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Case Control Study of Impulsivity, Aggression, Pesticide Exposure and Suicide Attempts Using Pesticides among Farmers

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Abstract

A case-control study was conducted to investigate associations between organophosphate pesticide (OP) exposure, aggression, impulsivity, and attempted suicide. Questionnaires were used to collect information; impulsivity and aggression were measured by the Barratt Impulsivity Scale (BIS) and the Aggression Inventory (AI). A greater number of OP symptoms was associated with an increased odds of a suicide attempt after adjusting for marital status and income ($OR = 1.45$; CI 1.14–1.86). Attempted suicide was significantly associated with high impulsivity scores (means: 72.4 vs. 60.6, $P < 0.0001$) and high aggression scores (means: 38.5 vs. 26.1, $P < 0.0001$). Suicide attempters had a higher number of OP exposure symptoms than controls and scored higher on scales of impulsivity and aggression.

A number of occupational studies have reported high rates of suicide in occupational groups^[1]. Farmers are among the occupational groups that have been reported to have high rates of suicide^[1]. Suicide and suicide attempts are associated with lack of impulse control, feelings of hopelessness, social isolation, and auto-aggression^[2]. In clinical studies and case reports, there is consistent evidence that acute poisoning from organophosphate pesticides may be associated with affective disorders, including anxiety and aggression^[3]. If organophosphate poisoning decreases impulse control and increases aggression in sensitive

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Author Contributions REN Shu Ping and STALLONES Lorann designed the study; BESELER L. Cheryl analyzed the data; REN Shu Ping, LYU Chun Ping, STALLONES Lorann and BESELER L. Cheryl all contributed in the writing process. PEI Jian Ru, LI Yu Ling, LI Jian Hui, REN Ming collected the data and interviewed the participants and helped established database.

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populations the result may well be increased suicidal behaviors. The purpose of this study was to use a case-control approach to investigate the relationships between attempted suicide and levels of impulsivity and aggression. Our primary hypothesis was that suicide attempters would have higher impulsivity and aggression scores compared to non-attempters. We hypothesized that levels of impulsivity and aggression could be used to discriminate between attempters and non-attempters. Our secondary hypothesis was that high occupational exposure to organophosphate pesticide (OP) would be associated with a suicide attempt and with higher levels of impulsivity and aggression compared to non-attempters with no history of pesticide exposure.

This study was approved by both the Institutional Review Board of Jilin University and by Colorado State University. Informed consent was obtained from all participants. Cases were identified using records from the local hospital. The records indicated that the cases attempted to kill themselves using pesticides.

Criteria for inclusion of cases included: (1) being able to take care of themselves without help; (2) being able to understand the information given by the chief of the Disease Control office and being able to answer the questions about voluntary participation in the study; and (3) the mental status of the cases was stable enough to participate in the study, as assessed by a licensed psychological consultant. Controls were those who lived in another village which was the same distance to the hospital in which the case was admitted. Controls were matched to cases by gender and by age (+/-2 years). Questionnaires were used to collect information on demographic characteristics, OP exposure and 11 symptoms of high OP exposure in the past two years, family history of mental problems and other medical conditions, and impulsivity and aggression.

The sample was 74.4% male and 25.6% female. The mean age of the sample was 46.2 (cases = 46.1 ± 13.0 ; controls = 46.2 ± 13.0 , $P = 0.682$) and did not differ due to matching. There were significant differences in marital status, family income, and duration of farm work between cases and controls (Supplement Table 1 available in www.besjournal.com). Although cases and controls were similar in their reporting of having had a pesticide illness, the median number of OP-related symptoms reported significantly differed between cases and controls (cases: 3 symptoms, range = 8; controls: 0 symptoms, range = 11; $P < 0.0001$). Each of the 43 cases reported at least one pesticide exposure symptom; this one symptom was either nausea/vomiting OR excessive sweating, indicating a high exposure to pesticides. In fact, the most frequently reported symptoms after exposure to OP pesticides were nausea and excessive sweating (Supplement Table 2 available in www.besjournal.com).

