse, and dependence in the United States population.* Drug Alcohol Depend, 2008. **93**(1-2): p. 21-9.

 Schwinn, T.M., S.P. Schinke, and D.N. Trent, Substance use among late adolescent urban youths: Mental health and gender influences. Addictive behaviors, 2010. 35(1): p. 30-34.

- 20. Armstrong, T.D. and E.J. Costello, *Community studies on adolescent substance use, abuse, or dependence and psychiatric comorbidity.* Journal of Consulting and Clinical Psychology, 2002. **70**(6): p. 1224-1239.
- 21. Strandheim, A., et al., *Alcohol intoxication and mental health among adolescents a population review of 8983 young people, 13-19 years in North-Trøndelag, Norway: the Young-HUNT Study.* Child and Adolescent Psychiatry and Mental Health, 2009. **3**(18).
- 22. Trim, R.S., et al., *The relation between adolescent substance use and young adult internalizing symptoms: Findings from a high-risk longitudinal sample.* Psychology of Addictive Behaviors, 2007. **21**(1): p. 97-107.
- 23. McCarty, C.A., et al., *Developmental consistency in associations between depressive symptoms and alcohol use in early adolescence*. Journal of studies on Alcohol and Drugs, 2012. **73**: p. 444-453.
- 24. Verdurmen, J., et al., *Alcohol use and mental health in adolescents: Interactions with age and gender Findings from the Dutch 2001 Health Behaviour in School-Aged Children Survey.* Journal of Studies on Alcohol and Drugs, 2005. **66**(5): p. 605-609.
- 25. Laukkanen, E.R., et al., *Heavy drinking is associated with more severe psychosocial dysfunction among girls than boys in finland.* The Journal of adolescent health : official publication of the Society for Adolescent Medicine, 2001. **28**(4): p. 270-277.
- 26. Theunissen, M.-J., M. Jansen, and A. van Gestel, *Are mental health and binge drinking associated in Dutch adolescents? Cross-sectional public health study.* BMC Research Notes, 2011. **4**(1): p. 100.
- 27. Torikka, A., et al., *Depression, drinking, and substance use among 14- to 16-year-old Finnish adolescents.* Nord J Psychiatry, 2001. **55**(5): p. 351-7.
- McLeod, J.D., R. Uemura, and S. Rohrman, *Adolescent Mental Health, Behavior Problems, and Academic Achievement*. Journal of Health and Social Behavior, 2012. 53(4): p. 482-497.
- 29. Skogen, J.C., et al., *Concurrent validity of the CAGE questionnaire. The Nord-Trøndelag Health Study.* Addictive behaviors, 2011. **36**(4): p. 302-307.
- 30. Knight, J.R., et al., *A new brief screen for adolescent substance abuse*. Archives of Pediatrics & Adolescent Medicine, 1999. **153**: p. 591-596.
- 31. Knight, J.R., et al., *Validity of brief alcohol screening tests among adolescents: A comparison of the AUDIT, POSIT, CAGE, and CRAFFT.* Alcoholism: Clinical and Experimental Research, 2003. **27**(1): p. 67-73.
- 32. Skogen, J.C., et al., *Psychometric properties and concurrent validity of the CRAFFT among Norwegian adolescents. Ung@hordaland, a population-based study.* Addictive behaviors, 2013(Ahead of print).
- 33. Thapar, A. and P. McGuffin, *Validity of the shortened Mood and Feelings Questionnaire in a community sample of children and adolescents: a preliminary research note.* Psychiatry Res., 1998. **81**(2): p. 259-68.
- 34. Sharp, C., I.M. Goodyer, and T.J. Croudace, *The Short Mood and Feelings Questionnaire (SMFQ): a unidimensional item response theory and categorical data factor analysis of self-report ratings from a community sample of 7-through 11-yearold children.* J Abnorm Child Psychol, 2006. **34**(3): p. 379-91.
- 35. Lundervold, A.J., et al., Symptoms of depression as reported by Norwegian adolescents on the Short Mood and Feelings Questionnaire. Frontiers in Psychology, 2013. 4.

1		
2	26	
3	36.	Birmaher, B., et al., <i>Psychometric properties of the screen for child anxiety related</i>
4		emotional disorders scale (SCARED): A replication study. Journal of the American
5		Academy of Child and Adolescent Psychiatry, 1999. <b>38</b> (10): p. 1230-1236.
6 7	37.	Kessler, R.C., et al., Validity of the World Health Organization Adult ADHD Self-
/		report Scale (ASRS) Screener in a representative sample of health plan members.
8		International Journal of Methods in Psychiatric Research, 2007. 16(2): p. 52-65.
9	38.	Adler, L.A., et al., Preliminary examination of the reliability and concurrent validity
10		of the attention-deficit/hyperactivity disorder self-report scale v1.1 symptom checklist
11		to rate symptoms of attention-deficit/hyperactivity disorder in adolescents Journal of
12		Child and Adolescent Psychonharmacology 2012 <b>27</b> (3): n 238 44
13	20	State Corn Statistical Software: Palagea 12.0, 2012. State Corneration: College
14	59.	StataCorp, Sidiisticul Software. Release 12.0. 2012, Stata Corporation. Conege
15	40	
10	40.	Tapert, S.F., L. Caldwell, and C. Burke, Alcohol and the adolescent brain: Human
18		<i>studies</i> . Alcohol Research & Health, 2004. <b>28</b> (4): p. 205-212.
10	41.	Chassin, L., et al., A longitudinal study of children of alcoholics: Predicting young
20		adult substance use disorders, anxiety, and depression. Journal of Abnormal
20		Psychology, 1999. 108(1): p. 106-119.
22	42.	Sher, K.J., et al., <i>Characteristics of children of alcoholics: Putative risk factors</i> ,
23		substance use and abuse, and psychopathology. Journal of Abnormal Psychology.
24		1991 <b>100</b> (4): p 427-448
25	43	Willoughby T et al <i>Examining the link between adolescent brain development and</i>
26	15.	risk taking from a social developmental perspective Brain and Cognition 2013
27		<b>93</b> (2): n 215 222
28	4.4	<b>03</b> (5). p. 513-525.
29	44.	Morton, S.M.B., et al., In the 21st Century, what is an acceptable response rate?
30		Australian and New Zealand Journal of Public Health, 2012. <b>36</b> (2): p. 106-108.
31	45.	The Directorate of Integration and Diversity (IMDi), [Education participation in high
32		schools]. 2012, http://www.imdi.no/no/Fakta-og-statistikk/Fakta-og-
33		<u>statistikk/Utdanning/?tab=chr</u> .
34	46.	Stormark, K.M., et al., Predicting nonresponse bias from teacher ratings of mental
35		health problems in primary school children. J Abnorm Child Psychol, 2008. 36(3): p.
36		411-9.
37	47.	Knudsen, A.K., et al., The health status of nonparticipants in a population-based
38		health study. The Hordaland Health Study American Journal of Epidemiology 2010
39		<b>172</b> (11): n 1306-14
40	18	O'Neil K A B T Conner and B C Kendall Internalizing disorders and substance
41	40.	O Nell, K.A., D. I. Colliel, and I.C. Kendall, International autor and implications for
42		use disorders in youin. Comorbially, risk, temporal order, and implications for $(1, 1, 2, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3,$
43	40	intervention. Clinical Psychology Review, 2011. <b>31</b> (1): p. 104-112.
44	49.	Loeber, R. and K. Keenan, Interaction between Conduct Disorder and Its Comorbid -
45		<i>Effects of Age and Gender</i> . Clinical Psychology Review, 1994. 14(6): p. 497-523.
46	50.	Danielsson, AK., et al., Alcohol use, heavy episodic drinking and subsequent
47		problems among adolescents in 23 European countries: does the prevention paradox
40		<i>apply</i> ? Addiction, 2012. <b>107</b> (1): p. 71-80.
49 50	51.	Hibell, B., et al., Summary: The 2007 ESPAD report among students in 35 countries.
51		(2009), European Monitoring Centre for Drugs and Drug Addiction: Luxembourg.
52	52.	Hoel, S., et al., Adolescent alcohol use, psychological health, and social integration.
53		Scandinavian Journal of Public Health 2004 <b>32</b> n 361-367
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55		
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57		
58		
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3 4 5 6 (girls n=4,995, boys n=4,208). 7 Born 1995 Total sample 8 % (95% CI) % (95% CI) 9 Never consumed alcohol (% yes) 10 11 Girls 31.4 (29.4-33.5) 20.7 (19.6-21.8) 12 Boys 27.1 (25.7-28.4) 39.7 (37.4-42.1) 13 14 Ever tried illicit drugs (% yes) 15 8.8 (8.1-9.6) 6.1 (5.1-7.3) Girls 16 17 12.5 (11.5-13.5) Boys 10.8 (8.5-11.2) 18 90<sup>th</sup> percentile alcohol consumption<sup>a</sup> (% 19 yes) 20 6.0 (4.8-7.4) Girls 7.7 (6.9-8.6) 21 5.9 (4.6-7.6) 7.4 (6.5-8.4) Boys 22 23 Frequent intoxication (>10; % yes)<sup>b</sup> 24 25 Girls 20.5 (19.4-21.6) 10.6 (9.3-12.0) 26 10.7 (9.3-12.3) Boys 21.3 (20.1-22.6) 27 28 CRAFFT positive (≥2; % yes) 29 Girls 23.3 (22.2-24.5) 20.0 (18.3-21.8) 30 31 20.3 (19.1-21.5) 15.8 (14.1-17.6) Boys 32 95 confidence intervals in parentheses 33 <sup>a</sup>Only includes those who have tried alcohol (n=7,031). 34 <sup>b</sup>Only includes those with valid response on frequency of intoxication (n=9,056). 35 CRAFFT: Screening tool for identification of problematic alcohol and drug use among adolescents. 36 37 38

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47 48 10 Table 1: Patterns of drinking and drug use, and problematic alcohol and drug use (CRAFFT positive) in adolescents born from 1993-1995, stratified for gender. N=9,203

Born 1994

% (95% CI)

19.4 (17.6-21.4)

26.4 (24.2-28.7)

8.2 (7.0-9.6)

10.2 (8.8-11.9)

7.7 (6.4-9.2)

5.9(5.5-8.5)

19.7 (17.9-21.7)

21.9 (19.9-24.1)

22.4 (20.5-24.4)

19.3 (17.4-21.4)

Born 1993

% (95% CI)

6.3 (5.1-7.7)

7.6 (6.1-9.4)

13.7 (11.9-15.6)

18.5 (16.3-21.0)

9.7 (8.2-11.5)

9.6 (7.9-11.6)

36.1 (33.6-38.8)

37.1 (34.2-40.1)

29.5 (27.1-32.1)

29.0 (26.3-31.8)

Odds ratio for age

trend (95% CI)

0.42 (0.38-0.47)

0.41 (0.37-0.45)

1.57 (1.39-1.77)

1.37 (1.22-1.54)

1.30 (1.13-1.51)

1.31 (1.10-1.55)

2.20 (2.01-2.41)

2.21 (2.00-2.44)

1.29 (1.19-1.40)

1.47 (1.34-1.62)

p-value

<.001

<.001

<.001

<.001

<.001

=.002

<.001

<.001

<.001

<.001

 Table 2: Logistic regression models of use of alcohol and illicit drugs as risk factor for mental health problems. Mental health variables analysed separately. N=9,203 (girls n=4,995, boys n=4,208).

	ASRS Inattention Odds ratio (OR) (95CI)	ASRS Hyperactivity OR (95CI)	SMFQ OR (95CI)	SCARED OR (95CI)
Ever consumed alcohol				
Crude	2.36 (1.90-2.93)	2.08 (1.68-2.58)	1.73 (1.43-2.09)	1.13 (0.94-1.37)
Adjusted for age and gender	2.21 (1.77-2.76)	2.07 (1.66-2.58)	1.64 (1.35-1.99)	1.03 (0.84-1.25)
Adjusted for age, gender and socioeconomic status (SES) <b>Ever tried illicit drugs</b>	2.15 (1.72-2.69)	2.03 (1.62-2.53)	1.59 (1.31-1.93)	0.99 (0.81-1.21)
Crude	2.38 (1.98-2.87)	2.26 (1.86-2.74)	1.69 (1.39-2.05)	1.33 (1.05-1.68)
Adjusted for age and gender	2.49 (2.06-3.02)	2.36 (1.94-2.89)	1.98 (1.62-2.42)	1.55 (1.23-1.98)
Adjusted for age, gender and SES 90 <sup>th</sup> percentile alcohol consumption <sup>a</sup>	2.42 (2.00-2.93)	2.31 (1.90-2.82)	1.91 (1.56-2.34)	1.50 (1.18-1.91)
Crude	1.97 (1.55-2.51)	2.70 (2.14-3.40)	2.19 (1.74-2.75)	1.61 (1.21-2.14)
Adjusted for age and gender	1.96 (1.54-2.50)	2.76 (2.19-3.48)	2.27 (1.79-2.88)	1.64 (1.22-2.19)
Adjusted for age, gender and SES Intoxication >10 times <sup>b</sup>	1.98 (1.55-2.53)	2.77 (2.19-3.49)	2.29 (1.80-2.90)	1.64 (1.22-2.20)
Crude	1.83 (1.56-2.15)	2.08 (1.77-2.45)	1.30 (1.10-1.52)	0.95 (0.78-1.16)
Adjusted for age and gender	1.80 (1.52-2.12)	2.18 (1.84-2.59)	1.36 (1.15-1.61)	0.97 (0.79-1.20)
Adjusted for age, gender and SES CRAFFT-positive (≥2)	1.80 (1.52-2.13)	2.17 (1.83-2.58)	1.35 (1.14-1.61)	0.96 (0.78-1.18)

Crude	3.15 (2.71-3.66)	3.24 (2.78-3.78)	2.84 (2.46-3.28)	1.68 (1.42-1.99)
Adjusted for age and gender	3.06 (2.63-3.56)	3.22 (2.76-3.77)	2.84 (2.45-3.29)	1.62 (1.38-1.94)
Adjusted for age, gender and SES	3.01 (2.59-3.51)	3.18 (2.72-3.71)	2.80 (2.41-3.25)	1.59 (1.34-1.90)

95 confidence intervals in parentheses.

