



UNIVERSITÀ DEGLI STUDI DI TORINO

AperTO - Archivio Istituzionale Open Access dell'Università di Torino

Short- term mediating factors of a school-based intervention to prevent youth substance use in Europe

This is the author's manuscript Original Citation: Availability: This version is available http://hdl.handle.net/2318/147142 since 2016-07-14T10:06:20Z Published version: DOI:10.1016/j.jadohealth.2013.10.009 Terms of use: Open Access Anyone can freely access the full text of works made available as "Open Access". Works made available under a Creative Commons license can be used according to the terms and conditions of said license. Use of all other works requires consent of the right holder (author or publisher) if not exempted from copyright

(Article begins on next page)

protection by the applicable law.





This Accepted Author Manuscript (AAM) is copyrighted and published by Elsevier. It is posted here by agreement between Elsevier and the University of Turin. Changes resulting from the publishing process - such as editing, corrections, structural formatting, and other quality control mechanisms - may not be reflected in this version of the text. The definitive version of the text was subsequently published in JOURNAL OF ADOLESCENT HEALTH, 54 (5), 2014, 10.1016/j.jadohealth.2013.10.009.

You may download, copy and otherwise use the AAM for non-commercial purposes provided that your license is limited by the following restrictions:

(1) You may use this AAM for non-commercial purposes only under the terms of the CC-BY-NC-ND license.

(2) The integrity of the work and identification of the author, copyright owner, and publisher must be preserved in any copy.

(3) You must attribute this AAM in the following format: Creative Commons BY-NC-ND license (http://creativecommons.org/licenses/by-nc-nd/4.0/deed.en), 10.1016/j.jadohealth.2013.10.009

The publisher's version is available at: http://linkinghub.elsevier.com/retrieve/pii/S1054139X13005442

When citing, please refer to the published version.

Link to this full text: http://hdl.handle.net/2318/147142

This full text was downloaded from iris - AperTO: https://iris.unito.it/

Short-Term Mediating Factors of a School-Based Intervention to Prevent Youth Substance Use in Europe

Fabrizia Giannotta, Ph.D. a,*, Federica Vigna-Taglianti, Ph.D. b,c, Maria Rosaria Galanti, M.D., Ph.D. d,e, Maria Scatigna, Ph.D. f, and Fabrizio Faggiano, M.D., Ph.D. c,g

aCenter for Developmental Research, Örebro University, Örebro, Sweden

b Department of Clinical and Biological Sciences, University of Torino, Torino, Italy

c Piedmont Centre for Drug Addiction Epidemiology, ASL TO3, Torino, Italy

d Department of Public Health Sciences, Karolinska Institutet, Stockholm, Sweden

e Centre for Epidemiology and Community Medicine, Stockholm Health Care District, Stockholm, Sweden

f Department of Life, Health and Environmental Sciences, University of Aquila, L'Aquila, Italy

g Department of Translational Medicine, Avogadro University, Novara, Italy

ABSTRACT

Purpose: To investigate factors mediating the effects of a European school-based intervention (Unplugged) based on a social influence approach to youths' substance use.

Methods: Schools in seven European countries (n = 143, including 7,079 pupils) were randomly assigned to an experimental condition (Unplugged curriculum) or a control condition (usual health education). Data were collected before (pre-test) and 3 months after the end of the program (post-test). Multilevel multiple mediation models were applied to the study of effect mediation separately for tobacco, alcohol, and cannabis use. Analyses were conducted on the whole sample, and separately on baseline users and nonusers of each substance.

Results: Compared with the control group, participants in the program endorsed less positive attitudes toward drugs; positive beliefs about cigarettes, alcohol, and cannabis; and the normative perception of peers using tobacco and cannabis. They also increased in knowledge about all substances and refusal skills toward tobacco. Decreased positive attitudes toward drugs, increase in refusal skills, and reappraisal of norms about peer using tobacco and cannabis appeared to mediate the effects of the program on the use of substances. However, mediating effects were generally weak and some of them were only marginally significant. Conclusions: This study lends some support to the notion that school-based programs based on a social influence model may prevent juvenile substance use through the modification of attitudes, refusal skills, and normative perceptions.

School-based interventions are considered effective in the prevention of substance use in adolescence. Recent reviews suggest that even long-term prevention of alcohol, cigarettes, and marijuana use among youths is possible (e.g., [1e3]). In particular, interventions based on a social influence approach, which aims to reinforce personal and interpersonal skills using interactive techniques, and including a normative education component, are more likely to be effective than others [1,2,4].

Whereas evidence of effectiveness of social influence interventions is well established [2], evidence about potential mediating mechanisms is still lacking. Identification of mediators of the intervention's effects, if any, is important to support the theoretical model behind the program [5]. This, in turn, allows the individuation of effective components and guides the efforts of development of new interventions [6].

Components of theories on adolescents' behaviors have been tested through mediation analyses in randomized trials. Specifically, school-based interventions have been shown to modify youths' substance use mainly by changing normative expectations or perceived norms about the use [7e14], intentions [8,15], refusal and resistance skills [12,16,17], risk-related attitudes and behaviors [12,14,16,18], positive and negative beliefs about consequences [15], reasons to use [13,18], and perceived peers' influence and friends' use [14,15,18].

However, only few studies in the past adopted up-to-date methods to investigate mediation: namely, a multiple mediation approach [19] and calculation of the statistical significance of mediating effects [6]. Moreover, all these studies were conducted in the United States. To understand whether the identified mediating mechanisms can be generalized to other populations, replication of similar approaches is required.