Cases were significantly more likely to report having been exposed to pesticides in the past two years than controls ($P = 0.021$). Only 44.2% of controls reported mixing, loading and applying pesticides in the past two years compared to 68.3% of cases ($P = 0.026$). Controls were significantly more likely to report removing protection (clothes, gloves, hats) before going home (50.0% vs. 4.7%, $\chi^2 = 22.1$, $P < 0.0001$) (Supplement Table 3 available in www.besjournal.com).

Suicide attempters had significantly higher impulsivity scores (means: 72.4 vs. 60.6, $P < 0.0001$) and aggression scores (means: 38.5 vs. 26.1, $P < 0.0001$) than controls in bivariate analyses. Impulsivity and aggression were significantly correlated ($\rho = 0.58$, $P < 0.0001$). Unmarried respondents had significantly higher impulsivity scores than married respondents ($P = 0.002$) and lower income respondents had higher impulsivity scores than those in the higher income category ($P = 0.002$). The same trend held for aggression scores; unmarried ($P = 0.005$) and low income ($P = 0.0003$) respondents scored significantly higher. Since being unmarried and having a low income were both associated with suicide attempts, models were adjusted for these two potential confounders. The number of OP-related symptoms was not associated with either impulsivity *OR* aggression, however, Barratt Impulsivity Scale (BIS) and Aggression Inventory (AI) scores were significantly higher in those that reported at least one pesticide exposure symptom compared to those who reported none (Supplement Tables 4–5 available in www.besjournal.com).

Both BIS and AI did significantly better than chance at predicting suicide attempters (BIS: $AUC = 0.78$, $se = 0.05$, 95% $CI 0.68–0.88$, $\chi^2 = 29.9$, $P < 0.0001$; AI: $AUC = 0.80$, $se = 0.05$, 95% $CI 0.70–0.89$, $\chi^2 = 37.4$, $P < 0.0001$) (Figure 1A–B). The optimal cutoff point was 69 for BIS and 33 for AI, where both sensitivity and specificity curves intersect, although this is only one approach to identifying optimal cut points. Adjusting the Receiver Operating Characteristic (ROC) models for gender, income *OR* number of reported pesticide-related symptoms did not alter these results. The number of pesticide symptoms was significantly higher in those who scored greater than 69 on the impulsivity scale ($P = 0.01$); symptom counts were not associated with the higher category of aggression.

Farm residents reporting a higher number of OP-related symptoms had a significantly higher odds of a suicide attempt ($OR = 1.27$, $CI 1.05–1.53$), and the effect was stronger after adjusting for marital status and income ($OR = 1.45$, $CI 1.14–1.86$) (Table 1). Suicide attempts were associated with increased impulsivity ($OR = 1.09$, $CI 1.04–1.14$). After adjusting for being unmarried and having a low income, the effect size was only slightly reduced ($OR = 1.07$, $CI 1.01–1.12$) (Table 1). Similarly, the odds of a suicide attempt were significantly elevated in those with higher aggression scores ($OR = 1.10$; $CI 1.05–1.16$); and remained significant after adjustment ($OR = 1.08$; $CI 1.02–1.14$).

In impulsivity models adjusted for the number of pesticide poisoning symptoms reported, the *OR* for pesticide symptoms and suicide attempt was 1.23 ($CI 1.01–1.50$) and 1.09 ($CI 1.04–1.14$) for impulsivity when entered in the same model (Table 1). Similarly, the aggression model adjusted for the number of pesticide exposure symptoms had an *OR* of 1.23 (1.01–1.49) for symptoms and an *OR* of 1.10 (1.05–1.15) for aggression (Table 2). Adjusting for marital status and income strengthened the association between symptoms and suicide attempt and only slightly decreased the effect size between the personality trait and suicide attempt (Table 2). Lastly, we tested for an interaction between the main effects (OP symptoms and personality trait) but with the addition of the interaction term in the model, the main effects were not significant (Table 2). A larger sample size is needed to explore the possibility of interactions between these associations in greater depth.