 <sup>a</sup>Only includes those who have tried alcohol (n=7,031).

<sup>b</sup>Only includes those with valid response on frequency of intoxication (n=9,056).

CRAFFT: Screening tool for identification of problematic alcohol and drug use among adolescents.

ASRS: Adult ADHD Self-report Scale, screening inventory for attention-deficit-hyperactivity disorder. ASRS Inattention: the inattention sub-scale of the ASRS (9 items). ASRS

Hyperactivity: the hyperactivity-impulsivity subscale of the ASRS (nine items).

SMFQ: The short version of the Mood and Feelings Questionnaire, screening inventory for symptoms of depression and depressive disorders.

SCARED: five-item screening inventory for symptoms of anxiety and anxiety disorders.

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Table 3: Table 2: Logistic regression models of use of alcohol and illicit drugs as risk factor for mental health problems, adjusted for age and gender. Mental health variables analysed collectively. N=9,203 (girls n=4,995, boys n=4,208).

	ASRS inattention Odds ratio (OR) (95CI)	ASRS Hyperactivity Odds ratio (OR) (95CI)	SMFQ Odds ratio (OR) (95CI)	SCARED Odds ratio (OR) (95CI)
Ever consumed alcohol	1.83 (1.44-2.32)	1.71 (1.35-2.16)	1.45 (1.17-1.78)	0.79 (0.64-0.98)
Ever tried illicit drugs	1.87 (1.50-2.32)	1.74 (1.40-2.17)	1.48 (1.18-1.86)	1.12 (0.86-1.46)
90 <sup>th</sup> percentile alcohol consumption <sup>a</sup>	1.24 (0.94-1.63)	2.30 (1.78-2.98)	1.88 (1.45-2.44)	1.18 (0.86-1.62)
Intoxication >10 <sup>b</sup>	1.45 (1.20-1.75)	1.95 (1.62-2.35)	1.18 (0.98-1.43)	0.79 (0.63-0.99)
CRAFFT-positive (≥2)	1.97 (1.66-2.34)	2.27 (1.91-2.69)	2.17 (1.84-2.55)	0.99 (0.82-1.21)

Significant associations in bold.

95 confidence intervals in parentheses.

<sup>a</sup>Only includes those who have tried alcohol (n=7,031).

<sup>b</sup>Only includes those with valid response on frequency of intoxication (n=9,056).

CRAFFT: Screening tool for identification of problematic alcohol and drug use among adolescents.

ASRS: Adult ADHD Self-report Scale, screening inventory for attention-deficit-hyperactivity disorder. ASRS Inattention: the inattention sub-scale of the ASRS (9 items). ASRS

Hyperactivity: the hyperactivity-impulsivity subscale of the ASRS (nine items).

SMFQ: The short version of the Mood and Feelings Questionnaire, screening inventory for symptoms of depression and depressive disorders.

SCARED: five-item screening inventory for symptoms of anxiety and anxiety disorders.

Table 4: Logistic regression models across alcohol consumption and frequency of intoxication categories, "Never tried alcohol" as reference. Adjusted for age and gender (N=9,203 for alcohol consumption and N=9,056 for frequency of intoxication).

	ASRS Inattention	ASRS Hyperactivity	SMFQ	SCARED	
	Odds ratio (OR) (95CI)	OR (95CI)	OR (95CI)	OR (95CI)	
Categories	Alcohol consumption categories as independent variable				
Never consumed alcohol	1.00 (Reference)	1.00 (Reference)	1.00 (Reference)	1.00 (Reference	
Non-consumption	1.90	1.46	1.43	1.01	
	(1.45-2.48)	(1.10-1.93)	(1.12-1.82)	(0.78-1.30)	
0.1-19.9 <sup>th</sup> percentile	1.61	1.69	1.44	1.00	
	(1.19-2.17)	(1.25-2.27)	(1.11-1.88)	(0.75-1.32)	
20.0-79.9 <sup>th</sup> percentile	2.15	1.99	1.51	0.90	
	(1.69-2.74)	(1.56-2.55)	(1.21-1.88)	(0.72-1.14)	
80.0-89.9 <sup>th</sup> percentile	3.77	3.56	2.21	1.39	
	(2.75-5.17)	(2.59-4.91)	(1.62-3.02)	(0.99-1.96)	
90.0-100 <sup>th</sup> percentile	4.16	5.28	3.47	1.60	
	(3.05-5.68)	(3.91-7.12)	(2.60-4.62)	(1.15-2.23)	
p-value for trend	p<0.001	p<0.001	p<0.001	p=0.061	
Categories	Frequency of	f intoxication categorie	es as independent v	ariable	
Never consumed alcohol	1.00 (Reference)	1.00 (Reference)	1.00 (Reference)	1.00 (Reference	
Never intoxicated	1.65	1.35	1.30	1.10	
	(1.24-2.20)	(1.00-1.81)	(1.00-1.67)	(0.85-1.43)	
Once	1.71	2.06	1.55	1.02	
	(1.23-2.38)	(1.50-2.82)	(1.16-2.08)	(0.74-1.39)	
2-3 times	2.00	1.62	1.61	0.88	
	(1.51-2.63)	(1.21-2.15)	(1.26-2.07)	(0.67-1.16)	
4-10 times	2.49	2.05	1.85	1.05	
	(1.90-3.26)	(1.55-2.72)	(1.44-2.36)	(0.81-1.38)	
>10 times	3.14	3.36	1.95	0.98	
	(2.43-4.05)	(2.61-4.33)	(1.54-2.47)	(0.76-1.27)	
p-value for trend	p<0.001	p<0.001	p<0.001	p=0.679	

Significant associations in comparison to "Never tried alcohol"-category in bold.

ASRS: Adult ADHD Self-report Scale, screening inventory for attention-deficit-hyperactivity disorder. ASRS Inattention: the inattention sub-scale of the ASRS (9 items). ASRS Hyperactivity: the hyperactivity-impulsivity subscale of the ASRS (nine items).

SMFQ: The short version of the Mood and Feelings Questionnaire, screening inventory for symptoms of depression and depressive disorders.

SCARED: five-item screening inventory for symptoms of anxiety and anxiety disorders.

# **Figure legends**

Figure 1: Proportion of participants having never consumed alcohol across gender-stratified age groups (N=9,203, girls n=4,995, boys n=4,208). Error bars represent 95% confidence intervals.

Figure 2: Proportion of participants having ever tried illicit drugs across gender-stratified age groups (N=9,203, girls n=4,995, boys n=4,208). Error bars represent 95% confidence intervals.



## ABSTRACT

**Objectives:** The use of alcohol and drugs is prevalent among adolescents, but <u>too</u> little is known about the association between debut of alcohol and drug<u>use</u>, <u>problematic</u> use and concurrent mental health. The aim of the study was to investigate the cross-sectional association between debut of any alcohol or drug use and alcohol- and drug-related problems, and mental health. We also wanted to examine potential interactions between gender and age, and alcohol- and drug-related <u>exposuresvariables</u>.

Design: Cross-sectional study.

Setting: Population-based sample of Norwegian adolescents.

**Participants:** Data stem from the large population-based ung@hordaland study (N=9,203), where all adolescents aged 16-18 years living in Hordaland county (Norway) were invited to participate. The main exposures independent variables were debut of alcohol and drug use, alcohol consumption, and presence of alcohol and drug problems as measured by CRAFFT.

**Outcomes:** The outcomesdependent variables were self-reported symptoms of anxiety, depression, inattention and hyperactivity. <u>Statistical analyses included logistic regression models.</u>

**Results:** Both debut of alcohol and drug use were associated with symptoms of depression, inattention and hyperactivity (crude ORs 1.69-2.38, Ps<0.001), while only debut of drug use was associated with increased symptoms of anxiety (OR=1.33, CI95%1.05-1.68, P=0.017). Alcohol- and drug-related problems as measured by CRAFFT were associated with all mental health problems (crude ORs 1.68-3.24, Ps<0.001). There was little evidence of any substantial age- or gender confounding on the estimated associations between alcohol- and drug-related measures and the mental health outcomesproblems.

**Conclusion:** Early debut of alcohol and drug use and drug problems are consistently associated with more symptoms of mental health problems, indicating that these factors are an important general indicator of mental health in adolescence.

### STRENGTHS AND LIMITATIONS OF THIS STUDY

- This study employed a large well-defined population-based sample of adolescents.
- The data employed in this study is from a recent data collection.
- This study included several different measures of alcohol- and drug use, as well as measures of both internalizing and externalizing problems.
- The cross-sectional design of this study precluded any causal inference.
- This study only had a limited age-range; a wider range would enable a more thorough investigation of age-related effects of alcohol and drug use.

### BACKGROUND

burden attributed to alcohol and illicit drug use is considerable worldwide.[1, 2] Alcohol and illicit drug use tend to begin and escalate during adolescence and early adulthood.[3, 4] Early life alcohol and drug are predictive of later use and abuse,[5] and for alcohol it has been shown that a 10% time delay in debut would lead to a 35% decrease in the subsequent expected alcohol consumption .[6] DrugPrevious studies of adolescents have found that a considerable proportion with one mental disorder also meets the criteria for at least one other mental disorder, including substance-use disorder.[1-3] Furthermore, drug use and abuse have been related to a range of mental health problems, most consistently to externalizing problems such as those associated with conduct disorder (CD) and attention-deficit-hyperactivity disorder (ADHD), [7]), [4] while the increased risk for anxiety disorders are lower. Also of note, several previous studies in adult populations have found that there is a J- or U-shaped relationship between alcohol consumption and mental health outcomes, [8-12][5-9] where both abstainers and high-level alcohol consumers have increased risk for mental health problems. Several explanations of this curvilinear relationship have been suggested: moderate alcohol consumers may directly benefit from a protective effect on mental health from alcohol, or their drinking habits may be a mere reflection of their psychological wellbeing.[9][6] Also, in cultures where alcohol use is the norm, moderate consumers may be more well-adjusted than individuals at the extremes of the consumption spectrum.[13][10] To date, few studies have investigated the relationship between the full spectrum of alcohol consumption (from non-debut to high-level alcohol consumption) and symptoms of mental health problems in a population-based sample of adolescents

Boys have an earlier debut of both alcohol consumption and intoxication according to most studies,[3] while the gender-pattern for onset of drug use is more unclear.Boys have an earlier debut of both alcohol consumption and intoxication according to most studies,[11] while the gender-pattern for onset of drug use is more unclear.[14-19][12-17] The established gender differences with males using more alcohol and drugs seems, however, to be closing [20] with recent studies not finding a higher rate of drug use among males.[21] If the relation between alcohol and drug use and mental health is gender specific is less clear. A lack of gender difference has been documented in population based studies where the use of alcohol and drugs is associated with worse mental health. The established gender differences with males using more alcohol and drugs seems,

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however, to be closing [18] with recent studies not finding a higher rate of drug use among males.[19] Whether the relation between alcohol and drug use and mental health is gender specific is less clear. A lack of gender difference has been documented in population-based studies where the use of alcohol and drugs is associated with worse mental health.[22-25][20-23] In contrast, some studies have reported a modifying effect of gender and age in relation to the association between alcohol use and mental health.[23, 26, 27][21, 24, 25] For instance, a recent study investigating the association between alcohol intoxications and mental health problems among 13-19-year-olds found that high level alcohol consumption was associated with conduct and attention problems for both genders, but only the girls who reported frequent intoxicationsintoxication reported more symptoms of anxiety and depression.[23][21] On the other hand, Verdurmen and colleagues reported a modifying effects of age on the association between alcohol use and mental health, but found no evidence of any modifying effects of gender.[26][24] The association between alcohol use and mental health is, however, not quite consistent, as a Dutch study did not confirm the expected relation between binge drinking and mental health problems among 16-18-years-olds in a population based study.[<del>28]</del>[26]. Furthermore, most of the previous studies have focused on drug and alcohol use, whereas less is known about the relationship between mental health and debut in adolescence.