This study aimed to analyze short-term mediating factors of a school-based curriculum (Unplugged) for the prevention of tobacco, alcohol, and drugs use evaluated in seven European countries. Unplugged is a social influence curriculum consisting of 12 1-hour sessions delivered weekly by class teachers (for more details, see [20]). It has been shown to reduce the use of tobacco, alcohol, and cannabis in the short term [21]. According to the leading theories of the social influence approach (social learning and social norms), drug use initiation is the result of social influence, from which adolescents can derive erroneous perceptions of frequency and acceptability of use. Normative education and resistance skills training included in prevention curricula are thought to reduce the effect of social influence by modifying attitudes, beliefs, and normative perceptions, finally supporting the development of general social skills and skills to resist social pressures. Accordingly, we hypothesized that beliefs about consequences of substance use; attitudes toward drug use; ability to resist an offer of alcohol, cigarettes, or cannabis (refusal skills); and perception of prevalence of use among peers would mediate the effects of the program on behavioral outcomes (Figure 1). Because previous studies have shown that school climate and school bonding were positively affected by school-based interventions [22e24] and could act as protective factors for substance use [23e26], we also included perceived social climate in the class, to investigate whether it mediates interventions' effects . Finally, we were interested in examining whether the Unplugged curriculum works differently for students who were users versus those who were not, as suggested in a previous study [18].

Methods

Study design

The present study was based on a four-arm, cluster randomized, controlled trial (the EUropean Drug Addiction Prevention [EU-DAP] trial), in which schools were randomly assigned either to one of three experimental arms (Unplugged curriculum alone, or Unplugged complemented by parents' seminars or peer sessions) or to a control group receiving the usual health education curriculum. Of 323 eligible schools, 33 were excluded because they did not meet the inclusion criteria, which were

following the national educational curricula, encompassing at least two classes in the target grade, and not concurrently conducting other comprehensive interventions against substance use. An additional 120 schools declined participation, mainly because they were unable to schedule the intervention during the 2004e2005 school year (74%). Thus, 170 schools were randomized, after stratification into three levels of neighborhood average socioeconomic status.

Computer-generated blocks of four digits were used to allocate schools within centers. Every five schools, two were assigned to the control group, and the remaining three were assigned to one of the intervention arms. Sixteen percent of schools (n = 27) withdrew from the study before the baseline survey (23.5% in the intervention arms and 4.4% in the control arm). This withdrawal occurred mainly during or just before the training of teachers, was comparable in all centers, and was similar across the three levels of area social stratification. Details on the study design and participation rates have been published elsewhere [27]. The program was conducted in 78 schools, whereas 65 acted as controls.

Because the added component of parents' education and the peer sessions was not implemented in practice [20], analyses were conducted pooling together the three intervention arms, in line with previous reports from this trial [21].

Participants

Pre-test data were collected from 7,079 students in seven different countries (Austria, Belgium, Germany, Greece, Italy, Spain, and Sweden). For 6,370 of them (48% females; mean age, 13.25 years; standard deviation [SD], .99 years), valid post-test information was obtained 3 months after the end of the program.

Students with both assessments did not differ significantly from those who dropped out, with respect to socio-demographic variables and substance use at baseline.

A general policy of informed consent was not adopted in the trial. Each participating center followed the practice of ethical clearance required locally.

Fidelity

The curriculum consisted of 12 1-hour units taught by class teachers who had previously attended a 2.5-day training course. Fidelity of the program's implementation was assessed across the different centers (for details on the procedure, see [20]). A total of 56% of the enrolled classes implemented all the units in the curriculum, 66% at least 10, and 77% at least six.

Fewer than 5% of classes failed to implement any part of the curriculum. On average, each unit was taught to 78% of the target population. This level of program implementation is comparable to that of other curricula administered in a European setting [28].

Measures

The questionnaire items were drawn primarily from existing international surveys included in the European Monitoring Centre for Drugs and Drug Addiction Evaluation Instrument Bank database (http://eib.emcdda.europa.eu/index.cfm?nNodeID=3267). Table 1 presents the measures adopted in this analysis, along with their psychometric properties. For all constructs (except knowledge), higher scores indicate a higher level of risk (see score points), so that negative values of Path a in Tables 2e4 indicate a reduction of the risk level.

Because of the statistical models adopted for the analysis in this study (i.e., linear regression models), the measures were used as continuous, whereas in the previous papers they were dichotomized [21]. In this analysis, we did not employ the frequency of drunkenness and cannabis use in the past 30 days because of the low variance of these variables at baseline. Statistical analyses We fitted multilevel multiple mediation models in Mplus 6 (Muthén & Muthén, Los

Angeles, CA), entering all hypothesized mediators simultaneously [19]. The conceptual model of mediation is depicted in Figure 1. We calculated the intervention effect for each mediator (Path a in Figure 1), the effect of each mediator on the outcome (Path b), and the intervention effect that was mediated by each mediator (Path a*b in the tables or "indirect effect" in the text). For each model, we computed the total indirect effects of the intervention (i.e., the total effect of intervention mediated by the sum of the mediating factors) and also the single indirect effect for each mediator, as suggested by Preacher and Hayes [19]. Moreover, we calculated the program's direct effect (i.e., the effect that was not explained by the hypothesized mediators). Because the randomization occurred at the school level, we entered school as the second level, and individuals as the first level. To control for variability across centers, we used the stratification option in Mplus. We fitted separate models for tobacco, alcohol, and cannabis. We performed separate analyses among students who were nonusers (had not smoked in the past 30 days, never got drunk, never used cannabis) and among users at baseline. In all models, we controlled for gender, age, and the initial levels of the variables at baseline. We allowed covariations among all variables within time. Concerning model fit, we examined the chi-square, comparative fit index (recommended values >.94), root mean square error of approximation (recommended values <.08) and standardized rootmean-square residual (recommended values <.08 [29,30]). All tests were two-tailed. Given the reduction of statistical power as a result of mediation analyses [19], we also presented effects with .05 as marginally significant.

Results

Whole sample

All models showed a good fit to the data (Tables 2e4). Participants in the intervention significantly reduced their positive attitudes toward drugs; positive beliefs about consequences of tobacco, alcohol, and cannabis use; and the perception of having many friends who smoke cigarettes and cannabis compared with the control group (Path a of Tables 2e4). They also increased their ability to refuse peers' invitations to smoke cigarettes and their knowledge about tobacco, cannabis, and alcohol, and reported an improvement in class climate. Although only marginally significant (p < .10), Unplugged also increased refusal skills for alcohol and cannabis, and negative beliefs about the consequences of tobacco use.