In general, the mental health problems show a consistent gender pattern in adolescence, with girls having a higher rate of depression and internalizing problems, and boys a higher rate of externalising disorders. [1] Less is known whether the associations between the drug use and mental health is gender-specific, but there are some indications that co-occurrence of other mental health problems are higher for girls than boys.[7][4] However, in a sample of late adolescent urban youth gender did not have a moderating effect on the relationship between substance abuse and mental health, and a Finnish study did not support a gender specific association between depression and substance use.[21] These studies are rather small, and the results may have been hampered by lack of statistical power. It is also uncertain if the co-occurrence is specifically related to alcohol or drug dependence or[19, 27]

Also, it is uncertain if the co-occurrence is specifically related to alcohol or drug dependence or if it is also evident for mere alcohol or drug use.[7][4] Overall, there is still a need for further investigation of the association between early alcohol and drug use and mental health in a general population of adolescents.[29][28] To that end, the aim of the present study was to

 Describe the debut of alcohol and drug use, alcohol use patterns and potential drug problems among Norwegian adolescents using data from a recent population-based study.

To investigate the cross-sectional relationship between a) alcohol and illicit drug use and b) alcohol and drug problems and mental health among adolescents. We also investigated any interactions with age and gender on the associations of interest, as well as the association between the full spectrum of alcohol consumption and <u>bingefrequency of</u> drinking <u>frequencyto intoxication</u> in relation to mental health.

#### METHODS

ii)

#### Study population

This study employed information from the ung@hordaland-survey ("youth@hordaland")study of adolescents in the county of Hordaland in western Norway. All adolescentadolescents born 1993-1995 living in Hordaland were invited to participate. (n=19,430), and 10,220 participated, yielding a participation rate of 53%. Ung@hordaland is a population-based study, and carried out in close collaboration with the county council administration. The majority fourth wave of participants responded to the questionnaires during school hourslongitudinal population-based study "Bergen Child Study" is also nested within ung@hordaland. The general aim of the ung@hordaland-survey was to assess mental health, lifestyle, school performance and healthservice use in adolescents, with a special emphasis on the prevalence of mental health problems. The data was collected during January-February 2012. Adolescents in upper secondary education received information per email, and one school hour was allocated for them to complete the questionnaire at school. Those not in school received information by mail to their home addresses. In addition, mental health facilities, juvenile detention centres and other institutions were contacted and the adolescents had the opportunity to participate. The questionnaire was web-based and covered a broad range of mental health issues, daily life functioning, use of health and social services, and demographic background variables, and a request for permission to obtain school data, and to link the information with national registries and parental questionnaires. Uni Health and Hordaland County Council are collaborating partners in conducting the study. Upon participation electronic informed consent was obtained. The study was approved by The Regional Committee for Medical Research Ethics in Western Norway. After complete descriptionHordaland County is generally regarded representative of Norway Formatted: English (U.S.) as a whole, comprising both the second largest city of Norway (Bergen), as well as large rural areas. Official statistics of the general population (not just adolescents) show that Hordaland County does not deviate in any

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arameters, including both socio-demographic indicators and several health indicators	<b>Formatted:</b> English (U.S.)
The current study is based on the first version of data files released in May 2012, with a total of 10,220	
articipants. Based on list-wise deletion of those with missing information on both exposures independent and	
utcomesdependent variables, n=1,017 were excluded, leaving 9,203 in the final study sample. Those with	
nissing information on the exposures and the outcomes, variables of interest were slightly younger than those	
vith valid information on the included variables (mean difference 0.12 years, p<.001) and more likely to be	
nales (61.6% versus 45.7% in the analysed sample, p<.001)	Formatted: Font: Not Italic
Demographic information	
nformation about age and gender was obtained on all participants. Furthermore, the participants indicated the	
erceived family socioeconomic status (SES) as i) "about the same as others" (67.9%), ii) "better than others"	
25.1%) or iii) "worse than others" (7.1%), with n=172 missing responses recoded to "about the same as others"	-
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dentification of alcohol problems and illicit drug use	
elf-reported debut, frequency and amount of alcohol consumption and illicit drug use were included as main	
xposureindependent variables. We included a binary measure of alcohol debut "Have you ever tried alcohol?"	
Yes/No), and illicit drug use debut: "Have you ever tried hash, marihuana or other narcotic substances?"	
Yes/No). Out of the total study sample, n=11 individuals did not answer the question on illicit drug use, and	
ney were recoded to the "No" category. Alcohol use was measured using the self-reported units of beer, cider,	
vine, spirits and illegally distilled spirits usually consumed during the last 14 days. Based on this information	
bout consumption and alcohol debut, a summed variable including the followingon gender-specific	
istribution <u>distributions</u> was constructed: "Never tried", "Non-consumption" (if reported consumption was "0"),	,
0.1-19.9 <sup>th</sup> ", "20.0- <del>39.9<sup>th</sup>", "40.0-59.9<sup>th</sup>", "60.0-</del> 79.9 <sup>th</sup> ", "80.0-89.9 <sup>th</sup> ", "90.0-100 <sup>th</sup> ". Excessive alcohol	
onsumption was defined as above the 90 <sup>th</sup> percentile sum score.[11, 30] Frequent binge drinking[8, 29]	
requent intoxication (binge drinking) was defined as drinking so much that one was clearly intoxicated more	
nan 10 times, based on the question: "Have you ever consumed so much alcohol that you were clearly	
ntoxicated (drunk)?", with five categories ranging from "No, never" to "Yes, more than 10 times". There were	
=147 missing responses on the binge drinkingintoxication item, leaving a sample of N=9,056 for this measure.	

CRAFFT.[31][30] CRAFFT was specifically designed to identify alcohol and illicit drug problems among adolescents,[31][30] and have been shown to have acceptable sensitivity and specificity at a cut-off of >=2 in international studies.[32, 33][31, 32] In relation to the ung@hordaland sample, a linear relationship between CRAFFT-score and excessive alcohol consumption, binge drinkingfrequency of intoxication and debut of illicit drug use has previously been reported.[33][32] In the current study, 28.4% of those having tried alcohol were CRAFFT-positive, while 61.1% of those having tried illicit drugs were CRAFFT-positive.

#### Mental health problems

Symptoms of depression, anxiety and ADHD were included as the main dependent variables. Depression was assessed using the short version of the Mood and Feelings Questionnaire (SMFQ).[34][33] The SMFQ comprises 13 items assessing depressive symptoms rated on a 3-point scale. The wordings of the response categories in the Norwegian translation equal the original categories of "Not true", "Sometimes true" and "True". High internal consistency between the items and a strong unidimensionality have been shown in population-based studies,[35][34] and was recently confirmed in a study based on the sample included in the present study.[36][35] For the purposes of the current study, depression was defined as a score above the 90<sup>th</sup> percentile of the Total SMFQ-score. It should be noted that term depression as used in the current study does not imply existence of a clinical diagnosis, such as MDD. Also, being a relatively brief self-report questionnaire, the SFMQ does not differentiate between different types of depressive disorders/conditions. The Cronbach's alpha of the SMFQ in the ung@hordaland-sample was 0.91.

Symptoms of inattention and hyperactivity were measured using sub-scores from the official Norwegian translation of the Adult ADHD Self-report Scale (ASRS),[38][37] originally constructed for use in adults, but recently validated among adolescents.[39][38] ASRS is an 18-item self-report scale, comprising 9 items on a hyperactivity-impulsivity subscale and 9 items on an inattention-subscale. The responses are given on a 5-point scale (ranging from "Never" to "Very often").

#### Statistical analysis

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Five different alcohol and drug-related binary variables were constructed as main exposures: Everindependent variables: ever consumed alcohol (Yes/No), ever tried illicit drugs (Yes/No), those reporting  $\geq$ 90<sup>th</sup> genderspecific percentile alcohol consumption (Yes/No), binge drinkingintoxication >10 times (Yes/No) and CRAFFTcasenesspositive according to the  $\geq$ 2 cutoff (Yes/No). For outcomedependent variables the mental health questionnaires were dichotomised at the 90<sup>th</sup> percentile, identifying those with high sum scores on the ASRS subscales of inattention (raw score range for cut-point 24-36, mean 27.2) and hyperactivity-impulsivity<sub>7</sub> (raw score range for cut-point 20-36, mean 22.9), the SMFQ scale of depression (raw score range for cut-point 15-26, mean 19.3) and the SCARED scale of anxiety- (raw score range for cut-point 5-10, mean 6.2).

First, the sample was described in relation to exposures stratified for age and gender. Second, odds ratios were computed using simple logistic regression models for the associations between exposures and outcomes, as well as logistic regression models adjusted for age and gender. In addition, First, the sample was described in relation to the independent variables, stratified for age and gender. Second, odds ratios were computed using crude logistic regression models for the associations of interest, as well as logistic regression models adjusted for age and gender, and then adjusted for age, gender and SES, with mental health variables both analysed separately, and collectively (i.e. entered into the same logistic regression model). In addition, as previous studies have shown differential age- and gender effects in the development of both mood disorder, anxiety disorder, behaviour disorder and substance use disorders among adolescents,[1] we investigated the potential two-way interaction with age and gender for the alcohol- and drug-related exposuresvariables in relation to mental health problems. The presence of significant interactions was assessed using likelihood-ratio tests (Wald), and when). When significant interactions were identified, the associations of interest were presented stratified for age or gender. In secondary analyses, we investigated the association between the spectrum of alcohol consumption ("Never tried alcohol", "Non-consumption", "0.1-19.9<sup>th</sup>, "20.0-39.9<sup>th</sup>," "40.0-59.9<sup>th</sup>, "60.0-79.9<sup>th</sup>", "80.0-89.9<sup>th</sup>", "90.0-100<sup>th</sup>") and binge drinkingfrequency of intoxication ("Never tried alcohol", "Never intoxicated", "Once", "2-3 times", 4-10 times" and ">10 times") categories separately for all mental health variables, employing age- and gender-adjusted logistic regression models with "Never tried" as reference group. All analyses were performed using Stata 12.0.[40][39] Due to missing information on the frequency of intoxication question, the analyses involving this variable only included n=9,056 participants. Frequency of intoxication was included as an independent variable in order to retain as much information as possible.

#### RESULTS

The final sample consisted of N=9,203 participants, where 54.3% were female and the mean age was 17.9 (SD 0.8) years. Table 1 and Figure 1 show the use of alcohol and illicit drugs stratified by age and gender. A total of 20.7% of the girls and 27.1% of the boys had *not* tried alcohol, whereas 8.8% of the girls and 12.5% of the boys *had* tried any illicit drug. Among the participants aged 17, 35.3% had *not* tried alcohol, compared to 16.0% aged 18 or more (p<0.001). Also, ~22% percent of the sample waswere CRAFFT-casespositive, indicating some alcohol- or drug-related problem. There was a significant age-trend for all exposures independent variables for both genders, where increasing age was associated with increased proportion of participants having tried alcohol or illicit drugs, as well as reporting alcohol- and drug-related problems (p-values ranging from: 0.002 to <0.001).

Table 2 depicts the crude-and, age- and gender-adjusted, and age-, gender- and SES-adjusted associations between having tried alcohol, illicit drugs or reporting alcohol- and drug-related problems, in relation to mental health problems. Having ever consumed alcohol was associated with increased odds of all mental health problems in both crude and age- and gender-adjusted models, except for symptoms of anxiety (crude ORs ranging from 1.13 to 2.36). Having ever tried illicit drugs was associated with increased odds of all mental health problems in both crude and age- and gender-adjusted models (crude ORs ranging from 1.33 to 2.38). In relation to alcohol- and drug-related problems, <u>being</u>\_CRAFFT-<u>easenesspositive</u> was associated with increased odds of all mental health problems in both the crude and the age- and gender-adjusted model (crude ORs ranging from 1.68 to 3.24). Excessive alcohol consumption was associated with all mental health problems in both crude and age- and gender-adjusted models (crude ORs ranging from 1.68 to 3.24). Excessive alcohol consumption was associated with all mental health problems in both crude and age- and gender-adjusted models (crude ORs ranging from 1.61 to 2.70). <u>Binge-drinking more</u> than 10 timesFrequent intoxication was associated with increased symptoms of depression, inattention and hyperactivity in both regression models, but not with symptoms of anxiety (ORs ranging from 0.95 to 2.08).

Table 3 shows the p-values of the Table 3 shows the age- and gender-adjusted associations between having tried alcohol, illicit drugs or reporting alcohol- and drug-related problems, in relation to mental health problems analysed collectively. Entering all of the mental health variables into the same logistic regression model substantially attenuated the associations of interest, but the pattern of associations were similar to entering them separately.

For the most part, no two-way age- and gender-interactions across use of alcohol and illicit drugs and mental health problems. For the most part, no interactions were found; only seven out 40 (17.5%) investigated

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interactions were statistically significant-<u>(data not shown)</u>. For age, significant interactions were found with having ever consumed alcohol and frequent <u>binge drinkingintoxication</u> in relation to hyperactivity (p=0.002 and p=0.025, respectively). In the age-stratified analyses having ever tried alcohol was associated with increased levels of hyperactivity for 17-year-olds (OR=2.53, CI95% 1.88-3.43) and 18-year-olds (OR=2.34, CI95% 1.57-3.48), but not for 19-year-olds (OR=0.79, CI95% 0.47-1.34). Frequent <u>binge drinkingintoxication</u> was associated with increased levels of hyperactivity for all age-groups, <u>but</u>-more strongly among 17<sub>-</sub> (OR=2.56, CI95% 1.88-3.47) and 18-year-olds (OR=2.54, CI95% 1.93-3.34) than for 19-year-olds (OR=1.55, CI95% 1.16-2.08).

For gender, significant interactions were found with having ever tried illicit drugs and excessive alcohol consumption in relation to symptoms of anxiety (p=0.021 and p<0.001, respectively), excessive alcohol consumption and symptoms of hyperactivity (p=0.021), and with binge drinkingfrequent intoxication and excessive consumption and symptoms of depression (p=0.027 and p=0.018, respectively). In the genderstratified analyses, having ever tried illicit drugs and reporting excessive consumption were associated with increased symptoms of anxiety for boys (OR=2.44, [CI95% 1.57-3.78] and OR=4.36 [CI95% 2.52-7.57], respectively), but not for girls (OR=1.30 [CI95% 0.98-1.73] and OR=1.23 [CI95% 0.87-1.73], respectively). The same pattern was evident for depression, where frequent binge drinkingintoxication was associated with increased symptoms inamong boys (OR=1.83, CI95%1.32-2.52) but not for girls (OR=1.19, CI95% 0.98-1.44). Excessive alcohol consumption was associated with increased symptoms of hyperactivity and depression, more strongly for boys (OR=3.78 [CI95% 2.65-5.40] and OR=3.57 [CI95% 2.32-5.51], respectively) than girls (OR=2.17 [CI95% 1.60-2.94] and OR=1.89 [CI95% 1.43-2.49], respectively).

Table 4 shows the associations between alcohol consumption and <u>binge drinkingfrequency of</u> <u>intoxication</u> categories with never having tried alcohol as reference. For symptoms of inattention, hyperactivity and depression there was a trend towards more symptoms of mental health problems with increasing levels of usual consumption and higher frequency for binge drinkingof intoxication (all p<0.001). This trend was not found for symptoms of anxiety where there were no associations with <u>binge drinkingintoxication</u> categories (pvalue for trend: 0.679) and alcohol consumption categories (p-value for trend: 0.058061). The only significant difference was found with excessive alcohol consumption (90<sup>th</sup> percentile) compared to never having tried alcohol (OR=1.60, CI95%1.15-2.23).

#### DISCUSSION

#### Main findings

In sum, most adolescents aged 17 to 19 had tried alcohol and one-fifth of the girls and one-fourth of the boys had tried some illicit drug. Also, twenty percent of the adolescents reported some alcohol- or drug-related problems, as indicated by 2 or more points on the CRAFFT-questionnaire. There was a clear positive age-trend for having tried both alcohol and illicit drugs for both boys and girls, and debut of alcohol and illicit drug use was associated with mental health problems for both genders. Having tried illicit drugs, excessive alcohol consumption and <u>being</u>.CRAFFT-casenesspositive were associated with all mental health problems. No J- or U-shaped association was found for alcohol consumption or <u>binge drinkingfrequency of intoxication</u> in relation to any of the mental health problems included in this study. –There was <u>weakno clear</u> evidence of any substantial age–or\_s gender-or SES-confounding or interactionmodifying effect of age and gender on the estimated association between alcohol- and drug-related measures and mental health problem. Moreover, a closing gender gap on alcohol and drug use was supported.

#### Strengths and limitations

The current study holds several strengths. Firstly, it is a large well-defined population-based sample of adolescents. The sample size enabled the investigation of both main effects and potential interaction effects of age and gender. Secondly, the data collection is recent which and allows for an updated and contemporary insight into the current status of alcohol- and drug use, presence of potential alcohol- and drug use problems, and associations with mental health problems. Thirdly, the study included different measures of alcohol- and drug use, as well as the validated CRAFFT-questionnaire for alcohol- or drug-related problems. Fourthly, we included measures of internalizing and externalizing mental health problems, using well-validated questionnaires measuring symptoms of depression and anxiety, as well as hyperactivity and inattention symptoms that are part of the ADHD-diagnosis.

The current study has also some limitations. Firstly, the study is cross-sectional, and this precludes any causal inference of the estimated associations. Previous findings indicate that mental health problems can be a consequence of alcohol- and drug use, be a risk factor for alcohol- and drug use or that both are influenced by a common cause. [41-43][40-42] Secondly, the age-range included in the present study is limited, and a wider range would enable a more thorough investigation of age-related effects of alcohol and drug use, and probably also increase the chances of finding any moderating effects of age.[44] Thirdly, both data on exposures and outcomes were self-reported, which could lead to common methods bias and bias in reporting, especially

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underreporting for the exposures.[43] Thirdly, both data on the independent and dependent variables were selfreported, which could lead bias in reporting, especially underreporting for the independent variables. Fourthly, due to missing information, almost ten percent of the total eligible study population was not included in our analyses. The attrition from the current study could affect generalizability, and result in adolescents attending school being overrepresented. Although no data on non-responders were available in the current wave of the Bergen Child Study, previous research from the former waves of the Bergen Child Study have shown that nonparticipants tend The problem with non-participation in survey research seems unfortunately to be on the rise.[44] Official data show that in 2012, 92% of all adolescents in Norway aged 16-18 attended high school, [45] compared to 98% in the current study. Based on previous research from the former waves of the Bergen Child Study (the nested within the current study), non-participants have also been shown to have more psychological problems than participants, [45][46] It is therefore likely that the prevalence of mental health problems is underestimated in the current study. This could suggest that the non-responders also have a higher alcohol and drug consumption, which could bias our sample further. This non-participation bias is, however, not likely to reduce the associations of interest substantially. [46] [47] Finally, the assessment of illicit drug does not indicate type of drug tried and frequency, thus limiting conclusions drawn according to the relation between mental health and more specific drugs or consumption patterns. Lastly, the study is based on self-reported questionnaires assessing symptoms of both alcohol and drug problems and mental health problems, and multiple information would have been have been preferable to reduce informant bias and common method bias. Also, self-reported questionnaires do not provide information regarding the existence of a clinical diagnosis, and the lack of a clinical interview in confirming a formal diagnosis is a limitation of the present study.

#### Interpretation

The closing gender gap reported in previous studies; was partly confirmed by the present study.[20][18] Even though more girls in the younger age group had tried alcohol, boys showed a pattern of trying illegal drugs more often and there were more boys with problems related to alcohol and drug use, while there were no differences in patterns of problem drinking and binge drinking.frequency of intoxication. The expected age pattern was confirmed with both more alcohol and drug use and binge drinkingfrequent intoxication with increasing age. 18 years is the legal age when adolescents can buy alcohol in Norway, and thus an increase in alcohol use at this age is as expected. In the 19-years-olds more than a third reported binge drinking to intoxication. Even though

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this is a lower rate than reported in a previous population-based Norwegian study-,[23][21] our findings suggest that this negative alcohol use pattern is highly-frequent among Norwegian adolescents.

There was strong evidence for a co-occurrence of alcohol and drug problems and mental health problems among adolescents, consistent with findings in previous studies. [7, 25][4, 23] While most studies have focused on drug abuse, we also found that even having tried alcohol or drugs in this age group are potential indicators of mental health problems for both genders. Due to the cross-sectional design of this study, we could not investigate direction of causality. But the relationship between alcohol and drug use and mental health problems in adolescence is probably complex and multidirectional. [24, 47][22, 48] For instance, mental health problems like impulsivity and hyperactivity may possibly render an adolescent susceptible for using alcohol, but these problems may also be exacerbated by alcohol and drug use. There may also be common etiological factors that accounts for both an increased risk of mental health problems and alcohol consumption, such as parental alcohol use or social peer network. While both ADHD and depressive symptoms were consistently related to alcohol and drug use and abuse, the co-occurrence of these disorders may have impacted the association. In line with this the strength of the association was attenuated when including all of the mental health variables, while the pattern was similar and the associations were still significant. Thus, while co-occurrence of symptoms is frequent, both depressive and ADHD symptoms seems to have an independent contribution to the association between mental health and alcohol and drug use.

In general, we found little evidence of a substantial modifying effect of age or gender, which is contrary to some previous studies. [23, 26, 27][21, 24, 25] We did, however, find weak evidence for a modifying effect of age and gender for specific mental health domains: there was some evidence for an age-differential association between alcohol and drug exposures variables and symptoms of hyperactivity, where having tried alcohol or frequent binge drinking intoxication were more strongly associated with symptoms of hyperactivity for younger adolescents. This could be related to a lower level of cognitive and emotional control in early adolescence compared to at an older age, [44] increasing the susceptibility both to alcohol use, binge drinking and mental health problems. [43] increasing the susceptibility both to alcohol use, drinking to intoxication and mental health problems.

We also found a gender-differential association between alcohol and drug <u>exposuresvariables</u> and symptoms of internalizing mental health problems, where boys showed more consistent association between <u>exposuresalcohol and drug variables</u> and symptoms of anxiety and depression than girls. Interestingly, we found that, among girls, excessive alcohol consumption was not associated with anxiety and frequent <u>binge</u>

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drinkingintoxication was not associated with depression. This is contrary to results from a previous Finnish study which found stronger associations between heavy drinking and psychosocial dysfunction among girls than among boys.[27] One factor that may have impacted on this relation is the higher rate of depressive symptoms in girls,[25] One factor that may have impacted on this relationship is the higher rate of depressive symptoms in girls,[36][35] and this may be understood in light of the "gender paradox" where the gender with the lowest prevalence rate will be more severely affected.[48]-[49] The interpretation of the findings concerning interactions should, however, be done with caution as the number of tested interactions (N=40) were high, and only 17.5% were significant. Future studies should therefore investigate the reported interactions in other study samples.

In the secondary analyses, we found no indication of any curvilinear relationship between alcohol consumption and frequency of binge drinkingintoxication in relation to mental health problems. This is contrary to previous findings, [8-12][5-9] predominantly from adult populations, and may be an indication of a differential association between alcohol habits and mental health problems in adolescentadolescents versus in an adult populationadults. Also, it may be an indication of cohort-effects, as there is a tendency for lower levels of alcohol consumption and more abstention among adolescents in recent years. [4, 49][50, 51] Thus, as the abstention group grows, the group also includes more individuals who identify with being abstinent, who have a social network and actively chooses abstinence due to an increased focus on education, career or sports [5052]

Overall, there may be multiple reasons for discrepancies between our findings and previous studies. The recency of our data collection in comparison to a decade or so older comparison studies might have impacted the results, especially with regards to the pattern of drug and alcohol use. Also, the assessment of mental health differs across studies, with most other studies using brief general mental health measures.

#### **Clinical implications**

The high rate of alcohol and drug use in <u>underageunderaged</u> adolescents, 10 percent having tried illegal substances, as well as a pattern of negative alcohol and drug use characterised by <u>binge\_frequent intoxication and</u> drinking <u>and-large</u> quantities of alcohol, together represent a serious public health concern that emphasizes a need of preventive interventions. The strong co-occurrence of mental health and alcohol and drug related problems <u>suggestsuggests</u> that assessment of alcohol and drug use and abuse should become an integral part of mental health assessment and interventions, especially in <u>adolescentadolescents</u> with depression and ADHD.

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<text> alcohol consumption are important and relevant indicators of mental health problems among adolescents. Investigation of the causal nature between alcohol-and /drug use, abuse and mental health problems, as well as the access to health care and utilisation of health care among adolescents who use alcohol and drugs in hazardous ways should be addressed in future studies.

#### **COMPETING INTERESTS**

The authors declare that they have no competing interests.

### **AUTHORS' CONTRIBUTIONS**

JCS conceived the initial concept, and <u>performed</u> the initial analyses and literature search. JCS, BS, MH interpreted the initial analyses and wrote the first draft. AJL, KMS and RJ reviewed and revised the initial analyses and first draft, and suggested further analyses, as well as revisions of the first draft. The subsequent drafts were reviewed by all authors, and all authors read and approved the final manuscript. <u>All authors</u> contributed to the revision of the manuscript after peer-review.