Concerning tobacco use, the total indirect effect of intervention ($\beta = -.016$; p = .013) was significant, whereas the direct effect ($\beta = -.014$; p = .092) was only marginally significant (Table 2). Ability to refuse peers' invitation to smoke ($\beta = -.008$; p = .040) and perception of the number of smoking friends ($\beta = -.004$; p = .016) were significant mediators of intervention effects on cigarettes use (Path a*b), whereas the mediating effect of positive attitudes toward drugs was only marginally significant ($\beta = -.003$; p = .070).

For alcohol-related behavior, the total indirect effect of intervention ($\beta = -.013$; p = .050) was significant, whereas the direct effect was not (Table 3). Positive attitudes toward drug were mediators of the intervention effects on alcohol ($\beta = -.005$; p = .046) (Path a*b). The mediating effect of positive beliefs about consequences of alcohol use ($\beta = -.001$; p = .096) and of the ability to refuse peers' invitation to drink was instead only marginally significant ($\beta = -.005$; p = .078). Both total indirect effects ($\beta = -.013$; p = .066) and direct effects ($\beta = -.018$; p = .090) of the intervention toward cannabis use were only marginally significant (Table 4). The decrease in perception of having many smoking friends ($\beta = -.002$; p = .048) (Path a*b) was a significant mediator of the intervention effects on cannabis use, whereas the mediating effects of the decrease in positive attitudes toward drugs ($\beta = -.005$; p = .060) and the increase in the ability to refuse peers' invitation to smoke ($\beta = -.006$; p = .078) were only marginally significant.

Never-users at baseline

All models fit the data well (Tables 2e4). Students who did not smoke in the past 30 days at baseline and took part in the intervention significantly increased in knowledge about tobacco (β = .056; p = .010) and decreased in the perception of having many friends who smoked (β = -.047; p= .032) and in marginally positive beliefs about the consequences of tobacco smoking (β = -.039; p= .064), compared with controls (Path a, Table 2). Students in this group also reported an increase in positive class climate (β = -.062; p = .006). The total indirect effect of the intervention was significant (β = -.017; p = .033), whereas the direct effect was not significant (Table 2). Increased knowledge about tobacco (β = -.002; p = .070) and the reduction in the perception of having friends who smoke (β = -.003; p = .058) were marginally significant mediators of the intervention effects on smoking frequency among non smokers at baseline (Path a*b, Table 2).

Among students who never had never been drunk, taking part in the program strongly increased knowledge about alcohol ($\beta = .164$; p = .000) and the perception of a positive class climate ($\beta = .045$; p = .034), compared with the control group (Path a, Table 3). However, mediation analyses showed that neither total indirect effects nor direct effect of the intervention were significant (Table 3). None of the factors examined were revealed to be a mediator of the intervention's effects (Path a*b, Table 3).

Finally, among those who had never tried cannabis, participants in the intervention decreased their positive beliefs about the consequences of cannabis use ($\beta = -.045$; p = .014) and increased in knowledge about cannabis ($\beta = .141$; p = .000) and the perception of a positive class climate ($\beta = -.053$; p = .012) compared with controls (Path a, Table 4). The program also marginally reduced the perception of having friends who smoked cannabis ($\beta = -.035$; p = .066). Total indirect effects of the intervention were not significant. Only the perception of having friends who smoked cannabis was found to be a marginally significant mediator of the intervention's effects ($\beta = -.025$; p = .090) (Path a*b, Table 4).

Users at baseline

All models fit the data well (Tables 2e4). Participants in the intervention who at baseline reported cigarette smoking in the past 30 days decreased their positive attitudes toward drugs (β =-.075; p= .006) and the perception of having many friends who smoked cigarettes (β = -.055; p = .038), relative to the control group (Path a, Table 2). Moreover, they increased in negative beliefs about the consequences of smoking cigarettes (β = -.040; p = .084) and their ability to say no to peers' invitation to smoke cigarettes (β = -.041; p = .066), although these two effects were only marginally significant. Notwithstanding, there was a significant total indirect effect of the intervention (β =-.027; p = .013) and a marginally significant direct effect (β =-.027; p = .062) (Table 2). The decrease in positive attitudes toward drugs (β = -.006; p = .044) and in the perception of having many friends who smoked (β = -.007; p = .044) mediated the intervention's effects on frequency of tobacco use among smokers at baseline (Path a*b, Table 2). The increase in the ability to resist peer pressure to smoke mediated the intervention effects on tobacco use (β = -.013; p = .066), but it attained only marginal significance.

Intervention students who reported an experience of drunkenness at baseline decreased positive attitudes toward drugs ($\beta = -.091$; p = .002), positive beliefs about alcohol drinking($\beta = -.084$; p = .004), as well as the perception of having many friends who got drunk ($\beta = -.59$; p = .044), compared with controls (Path a, Table 3). They also increased in negative attitudes toward drugs ($\beta = -.069$; p = .020), knowledge about alcohol ($\beta = -.128$; p = .000), and marginally, the ability to say no to peer pressure to drink ($\beta = -.055$; p = .068), compared with controls. Mediation analyses revealed a significant total indirect effect of the intervention ($\beta = -.035$; p = .006), whereas the direct effect was no longer significant (Table 3). The decrease in positive attitudes toward drugs ($\beta = -.015$; p = .008) and in positive beliefs about the consequences of alcohol use ($\beta = -.006$;

p= .038) mediated the intervention's effect on the frequency of drunkenness among baseline drinkers, whereas the decreased perception of having many friends who got drunk (β = -.005; p= .076) and the increased ability to refuse a peer's invitation to drink (β = -.007; p = .094) were marginally significant mediators (Path a*b, Table 3).

Students who had already tried cannabis at baseline and who took part in the intervention decreased their positive attitudes toward drugs (β =-.137; p = .004), and they marginally decreased their positive beliefs about the consequences of cannabis use (β = -.085; p = .076) and the perception of having many friends who smoked (β =-.081; p = .094), compared with controls (Path a, Table 4). They also increased their negative attitudes toward drugs (β = -.111; p = .030) and their knowledge about cannabis (β = -.130; p = .010). Overall, the total indirect effects were significant and in the anticipated direction (β = -.048; p = .072), whereas the direct effect was no longer significant (Table 3). Single mediation analyses did not identify significant mediation effects of any of the proposed mediators. However, the increased negative attitudes toward drugs (β = -.010; p = .096) were marginally significant mediators of the intervention effects (Path a*b, Table 4).

Finally, increased knowledge about cannabis owing to the intervention was marginally significant related to an increase in cannabis use ($\beta = .007$; p = .090).

Discussion

The aim of this study was to investigate the short-term mediating mechanisms of a European school program based on social influence (Unplugged) on youths' substance use. Overall, Unplugged reduced cigarette smoking, drunkenness episodes, and cannabis use through three common mediating factors: attitudes, refusal skills, and perception of prevalence of the behavior among peers. Although the associations between intervention and putative mediators were generally in the expected direction, and often so even at the conventional threshold for statistical significance, the mediating effects were in many cases only marginally significant, especially for nonusers at baseline. Among users, change in attitudes toward drugs and possibly the perception of the number of friends who used drugs mediated the intervention effects on cigarette smoking and alcohol misuse, as predicted by theories [31,32].

Overall, this study highlights similarities in the mediating mechanisms for tobacco, alcohol, and cannabis use. To our knowledge, this is an original finding in the context of experimental studies, in which substances are often combined together (e.g., [9,14]). Modification of positive attitudes toward drugs, of refusal skills, and of normative perceptions emerged as possible mediating factors of the effect of the program on all three classes of substances.

Our study leads to two main considerations of importance for future prevention. First, the intervention effects were not completely explained by the reinforcement of the skills targeted by the intervention. Second, in many cases, mediating effects were weak and only marginal, which suggests that Unplugged prevented substance use through alternative pathways. However, we cannot exclude other plausible explanations, such as inaccuracy of measurements, weak overall effects of the program, and strong interaction with contextual variables (e.g., at the center's level) that may have masked mediation effects, which emphasizes the need for careful planning of future evaluation studies that will enable the analysis of mediating mechanisms.

Almost all putative mediating factors included in this analysis affected outcomes in the predicted direction, they explained part of the program's effect among the whole study population, and they were significant among students with experience of substance use at the intervention's start, but not among nonusers. A possible explanation is that the program's educational activities, such as role-plays, were more relevant to students with this experience. Relevance, and consequent identification processes, may affect the motivation to learn and the acquisition of skills (see, for instance, [33]). However, because the effects of Unplugged on perceived peers' prevalence and resistance skills are found also among those who did not drink alcohol at baseline (Caria MP, Faggiano F, Bellocco R, et al. Effects of a school-based prevention program on European adolescents' intentions,

perceptions and expectations towards alcohol drinking. In preparation), we do not exclude the possibility that the program might work differently for pupils at different stages of alcohol use. The weak indication of an increased risk for cannabis use conveyed by an increase in knowledge about this substance might be due to chance. However, the role of knowledge enhancement in the prevention of illicit drug use is far from settled, because no study has shown that knowledge mediates intervention effects. Thus, the role of knowledge in prevention should be further explored.

Limitations

This study has several limitations that could explain part of the imprecision of results. Refusal skills and normative beliefs are complex constructs, and they were assessed with a single item. Another limitation is the small effect sizes of mediational effects, which suggests that other factors not captured by this evaluation are at work. In addition, the geographical diversity of the sample might have moderated the intervention effects, concurring in reducing their size. A conceptual limit is related to the timing of assessments.

Despite the common praxis of interpreting a parallel change in the mediators and in the outcome as indicative of mediation effects, we acknowledge the inherent weakness of inferring causal relations with this approach [34], because the lack of lag time between the mediators change and the outcomes could make the direction of the effects uncertain. However, the experimental design and the explicit a priori hypothesis that manipulating individuals' attitudes, skills, and risk perceptions may subsequently affect behaviors make it more plausible that the observed cognitive and skills modifications led to modifications in drug use, rather than vice versa. The different rate of attrition in the experimental and control condition might have reduced the advantage of randomization. However, in a previous sensitivity analysis [21], we found no evidence that the attrition would explain the association between program and outcomes variables. For the results of mediation analyses to be biased, it would be necessary for attrition to be selectively associated with change in the mediators, which is unlikely.

Tests of significance were not corrected for multiple testing, using a Bonferroni adjustment. This increased the risk that some results may have arisen by chance. However, this correction has been criticized for being too conservative and for leading to Type II error [35e37]. Moreover, consistency between the observed mediating mechanisms and the theoretical model underpinning the development of the curriculum speaks against an important role of chance.

Implications

Our study suggests that positive attitudes toward drugs, normative perceptions of use among peers, and refusal skills might explain most of the effectiveness of interventions based on social influence. On the other hand, targeting other mediators, such as beliefs or knowledge, may not contribute to the effectiveness of such programs. These results should be taken into account to create cost-effective interventions.