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1. Rehm, J., et al., Alcohol use disorders in EU countries and Norway: An overview of
the epidemiology. European Neuropsychopharmacology, 2005. 15: p. 377-388.
2. Degenhardt, L. and W. Hall, Extent of illicit drug use and dependence, and their
contribution to the global burden of disease. The Lancet, 2012. 379(9810): p. 55-70.
3. Hellandsjø Bu, E.T., et al., Teenage alcohol and intoxication debut: The impact of
family socialization factors, living area and participation in organized sports. Alcohol
and Alcoholism, 2002. <b>37</b> (1): p. 74-80.
4. Hibell, B., et al., Summary: The 2007 ESPAD report among students in 35 countries.
2009, European Monitoring Centre for Drugs and Drug Addiction: Luxembourg.
5. Hingson, R.W., T. Heeren, and M.R. Winter, Age at drinking onset and alcohol
dependence: Age at onset, duration, and severity. Archives of Pediatrics & Adolescent
Medicine, 2006. 160(7): p. 739-746.
6. Pedersen, W. and A. Skrondal, Alcohol consumption debut: Predictors and
consequences. Journal of Studies on Alcohol, 1998. 59(1): p. 32-42.
7. Costello, E.J., T.D. Armstrong, and A. Erkanli, <i>Report on the developmental</i>
epidemiology of comorbid psychiatric and substance use disorders. 2000, National
Institute on Drug Abuse.
8. Rodgers, B., et al., Non linear relationships in associations of depression and anxiety
with alcohol use. Psychological Medicine, 2000. 30(2): p. 421-32.
9. Rodgers, B., et al., Risk factors for depression and anxiety in abstainers, moderate
drinkers and heavy drinkers. Addiction, 2000. 95(12): p. 1833-1845.
10. Caldwell, T.M., et al., Patterns of association between alcohol consumption and
symptoms of depression and anxiety in young adults. Addiction, 2002. 97: p. 583-594.
11. Skogen, J.C., et al., Anxiety and depression among abstainers and low level alcohol
consumers. The Nord Trøndelag Health Study. Addiction, 2009. 104(9): p. 1519-1529.
17

# **BMJ Open**

<del>12.</del>	Pape, H. and T. Hammer, <i>Sober adolescence – Predictor of psychosocial</i>
	maladjustment in young adulthood? Scandinavian Journal of Psychology, 1996. 37:
	<del>362-277.</del>
<del>13.</del>	Upmark, M., J. Möller, and A. Romelsjö, Longitudinal, population based study of s
	reported alcohol habits, high levels of sickness absence, and disability pensions.
	Journal of Epidemiology and Community Health, 1999. 53: p. 223-229.
<del>14.</del>	Brecht, M. L., et al., Methamphetamine use behaviors and gender differences.
	Addictive behaviors, 2004. 29(1): p. 89-106.
<del>15.</del>	Westermeyer, J. and A.E. Boedicker, Course, severity, and treatment of substance
	abuse among women versus men. American Journal of Drug and Alcohol Abuse,
	<del>2000. <b>26</b>(4): p. 523-535.</del>
<del>16</del>	Hser, Y. I., et al., Gender differences in treatment outcomes over a three year perie
	A path model analysis. Journal of Drug Issues, 2004. 34: p. 419-440.
<del>17.</del>	Hernandez Avila, C.A., B.J. Rounsaville, and H.R. Kranzler, Opioid, cannabis and
	alcohol-dependent women show more rapid progression to substance abuse treatme
	Drug and Alcohol Dependence, 2004. 74(3): p. 265-272.
<del>18.</del>	Haas, K., Relationship of gender to licit and illicit drug use among adolescents.
	Chrestomathy: Annual Review of Undergraduate Research at the College of
	Charleston, 2004. 3: p. 92-100.
<del>19.</del>	Opland, E.A., K.C. Winters, and R.D. Stinchfield, Examining gender differences in
	drug abusing adolescents. Psychology of Addictive Behaviors, 1995. 9(3).
<del>20.</del>	Keyes, K.M., B.F. Grant, and D.S. Hasin, Evidence for a closing gender gap in
	alcohol use, abuse, and dependence in the United States population. Drug Alcohol
	<del>Depend, 2008. <b>93</b>(1-2); p. 21-9.</del>

21.	Schwinn, T.M., S.P. Schinke, and D.N. Trent, Substance use among late adolescent
	urban youths: Mental health and gender influences. Addictive behaviors, 2010. 35(1):
	<del>p. 30-34.</del>
22.	Armstrong, T.D. and E.J. Costello, Community studies on adolescent substance use,
	abuse, or dependence and psychiatric comorbidity. Journal of Consulting and Clinical
	Psychology, 2002. 70(6): p. 1224-1239.
23.	Strandheim, A., et al., Alcohol intoxication and mental health among adolescents a
	population review of 8983 young people, 13-19 years in North Trøndelag, Norway:
	the Young HUNT Study. Child and Adolescent Psychiatry and Mental Health, 2009.
	<del>3(18).</del>
<del>24</del>	Trim, R.S., et al., The relation between adolescent substance use and young adult
	internalizing symptoms: Findings from a high risk longitudinal sample. Psychology of
	Addictive Behaviors, 2007. 21(1): p. 97-107.
<del>25.</del>	McCarty, C.A., et al., Developmental consistency in associations between depressive
	symptoms and alcohol use in early adolescence. Journal of studies on Alcohol and
	Drugs, 2012. <b>73</b> : p. 444-453.
<del>26.</del>	Verdurmen, J., et al., Alcohol use and mental health in adolescents: Interactions with
	age and gender Findings from the Dutch 2001 Health Behaviour in School Aged
	Children Survey. Journal of Studies on Alcohol and Drugs, 2005. 66(5): p. 605-609.
27.	Laukkanen, E.R., et al., Heavy drinking is associated with more severe psychosocial
	dysfunction among girls than boys in finland. The Journal of adolescent health :
	official publication of the Society for Adolescent Medicine, 2001. 28(4): p. 270-277.
<del>28.</del>	Theunissen, M. J., M. Jansen, and A. van Gestel, Are mental health and binge
	drinking associated in Dutch adolescents? Cross-sectional public health study. BMC
	Research Notes, 2011. 4(1): p. 100.
### **BMJ Open**

<del>29.</del>	McLeod, J.D., R. Uemura, and S. Rohrman, Adolescent Mental Health, Behavior
	Problems, and Academic Achievement. Journal of Health and Social Behavior, 2012
	<del>53(4): p. 482-497.</del>
<del>30.</del>	- Skogen, J.C., et al., Concurrent validity of the CAGE questionnaire. The Nord-
	Trondelag Health Study. Addictive behaviors, 2011. 36(4): p. 302-307.
<del>31.</del>	Knight, J.R., et al., A new brief screen for adolescent substance abuse. Archives of
	Pediatrics & Adolescent Medicine, 1999. 153: p. 591-596.
<del>32.</del>	Knight, J.R., et al., Validity of brief alcohol screening tests among adolescents: A
	comparison of the AUDIT, POSIT, CAGE, and CRAFFT. Alcoholism: Clinical and
	Experimental Research, 2003. 27(1): p. 67-73.
<del>33.</del>	Skogen, J.C., et al., Psychometric properties and concurrent validity of the CRAFF1
	among Norwegian adolescents. Ung@hordaland, a population based study. Addicti
	behaviors, 2013(Ahead of print).
<del>34.</del>	Thapar, A. and P. McGuffin, Validity of the shortened Mood and Feelings
	Questionnaire in a community sample of children and adolescents: a preliminary
	research note. Psychiatry Res., 1998. 81(2): p. 259-68.
<del>35.</del>	Sharp, C., I.M. Goodyer, and T.J. Croudace, <i>The Short Mood and Feelings</i>
	Questionnaire (SMFQ): a unidimensional item response theory and categorical date
	factor analysis of self report ratings from a community sample of 7 through 11 year
	old children. J Abnorm Child Psychol, 2006. 34(3): p. 379-91.
<del>36.</del>	Lundervold, A.J., et al., Symptoms of depression as reported by Norwegian
	adolascents on the Short Mood and Feelings Questionnaire Frontiers in Develop

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<del>37.</del>	Birmaher, B., et al., Psychometric properties of the screen for child anxiety related
	emotional disorders scale (SCARED): A replication study. Journal of the American
	Academy of Child and Adolescent Psychiatry, 1999. 38(10): p. 1230-1236.
<del>38.</del>	Kessler, R.C., et al., Validity of the World Health Organization Adult ADHD Self-
	report Scale (ASRS) Screener in a representative sample of health plan members.
	International Journal of Methods in Psychiatric Research, 2007. 16(2): p. 52-65.
<del>39.</del>	Adler, L.A., et al., Preliminary examination of the reliability and concurrent validity
	of the attention deficit/hyperactivity disorder self report scale v1.1 symptom checklist
	to rate symptoms of attention deficit/hyperactivity disorder in adolescents. Journal of
	Child and Adolescent Psychopharmacology, 2012. 22(3): p. 238-44.
4 <del>0.</del>	-StataCorp, Statistical Software: Release 12.0. 2012, Stata Corporation: College
	Station, TX.
41.	-Tapert, S.F., L. Caldwell, and C. Burke, Alcohol and the adolescent brain: Human
	studies. Alcohol Research & Health, 2004. 28(4): p. 205-212.
4 <u>2.</u>	Chassin, L., et al., A longitudinal study of children of alcoholics: Predicting young
	adult substance use disorders, anxiety, and depression. Journal of Abnormal
	Psychology, 1999. 108(1): p. 106-119.
4 <del>3.</del>	Sher, K.J., et al., Characteristics of children of alcoholics: Putative risk factors,
	substance use and abuse, and psychopathology. Journal of Abnormal Psychology,
	<del>1991. <b>100</b>(4): p. 427-448.</del>
44.	Willoughby, T., et al., Examining the link between adolescent brain development and
	risk taking from a social developmental perspective. Brain and Cognition, 2013.
	<b>83</b> (3): p. 315-323.

### **BMJ Open**

45.	Stormark, K.M., et al., Predicting nonresponse bias from teacher ratings of menta
	health problems in primary school children. J Abnorm Child Psychol, 2008. 36(3)
	4 <del>11-9.</del>
4 <del>6.</del>	Knudsen, A.K., et al., The health status of nonparticipants in a population based
	health study: The Hordaland Health Study. American Journal of Epidemiology, 20
	<del>172(11): р. 1306-14.</del>
47.	O'Neil, K.A., B.T. Conner, and P.C. Kendall, Internalizing disorders and substan
	use disorders in youth: Comorbidity, risk, temporal order, and implications for
	intervention. Clinical Psychology Review, 2011. 31(1): p. 104-112.
4 <del>8.</del>	- Loeber, R. and K. Keenan, Interaction between Conduct Disorder and Its Comorl
	Effects of Age and Gender. Clinical Psychology Review, 1994. 14(6): p. 497-523.
4 <del>9.</del>	Danielsson, A. K., et al., Alcohol use, heavy episodic drinking and subsequent
	problems among adolescents in 23 European countries: does the prevention para
	apply? Addiction, 2012. 107(1): p. 71-80.
<del>50.</del>	Hoel, S., et al., Adolescent alcohol use, psychological health, and social integration
	Scandinavian Journal of Public Health, 2004. <b>32</b> : p. 361–367.