Acknowledgments

The authors acknowledge the contribution of Gregor Burkhart of the European Monitoring Centre for Drugs and Drug Addiction, for constant support to the EU-Dap project and for precious advice on scientific publications. This article is dedicated to the memory of Professor Silvia Ciairano, who strongly supported the collaboration between researchers that made this article possible. Preliminary results of this report were presented at the XXIVth conference of the European Association for Research in Adolescence, 2012. The EU-Dap Study Group includes: Barbara Zunino, Federica Vigna-Taglianti, G. Luca Cuomo, Serena Vadrucci, Laura Vitale, Silena Salmaso (Piedmont Centre for Drug Addiction Epidemiology, Turin, Italy); Karl Bohrn, Sebastian Bohrn (Institut für Sozial und Gesundheitspsychologie, Wien, Austria); Erwin Coppens, YannickWeyts (De Sleutel, Merelbeke, Belgium); Peer van der Kreeft, Johan Jongbloet (University College, Ghent, Belgium); Juan Carlos Melero, Tatiana Perez, Laura Varona, Oihana Rementeria (EDEX, Bilbao, Spain); Gudrun Wiborg (IFT-Nord, Kiel, Germany); MaroVasara, Maria Kyriakidou, Gabriela Terzopoulou (Pyxida, Thessaloniki, Greece); Sara Sanchez, Charlotte Jansson, Rosaria Galanti (Department of Public Health Sciences, Karolinska Institutet, Sweden); FabrizioFaggiano (Department of Translational Medicine, Avogadro University, Novara, Italy); Leila Fabiani (Department of Life, Health and Environmental Sciences, University of Aquila, Italy); and Caterina Pesce (Department of Human Movement and Sport Sciences, "Foro Italico" University, Rome, Italy).

References

[1] Faggiano F, Vigna-Taglianti FD, Versino E, et al. School-based prevention for illicit drugs use: A systematic review. Prev Med 2008;46:385-96.

[2] Tobler NS, Roona MR, Ochshorn P, et al. School-based adolescent drug prevention programs: 1998 meta-analysis. J Primary Prevention 2000;20: 275-336.

[3] Catalano RF, Fagan AA, Gavin LE, et al. Worldwide application of prevention science in adolescent health. Lancet 2012;379:1653e64. [4] Cuijpers P. Three decades of drug prevention research. Drug Educ Prev Polic 2003;10:7-20.

[5] Faggiano F, Giannotta F, Allara E. The science of prevention: an international perspective. In: Sloboda Z, Petras H, eds. Advances in Prevention Science. Vol. 1: Defining Prevention Science. New York: Springer; 2014.

[6] Mackinnon DP, Dwyer JH. Estimating mediated effects in prevention studies. Evaluation Rev 1993;17:144-58.

[7] Botvin GJ, Dusenbury L, Baker E, et al. Smoking prevention among urban minority youth: Assessing effects on outcome and mediating variables. Health Psychol 1992;11:290-9.

[8] Botvin GJ, Griffin KW, Diaz T, Ifill-Williams M. Drug abuse prevention among minority adolescents: Posttest and one-year follow-up of a schoolbased preventive intervention. Prev Sci 2001;2:1-13.

[9] McNeal RB, Hansen WB, Harrington NG, Giles SM. How All Stars works: An examination of program effects on mediating variables. Health Educ Behav 2004;31:165-78.

[10] WynnSR, Schulenberg J, Maggs JL, Zucker RA. Preventing alcohol misuse: The impact of refusal skills and norms. Psychol Addict Behav 2000;14:36-47.

[11] Donaldson SI, Graham JW, Piccinin AM, Hansen WB. Resistance-skills training and onset of alcohol use: Evidence for beneficial and potentially harmful effects in public schools and in private Catholic schools. Health Psychol 1995;14:291-300.

[12] Teasdale B, Stephens PC, Sloboda Z, et al. The influence of program mediators on eleventh grade outcomes for seventh grade substance users and nonusers. Drug Alcohol Depend 2009;102:11-8.

[13] Bate S, Stigler M, Thompson M, et al. Psychosocial mediators of a schoolbased tobacco prevention program in India: Results from the first year of Project MYTRI. Prev Sci 2009;10:116-28.

[14] Liu L, Flay B, Aban Aya Investigators. Evaluating mediation in longitudinal multivariate data: Mediation effects for the Aban Aya Youth Project Drug Prevention Program. Prev Sci 2009;10:197-207.

[15] Orlando M, Ellickson PL, McCaffrey DF, Longshore DL. Mediation analysis of a school-based drug prevention program: Effects of Project ALERT. Prev Sci 2005;6:35-46.

[16] Botvin GJ, Schinke SP, Epstein JA, et al. Effectiveness of culturally focused and generic skills training approaches to alcohol and drug abuse prevention among minority adolescents: Two-year follow-up results. Psychol Addict Behav 1995;9:183-94.

[17] Wynn SR, Schulenberg J, Kloska DD, Laetz VB. The mediating influence of refusal skills in preventing adolescent alcohol misuse. J Sch Health 1997; 67:390-5.

[18] Komro KA, Perry CL, Williams CL, et al. How did Project Northland reduce alcohol use among young adolescents? Analysis of mediating variables. Health Educ Res 2001;16:59-70.

[19] Preacher K, Hayes A. Asymptotic and resampling strategies for assessing and comparing indirect effects in multiple mediator models. Behav Res Methods 2008;40:879-91.

[20] Kreeft PVD, Wiborg G, Galanti MR, et al. "Unplugged": A new European school programme against substance abuse. Drugs Educ Prev Policy 2009; 16:167-81.

[21] Faggiano F, Galanti MR, Bohrn K, et al. The effectiveness of a school-based substance abuse prevention program: EU-Dap cluster randomised controlled trial. Prev Med 2008;47:537-43.

[22] Hawkins JD, Jie G, Karl H, et al. Long-term effects of the Seattle Social Development Intervention on school bonding trajectories. Appl Dev Sci 2001;5:225-36.

[23] Catalano RF, Oesterle S, Fleming CB, Hawkins JD. The importance of bonding to school for healthy development: Findings from the Social Development Research Group. J Sch Health 2004;74:252-61.

[24] Wenzel V, Weichold K, Silbereisen RK. The life skills program IPSY: Positive influences on school bonding and prevention of substance misuse. J Adolesc 2009;32:1391-401.

[25] Catalano RF, Kosterman R, Hawkins JD, et al. Modeling the etiology of adolescent substance use: A test of the social development model. J Drug Issues 1996;26:429-55.