**BMJ Open** 



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1	Merikangas K R et al Lifetime Prevalence of Mental Disorders in US Adolescents.
	Results from the National Comorbidity Survey Replication–Adolescent Supplement
	(NCS-A). Journal of the American Academy of Child and Adolescent Psychiatry,
	<u>2010.</u> <b>49</b> (10): p. 980-989.
2.	Ormel, J., et al., Mental health in Dutch adolescents: a TRAILS report on prevalence,
	severity, age of onset, continuity and co-morbidity of DSM disorders. Psychological
	Medicine, 2014. FirstView: p. 1-16.
<u>3.</u>	Teesson, M., T. Slade, and K. Mills, Comorbidity in Australia: Findings of the 2007
	National Survey of Mental Health and Wellbeing. Australian and New Zealand Journal
	<u>of Psychiatry, 2009. 43(7): p. 606-614.</u>
4.	Costello, E.J., T.D. Armstrong, and A. Erkanli, <i>Report on the developmental</i>
	epidemiology of comorbid psychiatric and substance use disorders. 2000, National
_	Institute on Drug Abuse.
<u>5.</u>	Rodgers, B., et al., Non-linear relationships in associations of depression and anxiety
~	with alcohol use. Psychological Medicine, 2000. <b>30</b> (2): p. 421-32.
<u>6.</u>	<u>Rodgers, B., et al., Risk factors for depression and anxiety in abstainers, moderate</u>
7	arinkers and heavy arinkers. Addiction, 2000. 95(12): p. 1833-1845.
1.	Caldwell, 1.M., et al., Patterns of association between alcohol consumption and sumptoms of depression and arrists in young adults. Addiction, 2002, <b>07</b> : p. 582-504
8	Symptoms of depression and dantassion among abstainars and low layal alcohol
0.	consumers The Nord-Trandelag Health Study Addiction 2009 104(9): p 1519-1529
9	Pane H and T Hammer Sober adolescence - Predictor of psychosocial
2.	maladiustment in young adulthood? Scandinavian Journal of Psychology 1996 37 p
	362-277.
10.	Upmark, M., J. Möller, and A. Romelsjö, Longitudinal, population-based study of self
	reported alcohol habits, high levels of sickness absence, and disability pensions.
	Journal of Epidemiology and Community Health, 1999. 53: p. 223-229.
<u>11.</u>	Hellandsjø Bu, E.T., et al., Teenage alcohol and intoxication debut: The impact of
	family socialization factors, living area and participation in organized sports. Alcohol
	and Alcoholism, 2002. 37(1): p. 74-80.
<u>12.</u>	Brecht, ML., et al., Methamphetamine use behaviors and gender differences.
10	<u>Addictive behaviors, 2004. <b>29</b>(1): p. 89-106.</u>
<u>13.</u>	Westermeyer, J. and A.E. Boedicker, <i>Course, severity, and treatment of substance</i>
	abuse among women versus men. American Journal of Drug and Alcohol Abuse,
14	<u>2000.</u> <b>20</b> (4). p. 525-555.
<u>14.</u>	A nath model analysis Journal of Drug Issues 2004 34: p. 410 440
15	<u>A puil model analysis</u> . Journal of Drug Issues, 2004. <b>54</b> . p. 419-440. Hernandez-Avila C A B I Rounsaville and H R Kranzler Opioid- cannabis- and
<u>10.</u>	alcohol-dependent women show more rapid progression to substance abuse treatment
	Drug and Alcohol Dependence. 2004. <b>74</b> (3): p. 265-272.
16.	Haas, K., Relationship of gender to licit and illicit drug use among adolescents.
	Chrestomathy: Annual Review of Undergraduate Research at the College of
	Charleston, 2004. 3: p. 92-100.
<u>17.</u>	Opland, E.A., K.C. Winters, and R.D. Stinchfield, Examining gender differences in
	drug-abusing adolescents. Psychology of Addictive Behaviors, 1995. 9(3).
<u>18.</u>	Keyes, K.M., B.F. Grant, and D.S. Hasin, Evidence for a closing gender gap in
	alcohol use, abuse, and dependence in the United States population. Drug Alcohol
	<u>Depend, 2008.</u> <b>93</b> (1-2): p. 21-9.

## BMJ Open

- / •	<i>urban vouths: Mental health and gender influences.</i> Addictive behaviors 2010 <b>35</b> (1):
	p. 30-34.
20.	Armstrong, T.D. and E.J. Costello, <i>Community studies on adolescent substance use</i> ,
	abuse, or dependence and psychiatric comorbidity. Journal of Consulting and Clinical
	Psychology, 2002. <b>70</b> (6): p. 1224-1239.
21.	Strandheim, A., et al., Alcohol intoxication and mental health among adolescents - a
	population review of 8983 young people, 13-19 years in North-Trøndelag, Norway:
	the Young-HUNT Study. Child and Adolescent Psychiatry and Mental Health, 2009.
	<u><b>3</b>(18).</u>
22.	Trim, R.S., et al., The relation between adolescent substance use and young adult
	internalizing symptoms: Findings from a high-risk longitudinal sample. Psychology of
	<u>Addictive Behaviors, 2007. 21(1): p. 97-107.</u>
23.	McCarty, C.A., et al., <i>Developmental consistency in associations between depressive</i>
	symptoms and alcohol use in early adolescence. Journal of studies on Alcohol and
<b>1</b> 4	Drugs, 2012. 15: p. 444-455.
24.	verduimen, J., et al., Alconol use and mental health in adolescents: Interactions with
	uge and gender - r maings from the Dutch 2001 field in Benaviour in School-Agea Children Survey Journal of Studies on Alashal and Drugs 2005 66(5): n 605 600
25	Laukkanen F.R. et al. Heavy drinking is associated with more severe psychosocial
23.	dysfunction among girls than boys in finland. The Journal of adolescent health
	official publication of the Society for Adolescent Medicine 2001 <b>28</b> (4): p 270-277
26	Theunissen M - I M Jansen and A van Gestel Are mental health and hinge
-0.	drinking associated in Dutch adolescents? Cross-sectional public health study. BMC
	Research Notes, 2011. 4(1); p. 100.
27.	Torikka, A., et al., Depression, drinking, and substance use among 14- to 16-year-old
	Finnish adolescents. Nord J Psychiatry, 2001. 55(5): p. 351-7.
28.	McLeod, J.D., R. Uemura, and S. Rohrman, Adolescent Mental Health, Behavior
	Problems, and Academic Achievement. Journal of Health and Social Behavior, 2012.
	<u>53(4): p. 482-497.</u>
29.	Skogen, J.C., et al., Concurrent validity of the CAGE questionnaire. The Nord-
	Trøndelag Health Study. Addictive behaviors, 2011. 36(4): p. 302-307.
30.	Knight, J.R., et al., A new brief screen for adolescent substance abuse. Archives of
	Pediatrics & Adolescent Medicine, 1999. 153: p. 591-596.
31.	Knight, J.R., et al., Validity of brief alcohol screening tests among adolescents: A
	<u>comparison of the AUDI1, POSI1, CAGE, and CRAFF1. Alconolism: Clinical and</u>
22	Experimental Research, 2005. 27(1). p. 67-75.
52.	<u>Skogell, J.C., et al., <i>Fsychometric properties and concurrent valially of the CKAFF1</i> among Nonvegian adolescents. Ung@hordaland. a population based study. Addictive</u>
	hebayiors 2013(Abead of print)
33	Thanar A and P McGuffin Validity of the shortened Mood and Feelings
<u>.</u>	Questionnaire in a community sample of children and adolescents: a preliminary
	research note. Psychiatry Res., 1998. <b>81</b> (2): p. 259-68.
34.	Sharp, C., I.M. Goodver, and T.J. Croudace. <i>The Short Mood and Feelings</i>
	<u>Questionnaire (SMFQ): a unidimensional item response theory and categorical data</u>
	factor analysis of self-report ratings from a community sample of 7-through 11-year-
	old children. J Abnorm Child Psychol, 2006. 34(3): p. 379-91.
<u>35.</u>	Lundervold, A.J., et al., Symptoms of depression as reported by Norwegian
	adolescents on the Short Mood and Feelings Questionnaire. Frontiers in Psychology,
	2012 4

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36.	Birmaher, B., et al., Psychometric properties of the screen for child anxiety related
	emotional disorders scale (SCARED): A replication study. Journal of the American
	Academy of Child and Adolescent Psychiatry, 1999. 38(10): p. 1230-1236.
<u>37.</u>	Kessler, R.C., et al., Validity of the World Health Organization Adult ADHD Self-
	report Scale (ASRS) Screener in a representative sample of health plan members.
	International Journal of Methods in Psychiatric Research, 2007. 16(2): p. 52-65.
<u>38.</u>	Adler, L.A., et al., Preliminary examination of the reliability and concurrent validity
	of the attention-deficit/hyperactivity disorder self-report scale v1.1 symptom checklist
	to rate symptoms of attention-deficit/hyperactivity disorder in adolescents. Journal of
	Child and Adolescent Psychopharmacology, 2012. 22(3): p. 238-44.
<u>39.</u>	StataCorp, Statistical Software: Release 12.0. 2012, Stata Corporation: College
	Station, TX.
<u>40.</u>	Tapert, S.F., L. Caldwell, and C. Burke, <i>Alcohol and the adolescent brain: Human</i>
	studies. Alcohol Research & Health, 2004. 28(4): p. 205-212.
<u>41.</u>	Chassin, L., et al., A longitudinal study of children of alcoholics: Predicting young
	adult substance use alsoraers, anxiety, and depression. Journal of Abnormal
42	Psychology, 1999. 108(1): p. 100-119.
42.	Silet, K.J., et al., Characteristics of children of alcoholics. Futurive risk factors,
	1001 100(A): p. 427 448
43	Willoughby T et al Framining the link between adolescent brain development and
<del>4</del> 5.	risk taking from a social-developmental perspective Brain and Cognition 2013
	<b>83</b> (3): n 315-323
44.	Morton, S.M.B., et al., In the 21st Century, what is an acceptable response rate?
	Australian and New Zealand Journal of Public Health, 2012, 36(2); p. 106-108.
45.	The Directorate of Integration and Diversity (IMDi), [Education participation in high
	schools]. 2012, http://www.imdi.no/no/Fakta-og-statistikk/Fakta-og-
	statistikk/Utdanning/?tab=chr.
<u>46.</u>	Stormark, K.M., et al., Predicting nonresponse bias from teacher ratings of mental
	health problems in primary school children. J Abnorm Child Psychol, 2008. 36(3): p.
	<u>411-9.</u>
<u>47.</u>	Knudsen, A.K., et al., The health status of nonparticipants in a population-based
	health study: The Hordaland Health Study. American Journal of Epidemiology, 2010.
10	172(11): p. 1306-14.
<u>48.</u>	<u>O'Neil, K.A., B.I. Conner, and P.C. Kendall, Internalizing disorders and substance</u>
	<u>use disorders in youth: Comorbiaity, risk, temporal order, and implications for</u>
40	Intervention. Clinical Psychology Review, 2011. <b>31</b> (1): p. 104-112.
<u>49.</u>	Effects of Age and Conder Clinical Develology Paviay 1004 14(6): p. 407-522
50	Danielsson A K et al. Alcohol use heavy enisodic drinking and subsequent
<u>J0.</u>	problems among adolescents in 23 Furgeean countries: does the prevention naradox
	apply? Addiction 2012 107(1): p 71-80
51	Hibell B et al Summary: The 2007 ESPAD report among students in 35 countries
<u>01.</u>	(2009). European Monitoring Centre for Drugs and Drug Addiction: Luxembourg.
52.	Hoel, S., et al., Adolescent alcohol use, psychological health, and social integration.
	Scandinavian Journal of Public Health, 2004. <b>32</b> : p. 361-367.

audiescents doni from 1993-1993, strauffed for	gender. N=9,203 (girls	n=4,993, boys n=4,208	).			
	Total sample	Born 1995	<u>Born 1994</u>	<u>Born 1993</u>	Odds ratio for age	n-value
	<u>% (95% CI)</u>	<u>% (95% CI)</u>	<u>% (95% CI)</u>	<u>% (95% CI)</u>	<u>trend (95% CI)</u>	<u>p valao</u>
Never consumed alcohol (% yes)						
<u>_Girls</u>	<u>20.7 (19.6-21.8)</u>	<u>31.4 (29.4-33.5)</u>	<u>19.4 (17.6-21.4)</u>	<u>6.3 (5.1-7.7)</u>	<u>0.42 (0.38-0.47)</u>	<u>&lt;.001</u>
Boys	<u>27.1 (25.7-28.4)</u>	<u>39.7 (37.4-42.1)</u>	<u>26.4 (24.2-28.7)</u>	<u>7.6 (6.1-9.4)</u>	<u>0.41 (0.37-0.45)</u>	<u>&lt;.001</u>
Ever tried illicit drugs (% yes)						
<u>_Girls</u>	<u>8.8 (8.1-9.6)</u>	<u>6.1 (5.1-7.3)</u>	<u>8.2 (7.0-9.6)</u>	<u>13.7 (11.9-15.6)</u>	<u>1.57 (1.39-1.77)</u>	<u>&lt;.001</u>
Boys	<u>12.5 (11.5-13.5)</u>	10.8 (8.5-11.2)	<u>10.2 (8.8-11.9)</u>	<u>18.5 (16.3-21.0)</u>	<u>1.37 (1.22-1.54)</u>	<u>&lt;.001</u>
90 <sup>th</sup> percentile alcohol consumption <sup>a</sup> (% yes)						
Girls	<u>7.7 (6.9-8.6)</u>	<u>6.0 (4.8-7.4)</u>	<u>7.7 (6.4-9.2)</u>	<u>9.7 (8.2-11.5)</u>	<u>1.30 (1.13-1.51)</u>	<u>&lt;.001</u>
Boys	<u>7.4 (6.5-8.4)</u>	<u>5.9 (4.6-7.6)</u>	<u>5.9(5.5-8.5)</u>	<u>9.6 (7.9-11.6)</u>	<u>1.31 (1.10-1.55)</u>	<u>=.002</u>
Frequent intoxication (>10; % yes) <sup>b</sup>						
Girls	<u>20.5 (19.4-21.6)</u>	<u>10.6 (9.3-12.0)</u>	<u>19.7 (17.9-21.7)</u>	<u>36.1 (33.6-38.8)</u>	<u>2.20 (2.01-2.41)</u>	<u>&lt;.001</u>
Boys	<u>21.3 (20.1-22.6)</u>	<u>10.7 (9.3-12.3)</u>	<u>21.9 (19.9-24.1)</u>	37.1 (34.2-40.1)	<u>2.21 (2.00-2.44)</u>	<u>&lt;.001</u>
<u>CRAFFT positive (≥2; % yes)</u>						
Girls	<u>23.3 (22.2-24.5)</u>	<u>20.0 (18.3-21.8)</u>	22.4 (20.5-24.4)	<u>29.5 (27.1-32.1)</u>	<u>1.29 (1.19-1.40)</u>	<u>&lt;.001</u>
Boys	<u>20.3 (19.1-21.5)</u>	<u>15.8 (14.1-17.6)</u>	<u>19.3 (17.4-21.4)</u>	<u>29.0 (26.3-31.8)</u>	<u>1.47 (1.34-1.62)</u>	<u>&lt;.001</u>
95 confidence intervals in parentheses <sup>a</sup> Only includes those who have tried alcohol	<u>(n=7,031).</u>					
<sup>b</sup> Only includes those with valid response on	Total sample (95% CI)	<del>Born 1995</del> <del>(95% CI)</del>	<del>Born 1994</del> <del>(95% CI)</del>	<del>Born 1993</del> <del>(95% CI)</del>	OR for age trend (95% CI)	<del>p-value</del>
Never consumed alcohol (yes%)			. ,			
-Girls	<del>20.7% (19.6-21.8)</del>	<del>31.4% (29.4-33.5)</del>	<del>19.4% (17.6-21.4)</del>	<del>6.3% (5.1-7.7)</del>	<del>0.42 (0.38-0.47)</del>	<del>&lt;.001</del>
-Boys	<del>27.1% (25.7-28.4)</del>	<del>39.7% (37.4-42.1)</del>	<del>26.4% (24.2-28.7)</del>	<del>7.6% (6.1-9.4)</del>	<del>0.41 (0.37-0.45)</del>	<del>&lt;.001</del>

Table 1: DescriptionPatterns of abstention, drinking pattern, CRAFFT-casenessand drug use, and problematic alcohol and drug use across age cohorts(CRAFFT positive) in adolescents born from 1993-1995, stratified for gender, N=9 203 (girls n=4 995, boys n=4 208)

Ever tried illicit drugs (yes%)						
<del>Girls</del>	<del>8.8% (8.1-9.6)</del>	<del>6.1% (5.1-7.3)</del>	<del>8.2% (7.0-9.6)</del>	<del>13.7% (11.9-15.6)</del>	<del>1.57 (1.39-1.77)</del>	<del>&lt;.001</del>
Boys	<del>12.5% (11.5-13.5)</del>	<del>10.8% (8.5-11.2)</del>	<del>10.2% (8.8-11.9)</del>	<del>18.5% (16.3-21.0)</del>	<del>1.37 (1.22-1.54)</del>	<del>&lt;.001</del>
90 <sup>th</sup> -percentile alcohol consumption <sup>a</sup> (yes%)						
Girls	<del>7.7% (6.9-8.6)</del>	<del>6.0% (4.8-7.4)</del>	<del>7.7% (6.4-9.2)</del>	<del>9.7% (8.2-11.5)</del>	<del>1.30 (1.13-1.51)</del>	<del>&lt;.001</del>
– <del>Boys</del>	<del>7.4% (6.5-8.4)</del>	<del>5.9% (4.6-7.6)</del>	<del>5.9%(5.5-8.5)</del>	<del>9.6% (7.9-11.6)</del>	<del>1.31 (1.10-1.55)</del>	<del>=.002</del>
Binge drinking (>10; yes%) <sup>b</sup>						
-Girls	<del>20.5% (19.4-21.6)</del>	<del>10.6% (9.3-12.0)</del>	<del>19.7% (17.9-21.7)</del>	<del>36.1% (33.6-38.8)</del>	<del>2.20 (2.01-2.41)</del>	<del>&lt;.001</del>
-Boys	<del>21.3% (20.1-22.6)</del>	<del>10.7% (9.3-12.3)</del>	<del>21.9% (19.9-24.1)</del>	<del>37.1% (34.2-40.1)</del>	<del>2.21 (2.00-2.44)</del>	<del>&lt;.001</del>
CRAFFT positive (≥2; yes%)						
-Girls	<del>23.3% (22.2-24.5)</del>	<del>20.0% (18.3-21.8)</del>	<del>22.4% (20.5-24.4)</del>	<del>29.5% (27.1-32.1)</del>	<del>1.29 (1.19-1.40)</del>	<del>&lt;.001</del>
– <del>Boys</del>	<del>20.3% (19.1-21.5)</del>	<del>15.8% (14.1-17.6)</del>	<del>19.3% (17.4-21.4)</del>	<del>29.0% (26.3-31.8)</del>	<del>1.47 (1.34-1.62)</del>	<del>&lt;.001</del>
95% confidence intervals in parentheses frequency of intoxication (n=9,056).						
CRAFFT: Screening tool for identification of	f problematic alcohol a	and drug use among	adolescents.			
<sup>3</sup> Only includes these who have tried alcoho	1/(n-7.031)					
<sup>b</sup> Only includes those with valid response or	<del>r (n= r,00 r).</del> ⊨ <del>binge drinking (n=9.(</del>	<del>)56).</del>				
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Table 2: Logistic regressions regression models of use of alcohol and illicit drugs as risk factor for mental health problems. Mental health variables analysed separately. N=9,203 (girls n=4,995, boys n=4,208).

	ASRS Inattention OR (95%CI)	ASRS Hyperactivity OR (95%CI)	SMFQ O <del>R (95%Cl)</del>	SCARED OR (95%CI)	
	ASRS Inattention Odds ratio (OR) (95CI)	ASRS Hyperactivity OR (95CI)	<u>SMFQ</u> OR (95CI)	<u>SCARED</u> OR (95CI)	
Ever consumed alcohol					
Crude	2.36 (1.90-2.93)	2.08 (1.68-2.58)	1.73 (1.43-2.09)	1.13 (0.94-1.37)	<b>Formatted:</b> Font: Bold
Adjusted for age and gender	2.21 (1.77-2.76)	2.07 (1.66-2.58)	1.64 (1.35-1.99)	1.03 (0.84-1.25)	<b>Formatted:</b> Font: Bold
Adjusted for age, gender and socioeconomic status (SES) Ever tried illicit drugs	<u>2.15 (1.72-2.69)</u>	<u>2.03 (1.62-2.53)</u>	<u>1.59 (1.31-1.93)</u>	<u>0.99 (0.81-1.21)</u>	
Crude	2.38 (1.98-2.87)	2.26 (1.86-2.74)	1.69 (1.39-2.05)	1.33 (1.05-1.68)	Formatted: Font: Bold
Adjusted for age and gender	2.49 (2.06-3.02)	2.36 (1.94-2.89)	1.98 (1.62-2.42)	1.55 (1.23-1.98)	Formatted: Font: Bold
Adjusted for age, gender and SES 90 <sup>th</sup> percentile alcohol consumption <sup>a</sup>	<u>2.42 (2.00-2.93)</u>	<u>2.31 (1.90-2.82)</u>	<u>1.91 (1.56-2.34)</u>	<u>1.50 (1.18-1.91)</u>	
Crude	1.97 (1.55-2.51)	2.70 (2.14-3.40)	2.19 (1.74-2.75)	1.61 (1.21-2.14)	<b>Formatted:</b> Font: Bold
Adjusted for age and gender	1.96 (1.54-2.50)	2.76 (2.19-3.48)	2.27 (1.79-2.88)	1.64 (1.22-2.19)	Formatted: Font: Bold
Adjusted for age, gender and SES BingeIntoxication >10 times <sup>b</sup>	<u>1.98 (1.55-2.53)</u>	<u>2.77 (2.19-3.49)</u>	<u>2.29 (1.80-2.90)</u>	<u>1.64 (1.22-2.20)</u>	
Crude	1.83 (1.56-2.15)	2.08 (1.77-2.45)	1.30 (1.10-1.52)	0.95 (0.78-1.16)	Formatted: Font: Bold
Adjusted for age and gender	1.80 (1.52-2.12)	2.18 (1.84-2.59)	1.36 (1.15-1.61)	0.97 (0.79-1.20)	Formatted: Font: Bold
Adjusted for age, gender and SES	<u>1.80 (1.52-2.13)</u>	<u>2.17 (1.83-2.58)</u>	<u>1.35 (1.14-1.61)</u>	<u>0.96 (0.78-1.18)</u>	

CRAFFT- <del>casenesspositive</del> (≥2)					
	2 45 (0 74 2 66)	2 04 (2 70 2 70)	0.04 (0.46.0.00)	4 60 (4 40 4 00)	
Crude	3.15 (2.71-3.66)	3.24 (2.78-3.78)	2.84 (2.46-3.28)	1.68 (1.42-1.99)	Formatted: Font: Bold
Adjusted for age and gender	3.06 (2.63-3.56)	3.22 (2.76-3.77)	2.84 (2.45-3.29)	1.62 (1.38-1.94)	Formatted: Font: Bold
<del>95%</del>					
-confidence intervals in parentheses					
*Only includes those who have tried ald	<del>cohol (n=7,031).</del>				
<sup>e</sup> Only includes these with valid	3 01 (2 59-3 51)	3 18 (2 72-3 71)	2 80 (2 41-3 25)	1 59 (1 34-1 90)	
gender and SES	0.01 (2.03-0.01)	0.10 (2.12-0.11)	2.00 (2.41-0.20)	1.00 (1.04-1.00)	
Significant associations in bold.					
<u>95 contidence intervals in parentheses.</u> <sup>a</sup> Only includes those who have tried all					
<sup>b</sup> Only includes those with valid response	se on <del>binge drinking</del> frequency	of intoxication (n=9.056)			
CRAFFT: Screening tool for identification of	f problematic alcohol and drug u	se among adolescents.			
ASRS: Adult ADHD Self-report Scale, scree	ening inventory for attention-defi	cit-hyperactivity disorder. ASRS In	attention: the inattention sub-scale	e of the ASRS (9 items). ASRS	
Hyperactivity: the hyperactivity-impulsivity s	subscale of the ASRS (nine item	<u>s).</u>			
<u>SCARED: five-item screeping inventory for</u>	Feelings Questionnaire, screenil symptoms of anyiety and anyiet	ng inventory for symptoms of depr v disorders	ession and depressive disorders.		
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	ASRS in	attention	ASRS Hy	peractivity	SM	FQ	SCA	RED
	Gender	Age	Gender	Age	Gender	Age	Gender	Age
ver consumed alcohol	<del>p=0.187</del>	<del>p=0.595</del>	<del>p=0.466</del>	<del>p=0.002</del>	<del>p=0.257</del>	<del>p=0.255</del>	<del>p=0.052</del>	<del>p=0.182</del>
ver tried illicit drugs	<del>p=0.666</del>	<del>p=0.630</del>	<del>p=0.542</del>	<del>p=0.384</del>	<del>p=0.054</del>	<del>p=0.978</del>	<del>p=0.021</del>	<del>p=0.425</del>
<sup>th</sup> percentile alcohol consumption	<del>p=0.752</del>	<del>p=0.359</del>	<del>p=0.021</del>	<del>p=0.937</del>	<del>p=0.018</del>	<del>p=0.639</del>	<del>p&lt;0.001</del>	<del>p=0.198</del>
<del>nge &gt;10</del>	<del>p=0.083</del>	<del>p=0.427</del>	<del>p=0.514</del>	<del>p=0.025</del>	<del>p=0.027</del>	<del>p=0.889</del>	<del>p=0.245</del>	<del>p=0.452</del>
<del>RAFFT-caseness (≥2)</del>	<del>p=0.413</del>	<del>p=0.263</del>	<del>p=0.293</del>	<del>p=0.190</del>	<del>p=0.100</del>	<del>p=0.840</del>	<del>p=0.991</del>	<del>p=0.46</del> 4
			*					_
	ASRS inatter	tion	ASRS Hyperactiv	<u>vity</u>	SMFQ	) Odda rai	CARED	
ver consumed alcohol	<u>1.83 (1.44-2.</u>	<u>(9501)</u> <u>32)</u>	<u>1.71 (1.35-2.16</u>	$\frac{501}{1}$	45 (1.17-1.78)	<u>00081a</u> <u>0.79</u>	(0.64-0.98)	-
ver tried illicit drugs	<u>1.87 (1.50-2</u> .	<u>32)</u>	<u>1.74 (1.40-2.17</u>	) <u>1.</u>	4 <u>8 (1.18-1.86)</u>	<u>1.12</u>	<u>(0.86-1.46)</u>	
0 <sup>th</sup> percentile alcohol consumption <sup>a</sup>	<u>1.24 (0.94-1.</u>	<u>63)</u>	<u>2.30 (1.78-2.98</u>	<u>) 1.</u>	<u> 88 (1.45-2.44)</u>	<u>1.18</u>	<u>(0.86-1.62)</u>	
toxication >10 <sup>b</sup>	<u>1.45 (1.20-1</u>	<u>75)</u>	<u>1.95 (1.62-2.35</u>	<u>) 1.</u>	<u>18 (0.98-1.43)</u>	<u>0.79</u>	<u>(0.63-0.99)</u>	
RAFFT-positive (≥2)	<u>1.97 (1.66-2.</u>	<u>34)</u>	<u>2.27 (1.91-2.69</u>	<u>) 2.</u>	<u>17 (1.84-2.55)</u>	0.99	<u>(0.82-1.21)</u>	
gnificant interactionsassociations in t	old. Interactions	s assessed (	using Wald likeliho	od-ratio tests				

CRAFFT: Screening tool for identification of problematic alcohol and drug use among adolescents.