[26] Giannotta F, Özdemir M. School bonding and alcohol use in Italian early adolescents: What comes first? Merrill Palmer Quarterly 2013;59: 280-303.

[27] Faggiano F, Richardson C, Bohrn K, et al. A cluster randomized controlled trial of school-based prevention of tobacco, alcohol and drug use: The EUDap design and study population. Prev Med 2007;44:170-3.

[28] Stead M, Stradling R, Macneil M, et al. Implementation evaluation of the Blueprint multi-component drug prevention programme: Fidelity of school component delivery. Drug Alcohol Rev 2007;26:653-64.

[29] Brown MW, Cudeck R. Alternative ways of assessing model fit. In: Bollen K, Long S, eds. Testing structural equation models. Newbury Park (NJ): Sage; 1993, pp. 445-55.

[30] Hu L, Bentler PM. Fit indices in covariance structure modeling: Sensitivity to underparameterized model misspecification. Psychol Methods 1998;3: 424-53.

[31] Bandura A. Social Learning Theory. Englewood Cliffs (NJ): Prentice-Hall; 1977.

[32] Fishbein M. A theory of reasoned action: Some applications and implications. Nebraska Symposium on Motivation 1979;27:65-116.

[33] Avi K, Hanoch F. Motivation and identity: The relations of action and development in educational contextsdan introduction to the Special Issue. Educ Psychol 2009;44:73-7.

[34] MacKinnon DP, Fairchild AJ, Fritz MS. Mediation analysis. Ann Rev Psychol 2007;58:593.

[35] Nakagawa S. A farewell to Bonferroni: The problems of low statistical power and publication bias. Behav Ecol 2004;15:1044-5.

[36] Moran MD. Arguments for rejecting the sequential Bonferroni in ecological studies. Oikos 2003;100:403-5.

[37] Perneger TV. What's wrong with Bonferroni adjustments. BMJ 1998;316: 1236.

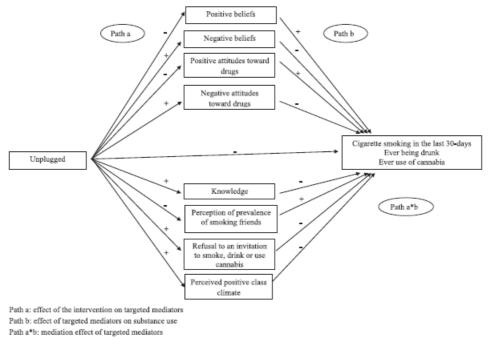




Table 1 Measures and their psychometric properties

Measure	Question and items	Answers	Cronbach α (at baseline and posttest	
Tobacco use	How many times (if any) have you smoked cigarettes during the past 30 days?	0		
Drunkenness	How many times (if any) have you been drunk from drinking alcoholic beverages in your	1-2		
	lifetime?	3-5		
Cannabis use	How many times (if any) have you used marijuana or hashish in your lifetime?	6-9		
		10-19		
		20-29		
		≥ 30		
Positive attitudes	Here are some statements that people have made about illegal substances. How much do	4-points Likert scale:	.7079	
toward drugs	you agree with the following opinions on drugs?	Strongly agree (= 4)		
	Using drugs can be a pleasant activity.	Agree (= 3)		
	Using drugs is fun.	Disagree $(=2)$		
	Many things are much more risky than trying drugs.	Strongly disagree (= 1)		
	Drugs help people to have experience life in full.			
Norativo attitudor	The police should not be annoying young people who are trying drugs. A young person should never try drugs.	A point Likest scale:	.7882	
Negative attitudes toward drugs	Everyone who tries drugs eventually regrets it.	4-point Likert scale: Strongly agree (= 1)	,70-,02	
toward drugs	The laws about drugs should be made stronger.	Agree (= 2)		
	Drug use is one of the biggest evils in the country.	Disagree (= 3)		
	Schools should teach about the real hazards of taking drugs	Strongly disagree (= 4)		
	To experiment with drugs is to give away control of your life.	Strongry usagree (= 4)		
Positive beliefs:	How likely is that each of the following would happen to you if you smoke cigarettes in the	4-point Likert scale:	.7880	
tobacco	next month?	Very likely (= 4)		
	Feel more relaxed	Likely $(= 3)$		
	Have more fun	Unlikely $(= 2)$		
	Be more popular	Very unlikely $(= 1)$		
	Be more confident and outgoing			
Negative beliefs:	Get into trouble with parents	4-point Likert scale:	.6667	
tobacco	Have problems with my friends	Very likely (= 1)		
	Become an addict	Likely $(=2)$		
	Have money problems	Unlikely $(= 3)$		
		Very unlikely $(= 4)$		
Positive beliefs:	Feel more relaxed	4-point Likert scale:	.8284	
alcohol	Have more fun	Very likely (= 4)		
	Be more popular	Likely (=3)		
	Forget my troubles	Unlikely $(= 2)$		
	Be more confident and outgoing	Very unlikely $(= 1)$		
Negative beliefs:	Do badly in school	4-point Likert scale:	.8284	
alcohol	Get into trouble with parents	Very likely (= 1)		
	Have problems with my friends	Likely $(= 2)$		
	Become an addict	Unlikely $(= 3)$		
	Have money problems	Very unlikely (= 4)		
Positive beliefs:	Feel more relaxed	4-point Likert scale:	.8486	
cannabis	Have more fun	Very likely $(= 4)$		
	Be more popular	Likely $(= 3)$		
	Be more confident and outgoing	Unlikely $(= 2)$		
		Very unlikely $(= 1)$		
Negative beliefs:	Get into troubles with police	4-point Likert scale:	.8687	
cannabis	Have problems in school	Very likely $(= 1)$		
	Get into trouble with parents	Likely $(= 2)$		
	Have problems with my friends	Unlikely $(= 3)$		
	Become an addict	Very unlikely $(= 4)$		
	Have money problems			
Knowledge about	Mark whether you think it is correct or not:	Yes		
tobacco	Nicotine is the substance in cigarettes that causes lung cancer.	No		
	One needs to smoke several cigarettes per day during many years to become addicted.	Don't know (Correct		
Knowledge about	Women have lower tolerance to alcohol than men.	answer = 1;		
alcohol	It takes about half an hour to eliminate from the body the amount of alcohol contained	wrong $= 0$).		
	in a can of strong beer.			
Knowledge about	Smoking marijuana does not cause physical dependence.			
cannabis Referent ability	High consumption of hash or marijuana decreases the production of sexual hormones.	A maintail Brook and		
Refusal skills:	You and the same friend are studying hard for an important test at school the day after. Both	4-point Likert scale;		
tobacco	of you feel stressed and need to calm down. Your friend suggests a cigarette would help,	Very likely $(= 4)$		
Defined shills	and offers one. Do you accept? The day after you both page the test and feel that it is now time to calebrate. You still have	Likely $(= 3)$		
Refusal skills: alcohol	The day after, you both pass the test, and feel that it is now time to celebrate. You still have	Unlikely $(= 2)$		
alconol	some pocket money left, and the liquor store is nearby. Would you buy some alcohol	Very unlikely (= 1)		
Refusal skills:	(beer, wine) to celebrate? You and your best friend are at a party where you meet new people, and you feel you really.			
cannabis	You and your best friend are at a party where you meet new people, and you feel you really			
cannabis	want to get to know them. Someone offers to smoke hash together. Your friend accepts. Do you?			
Perception of	When you answer this question, think about the friends with whom you spend most of your	None		
number of	leisure time.	Less than half		
friends who use	How many of them smoke cigarettes?	About half		
Contraction of the second second	How many of them get drunk?	More than half		
	How many of them use marijuana or other drugs?	All of them		
		I don't know		
Class climate	How much do you agree with the following descriptions of your school?	4-point Likert scale:	.6874	
	The students in my class enjoy being together.	Strongly agree $(= 1)$		
	Most of the students in my class are kind and helpful.	Agree $(=2)$		
	Other students accept me as I am.	Disagree (= 3)		
		Strongly disagree (= 4)		

Table 2

Standardized effects (β and standard errors) of Path a, Path b, and Path a*b of multilevel multiple mediation model on youth cigarettes use in the past 30 days (controlling for age, gender, and baseline levels of mediators and outcome), short-term follow-up

Mediator	Path a		Path b		Path a*b	
	β (SE)	p Value	β (SE)	p Value	β (SE)	p Value
Whole sample (n = 6,972), direct effect: β 014, S	E .008, p = .092			100		
Positive attitudes toward drugs	041 (.020)	.038	.075 (.014)	.000	003 (.002)	.070
Negative attitudes toward drugs		n.s.	.025 (.011)	.022		n.s.
Positive beliefs toward tobacco	044 (.021)	.034		n.s.		n.s.
Negative beliefs toward tobacco	029 (.017)	.086		n.s.		n.s.
Knowledge about tobacco	.049 (.021)	.022		n.s.		n.s.
Refusal skills for tobacco	030 (.015)	.040	280 (.016)	.000	008 (.004)	.040
Perception of number of friends who smoke	051 (.020)	.010	.071 (.009)	.000	004 (.002)	.016
Perception of positive class climate	047 (.021)	.022		n.s.		n.s.
No smokers (n = 4,511), direct effect: n.s.						
Positive attitudes towards drugs		n.s.	.101 (.027)	.000		п.s.
Negative attitudes towards drugs		n.s.	.048 (.020)	.014		n.s.
Positive beliefs tobacco	039 (.021)	.064		n.s.		n.s.
Negative beliefs tobacco		n.s.		n.s.		n.s.
Knowledge about tobacco	.056 (.022)	.010	035 (.015)	.018	002 (.001)	.070
Refusal skills tobacco	028 (.017)	n.s.	289 (.027)	.000		n.s.
Perception of number of smokers friends	047 (.022)	.032	.069 (.015)	.000	003 (.002)	.058
Perception of positive class climate	062 (.023)	.006		n.s.		n.s.
Smokers (n = 2,461), direct effect: β 027, SE .014	4, <i>p</i> = .062					
Positive attitudes towards drugs	075 (.027)	.006	.075 (.020)	.000	006 (.003)	.044
Negative attitudes towards drugs		n.s.		n.s.		n.s.
Positive beliefs tobacco		n.s.		n.s.		n.s.
Negative beliefs tobacco	040 (.023)	.084		n.s.		n.s.
Knowledge about tobacco		n.s.	.041 (.015)	.006		n.s.
Refusal skills tobacco	041 (.023)	.066	.324 (.018)	.000	013 (.007)	.066
Perception of number of smokers friends	055 (.026)	.038	.130 (.018)	.000	007 (.003)	.044
Perception of positive class climate		n.s.		n.s.		n.s.

Model fit. Total sample: χ^2 (84) = 741.154; comparative fit index = .96; root mean square error of approximation = .033, standardized root-mean-square residual = .061. No smokers: χ^2 (75) = 315.548; comparative fit index = .95; root mean square error of approximation = .027; standardized root-mean-square residual = .039. Smokers: χ^2 (84) = 343.310; comparative fit index = .96; root mean square error of approximation = .035; standardized root-mean-square residual = .050. n.s. = not significant; SE = standard error,

Table 3

Standardized effects (β and standard errors) of Path a, Path b, and Path a*b, of multilevel multiple mediation model on youths' ever being drunk (controlling for age, gender, and baseline levels of mediators and outcome), short-term follow-up