ASRS: Adult ADHD Self-report Scale, screening inventory for attention-deficit-hyperactivity disorder. ASRS Inattention: the inattention sub-scale of the ASRS (9 items). ASRS

Hyperactivity: the hyperactivity-impulsivity subscale of the ASRS (nine items).

SMFQ: The short version of the Mood and Feelings Questionnaire, screening inventory for symptoms of depression and depressive disorders.

SCARED: five-item screening inventory for symptoms of anxiety and anxiety disorders.

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Table 4: Logistic regression models across alcohol consumption and binge drinkingfrequency of intoxication categories, "Never tried alcohol" as reference. Adjusted for age	
and gender (N=9,203 for alcohol consumption and N=9,056 for binge drinking frequency of intoxication).	

	ASRS Inattention Odds ratio (OR (95%CI) (95CI)	ASRS Hyperactivity OR ( <del>95%Cl<u>95Cl</u>)</del>	SMFQ OR ( <del>95%Cl<u>95Cl</u>)</del>	SCARED OR ( <del>95%Cl<u>95Cl</u>)</del>
Categories	Alc	ohol consumption categories	as exposureindependent variab	ble
Never consumed alcohol	1.00 (Reference)	1.00 (Reference)	1.00 (Reference)	1.00 (Reference)
Non-consumption	<u>1.90</u> (1.45-2.48)	<u>1.46</u> (1.10-1.93)	<u>1.43</u> (1.12-1.82)	<u>1.01</u> (0.78-1.30)
).1-19.9 <sup>th</sup> percentile	<u>(1.19-2.17)</u>	<u>(1.25-2.27)</u>	<u>1.44</u> (1.11-1.88)	<u>1.00</u> (0.75-1.32)
20.0-79.9 <sup>th</sup> percentile	<u>2.15</u> (1.69-2.74)	<u>1.99</u> (1.56-2.55)	<u>1.51</u> (1.21-1.88)	<u>0.90</u> (0.72-1.14)
0.0-89.9 <sup>th</sup> percentile	<u>(2.75-5.17)</u>	<u>3.56</u> (2.59-4.91) 5.28	<u>2.21</u> (1.62-3.02) 2.47	<u>1.39</u> (0.99-1.96) <b>1.60</b>
00.0-100 <sup>th</sup> percentile	<u>4.16</u> (3.05-5.68)	<u>(3.91-7.12)</u>	<u>(2.60-4.62)</u>	<u>(1.15-2.23)</u>
-value for trend	<u>p&lt;0.001</u>	<u>p&lt;0.001</u>	<u>p&lt;0.001</u>	<u>p=0.061</u>
Categories	<u>F</u>	Frequency of intoxication cate	gories as independent variable	
lever consumed alcohol	1.00 (Reference)	1.00 (Reference)	1.00 (Reference)	1.00 (Reference)
lon-consumptionNever intoxicated	1. <del>90<sup></sup><u>65</u> (1.45<u>24</u>-2.48<u>20)</u></del>	(1.10 <u>00</u> -1. <u>9381</u> )		1. <u>0110</u> <u>- (0.<del>7885</del>-1.<del>3043),</del></u>
0.1-19.9 <sup>th</sup> percentileOnce	1. <del>61 71</del> (1. <del>19<u>23</u>-2.<del>17<u>38)</u></del></del>	<u>2.06</u> (1.69 <sup></sup> (1.25 <b>50-2</b> 2782)	<u>1-44<sup>=</sup>55</u> (1- <u>11-1-88<mark>16-2:08</mark>)</u>	1. <u>0002</u> 
20.0-39.9 <sup>th</sup> percentile	<del>1.98<sup>***</sup> (1.46-2<u>.69)</u></del>	<del>1.91</del> (1.40-2.60)	<del>1.53</del> <sup>#</sup> ( <del>1.16-2.03)</del>	<del>0.90</del> ( <del>0.66-1.23)</del>
10.0-59.9 <sup>th</sup> percentile	<del>1.92*** (1.42-2.59)</del>	<del>1.55</del> <sup>™</sup> (1.13-2 <u>.1</u> 3)	<del>1.16</del> (0.87-1.56)	<del>0.97</del> <del>(0.72-1.30)</del>
60.0-79.9 <sup>th</sup> percentile	<del>2.56</del> (1.92-3.40)	<del>2.55</del>	<del>1.85</del>	<del>0.84</del> <del>(0.62-1.14)</del>
1 <del>0.0-89.9<sup>th</sup> percentile</del>	<del>3.77 -</del> <del>(2.75 5.17)</del>	<del>3.56 (2.59 4.91)</del>	<del>2.21</del> <del>(1.62 <u>3.02)</u></del>	<del>1.39</del> <del>(0.99-1.96)</del>
90.0-100 <sup>th</sup> -percentile	4 <del>.16<sup></sup> (3.05-5.68)</del>	<del>5.28</del> <sup></sup> ( <del>3.91-7.12)</del>	<del>3.47 -</del> <del>(2.60-4.62)</del>	<del>1.60<sup></sup> (<del>1.15-2.23)</del></del>

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Categories         Binge drinking categories: as exposure           Never consumed alcohol         1.00 (Reference)         1.00 (Reference)         1.00 (Reference)           Never binge         1.435         1.395         1.496           Once         1.474         2.065         1.652         1.62           2.3 times         (1.512.63)         (1.212.15)         (1.262.07)         (0.67-1.1)           2.3 times         (1.512.63)         (1.212.15)         (1.262.07)         (0.67-1.1)           4.10 times         (1.90-3.26)         (1.552.72)         (1.442.36)         (0.81-1.3)           >10 times         (1.90-3.26)         (1.552.72)         (1.442.36)         (0.81-1.3)           >10 times         (2.434.05)         (2.614.33)         (1.642.47)         (0.76-1.2)           p-value for trend         p<0.001         p<0.001         p<0.001         p<0.001         p=0.675 $p < 0.005, " p < 0.01, " p < 0.001         p<0.001         p<0.001         p=0.675         0.675  $		
Never consumed alcohol       1.00 (Reference)       1.00 (Reference)       1.00 (Reference)       1.00 (Reference)         Never binge       1.66 <sup>±±</sup> 1.35 <sup>±±</sup> 1.30 <sup>±±</sup> 1.40 <sup>±</sup> Once       1.44 <sup>±±</sup> 2.06 <sup>±±±</sup> 1.46 <sup>±±</sup> 1.40 <sup>±</sup> Once       (1.24.2.20)       (1.00-1.81)       (1.00-1.67)       (0.85-1.4         Once       (1.23.2.33)       (1.60.2.92)       (1.46-2.08)       (0.74-1.3)         2-3 times       (1.51-2.63)       (1.21-2.15)       (1.26-2.07)       (0.87-1.1)         1-10 times       (1.90-3.26)       (1.55-2.72)       (1.44-2.36)       (0.81-1.3)         1-10 times       (2.43 <sup>±±</sup> )       2.05 <sup>±±</sup> 1.95 <sup>±±</sup> 0.98 <sup>±±</sup> 1-10 times       (2.43-4.05)       (2.61-4.33)       (1.54-2.47)       (0.76-1.2)         0-value for trend       p<0.001       p<0.001       p<0.001       p=0.679         p<4.005, <sup>±±</sup> p < 0.015ignificant associations in comparison to "Never tried alcohol"-category in bold.       0.98 <sup>±±</sup> 0.98 <sup>±±</sup>		
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ASRS: Adult ADHD Self-report Scale, screening inventory for attention-deficit-hyperactivity disorder. ASRS Inattention: the inattention sub-scale of the ASRS (9 items). ASRS Hyperactivity: the hyperactivity-impulsivity subscale of the ASRS (nine items).

SMFQ: The short version of the Mood and Feelings Questionnaire, screening inventory for symptoms of depression and depressive disorders.

SCARED: five-item screening inventory for symptoms of anxiety and anxiety disorders.

### **Figure legends**

Figure 1: Proportion of participants having evernever consumed alcohol across gender-stratified age groups (N=9,203, girls n=4,208). Error bars represent 95% confidence intervals.

#### and

.ed age groups (N=9,203, girls Figure 2: Proportion of participants having ever tried illicit drugs across gender-stratified age groups (N=9,203, girls n=4,995, boys n=4,208). Error bars represent 95% confidence intervals.



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25% 20% -5% -6.1% 10.8% -8.2% 10.2% -10.2% -13.7% 18.5% Born 1995 (aged 17) Born 1995 (aged 19)

■ Girls ■ Boys

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# STROBE Statement—checklist of items that should be included in reports of observational studies **"X" indicates checked**

	Item No	Recommendation
Title and abstract	1 X	(a) Indicate the study's design with a commonly used term in the title or the abstract
		(b) Provide in the abstract an informative and balanced summary of what was done
		and what was found
Introduction		
Background/rationale	2 X	Explain the scientific background and rationale for the investigation being reported
Objectives	3 X	State specific objectives including any prespecified hypotheses
Methods	JA	State specific objectives, meruang any prespectified hypotheses
Study design	4 X	Present key elements of study design early in the paper
Setting	5 X	Describe the setting locations and relevant dates including periods of recruitment
betting	5 1	exposure follow-up and data collection
Participants	6 X	(a) Cohort study—Give the eligibility criteria and the sources and methods of
i unopunto	0 11	selection of participants. Describe methods of follow-up
		<i>Case-control study</i> —Give the eligibility criteria and the sources and methods of
		case ascertainment and control selection. Give the rationale for the choice of cases
		and controls
		<i>Cross-sectional study</i> —Give the eligibility criteria and the sources and methods of
		selection of participants
		(b) Cohort study—For matched studies, give matching criteria and number of
		exposed and unexposed
		<i>Case-control study</i> —For matched studies, give matching criteria and the number of
		controls per case
Variables	7 X	Clearly define all outcomes, exposures, predictors, potential confounders, and effect
		modifiers. Give diagnostic criteria, if applicable
Data sources/	8* X	For each variable of interest, give sources of data and details of methods of
measurement		assessment (measurement). Describe comparability of assessment methods if there
		is more than one group
Bias	9 X	Describe any efforts to address potential sources of bias
Study size	10 X	Explain how the study size was arrived at
Quantitative variables	11 X	Explain how quantitative variables were handled in the analyses. If applicable,
		describe which groupings were chosen and why
Statistical methods	12 X	(a) Describe all statistical methods, including those used to control for confounding
		(b) Describe any methods used to examine subgroups and interactions
		(c) Explain how missing data were addressed
		(d) Cohort study—If applicable, explain how loss to follow-up was addressed
		<i>Case-control study</i> —If applicable, explain how matching of cases and controls was addressed
		<i>Cross-sectional study</i> —If applicable, describe analytical methods taking account of
		sampling strategy
		(e) Describe any sensitivity analyses
		<u> </u>

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Results		
Participants	13*	(a) Report numbers of individuals at each stage of study-eg numbers potentially eligible,
	Х	examined for eligibility, confirmed eligible, included in the study, completing follow-up, and
		analysed
		(b) Give reasons for non-participation at each stage
		(c) Consider use of a flow diagram
Descriptive	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and
data	Х	information on exposures and potential confounders
		(b) Indicate number of participants with missing data for each variable of interest
		(c) Cohort study—Summarise follow-up time (eg, average and total amount)
Outcome data	15	Cohort study—Report numbers of outcome events or summary measures over time
	X*	Case-control study-Report numbers in each exposure category, or summary measures of
		exposure
		Cross-sectional study—Report numbers of outcome events or summary measures
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their
	Х	precision (eg, 95% confidence interval). Make clear which confounders were adjusted for and
		why they were included
		(b) Report category boundaries when continuous variables were categorized
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a
		meaningful time period
Other analyses	17	Report other analyses done-eg analyses of subgroups and interactions, and sensitivity
	Х	analyses
Discussion		
Key results	18	Summarise key results with reference to study objectives
	Х	
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision.
	Х	Discuss both direction and magnitude of any potential bias
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations,
	Х	multiplicity of analyses, results from similar studies, and other relevant evidence
Generalisability	21	Discuss the generalisability (external validity) of the study results
	Х	
Other informati	on	
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable,
	Х	for the original study on which the present article is based

\*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

**Note:** An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.