Mediator	Path a		Path b		Path a*b	
	β (SE)	p Value	β (SE)	p Value	β (SE)	p Valu
Whole sample ($n = 6,972$), direct effect: n.s.						
Positive attitudes toward drugs	040 (.019)	.036	.126 (.017)	.000	005 (.003)	.046
Negative attitudes toward drugs		n.s.	.027 (.015)	.078		n.s.
Positive beliefs toward alcohol	038 (.018)	.040	.037 (.012)	.002	001 (.001)	.096
Negative beliefs toward alcohol		n.s.	.069 (.014)	.000		n.s.
Knowledge about alcohol	.153 (.017)	.000		n.s.		n.s.
Refusal skills alcohol	032 (.018)	.072	.151 (.016)	.000	005 (.003)	.078
Perception of number of drunk friends		n.s.	.067 (.010)	.000		n.s.
Perception of positive class climate	047 (.021)	.022	.022 (.010)	.034		n.s.
Never drunk lifetime (n = 5,295), direct effect	:: n.s.					
Positive attitudes toward drugs		n.s.	.128 (.023)	.000		n.s.
Negative attitudes toward drugs		n.s.	.034 (.020)	.096		n.s.
Positive beliefs toward alcohol		n.s.	.039 (.019)	.040		n.s.
Negative beliefs toward alcohol		n.s.	.057 (.020)	.004		n.s.
Knowledge about alcohol	.164 (.020)	.000		n.s.		n.s.
Refusal skills for alcohol		n.s.	.206 (.021)	.000		n.s.
Perception of number of drunk friends		п.s.	.077 (.013)	.000		n.s.
Perception of positive class climate	045 (.021)	.034	.035 (.015)	.018		n.s.
Ever drunk lifetime (n = 1677), direct effect:	n.s.					
Positive attitudes toward drugs	091 (.030)	.002	.159 (.030)	.000	015 (.008)	.008
Negative attitudes toward drugs	069 (.030)	.020		n.s.		n.s.
Positive beliefs toward alcohol	084 (.028)	.004	.072 (.024)	.002	006 (.003)	.038
Negative beliefs toward alcohol		n.s.	.131 (.022)	.000		n.s.
Knowledge about alcohol	.128 (.029)	.000		n.s.		n.s.
Refusal skills for alcohol	055 (.030)	.068	.132 (.025)	.000	007 (.004)	.094
Perception of number of drunk friends	059 (.031)	.044	.087 (.022)	.000	005 (.003)	.076
Perception of positive class climate	049 (.030)	n.s.		n.s.		n.s.

Model fit. Total sample: χ^2 (84) = 717.911; comparative fit index = .96; root mean square error of approximation = .033; standardized root-mean-square residual = .059. No drinkers: χ^2 (75) 427.78; comparative fit index = .95; root mean square error of approximation = .030; standardized root-mean-square residual = .043. Drinkers: χ^2 (84) = 269.39; comparative fit index = .96; root mean square error of approximation = .036; standardized root-mean-square residual = .053.

n.s. = not significant: SE = standard error.

Table 4 Standardized effects (β and standard errors) of Path a, Path b, and Path a*b, of multilevel multiple mediation model on youths' ever-use of cannabis (controlling for age, gender, and baseline levels of mediators and outcome), short-term follow-up

Mediator	Path a		Path b		Path a*b	
	β (SE)	p Value	β (SE)	p Value	β (SE)	p Value
Whole sample (n = 6,972), direct effect: β 013	8; SE .011; p = .090					
Positive attitudes toward drugs	041 (.021)	.044	.121 (.016)	.000	005 (.003)	.060
Negative attitudes toward drugs		n.s.	.042 (.012)	.000		n.s.
Positive beliefs toward cannabis	050 (.019)	.006		n.s.		n.s.
Negative beliefs toward cannabis		n.s.	.044 (.012)	.000		n.s.
Knowledge about cannabis	.137 (.022)	.000		n.s.		n.s.
Refusal skills for cannabis	033 (.019)	.074	.180 (.016)	.000	006 (.003)	.078
Perception of number of friends who use	042 (.020)	.034	.048 (.008)	.000	002 (.001)	.048
Perception of positive class climate	047 (.021)	.022		n.s.		n.s.
Never-users lifetime ($n = 6,358$), direct effect:	$\beta = .025; SE .001; p = .$	090				
Positive attitudes toward drugs		n.s.	.149 (.022)	.000		n.s.
Negative attitudes toward drugs		n.s.	.039 (.020)	.050		n.s.
Positive beliefs toward cannabis	045 (.019)	.014		n.s.		n.s.
Negative beliefs toward cannabis		n.s.	.060 (.019)	.002		n.s.
Knowledge about cannabis	.141 (.023)	.000		n.s.		n.s.
Refusal skills for cannabis		n.s.	.217 (.025)	.000		n.s.
Perception of number of friends who use	035 (.019)	.066	.053 (.011)	.000	002 (.001)	.084
Perception of positive class climate	053 (.021)	.012		n.s.		n.s.
Ever-users lifetime ($n = 614$), direct effect: n.s.						
Positive attitudes toward drugs	137 (.047)	.004	.106 (.054)	.026		n.s.
Negative attitudes toward drugs	111 (.051)	.030	.094 (.042)	.050	010 (.006)	.096
Positive beliefs toward cannabis	085 (.048)	.076		n.s.		n.s.
Negative beliefs toward cannabis	1. S.	n.s.		n.s.		n.s.
Knowledge about cannabis	.130 (.050)	.010	.054 (.030)	.070	.007 (.004)	.090
Refusal skills for cannabis		n.s.	.247 (.046)	.000		n.s.
Perception of number of friends who use	081 (.048)	.094	.109 (.036)	.002		n.s.
Perception of positive class climate		n.s.	and the second	n.s.		n.s.

No smokers; χ^2 (75) = 547.823; comparative fit index = .95; root mean square error of approximation = .031; standardized root-mean-square residual = .052. Smokers: χ^2 (84) = 174.555; comparative fit index = .96; root mean square error of approximation = .042; standardized root-mean-square residual = .070. n.s. = not significant; SE = standard error. Model fit. Total sample: χ^2 (84) = 896.618; comparative fit index = .95; root mean square error of approximation = .037; standardized root-mean-square residual = .071.