There were three important findings in our study. First, cases reported a significantly greater number of symptoms of pesticide exposure and greater occupational exposure to pesticides than controls. Each of the 43 cases had experienced at least a nausea or an excessive sweating episode associated with pesticide exposure. Clearly, the cases had higher exposure to pesticides than the controls, over half of whom ($n = 24$) reported no symptoms. This result is consistent with previous studies supporting a causal connection between pesticide exposure and mood disorders^[4], as well as ecological studies demonstrating higher suicide rates in agricultural communities^[5]. Further, results are consistent with studies in Sri Lanka and China that reported the storage of pesticides in homes was associated with suicide attempts^[6-8], indicating that pesticides are a potent risk factor for suicide attempts.

Second, cases scored significantly higher on the BIS and AI than controls. This result is consistent with Wesseling et al.^[9] findings among OP-poisoned banana workers who had increased suicidal ideation, anxiety and hostility compared with non-poisoned controls. Impulsivity is a strong predictor for suicide intent even when controlling for anxiety and depression severity^[10]. Therefore, OP exposure is likely to be exacerbating a trait already present in the farmer that increases the risk of a suicide attempt. Thus, the OP-poisoned participants may be at risk of future suicide attempts and may require ongoing screening to reduce this risk.

Third, the symptoms participants reported after being exposed to pesticides were nausea and excessive sweating. These symptoms are consistent with exposure to OPs, which suggests that cases had a history of OP exposure before they experienced their suicide attempt. Therefore, our results support the assertion that the OP exposure likely occurred prior to the suicide attempt.

Controls were significantly more likely to remove protective clothing after working with pesticides and more likely to remove their protections before going home than were cases. OPs can be absorbed through three routes: inhalation, skin absorption, and the gastrointestinal tract. Farmers check the weather forecast before applying OPs to plants or fruits. If the weather is windy, farmers will postpone the application of pesticides. Unless farmers consume OPs on purpose, or eat before washing their hands, the gastrointestinal tract is not a major route for OP exposure. Inhalation and skin absorption are the most likely routes of exposure. The greater care taken by controls when handling pesticides may be related to their lower level of impulsivity, the fact that they handle pesticides less frequently and are less comfortable using them, or because they were less depressed than the cases. This observation would benefit from further research because wearing protective equipment is a target of farm safety and health programs.

Although the mean scores on the BIS and AI were higher in those with pesticide symptoms, every case had one or more serious pesticide symptom. There was no way to disentangle the pesticide exposure effects from the attempted suicide. The higher scores may be related to the suicide attempt alone and not be due to the OP exposure, or the suicide attempt may be due to both the OP exposure and the presence of high impulsivity or aggression.

The study provided evidence for connections between impulse control disorders, pesticide exposures and suicide attempts. Due to local culture, potential cases not responding resulted in a low response rate, which is a serious limitation of the study. In this population, there is a strong stigma associated with suicide attempts, therefore, cases may not have been willing to reveal the reason for their hospitalizations to family and neighbors. A larger sample size would allow better control for other confounding factors, although we did not find evidence for significant confounding aside from marital status and income. In this study, OP exposure was assessed by exposure symptoms, such symptoms as excessively tired or headache which may be caused by other factors and therefore it is likely to overestimate the effects of OP exposure on suicide attempts. Moreover, questionnaire-based information was collected about OP exposure and OP-associated symptoms, which is also possible to result in recall bias. These are limitations of the study, a more physiological and toxicological assessment would improve certainty about actual exposures. The study did not assess the relationship between suicide attempts and amount of alcohol consumption nor did we assess substance abuse. Future work needs to assess these risk factors. Future work would also benefit from determining whether OP exposure mediates or moderates the association between impulsivity or aggression and suicide attempt. We could not assess BIS and AI independently from pesticide exposure and suicide attempt in this study. This information would help identify individuals who should be closely monitored for suicide ideation. Suicide attempters had a higher number of OP exposure symptoms than controls and scored higher on scales of impulsivity and aggression. Further work is needed to confirm these findings in a larger sample.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

Acknowledgments

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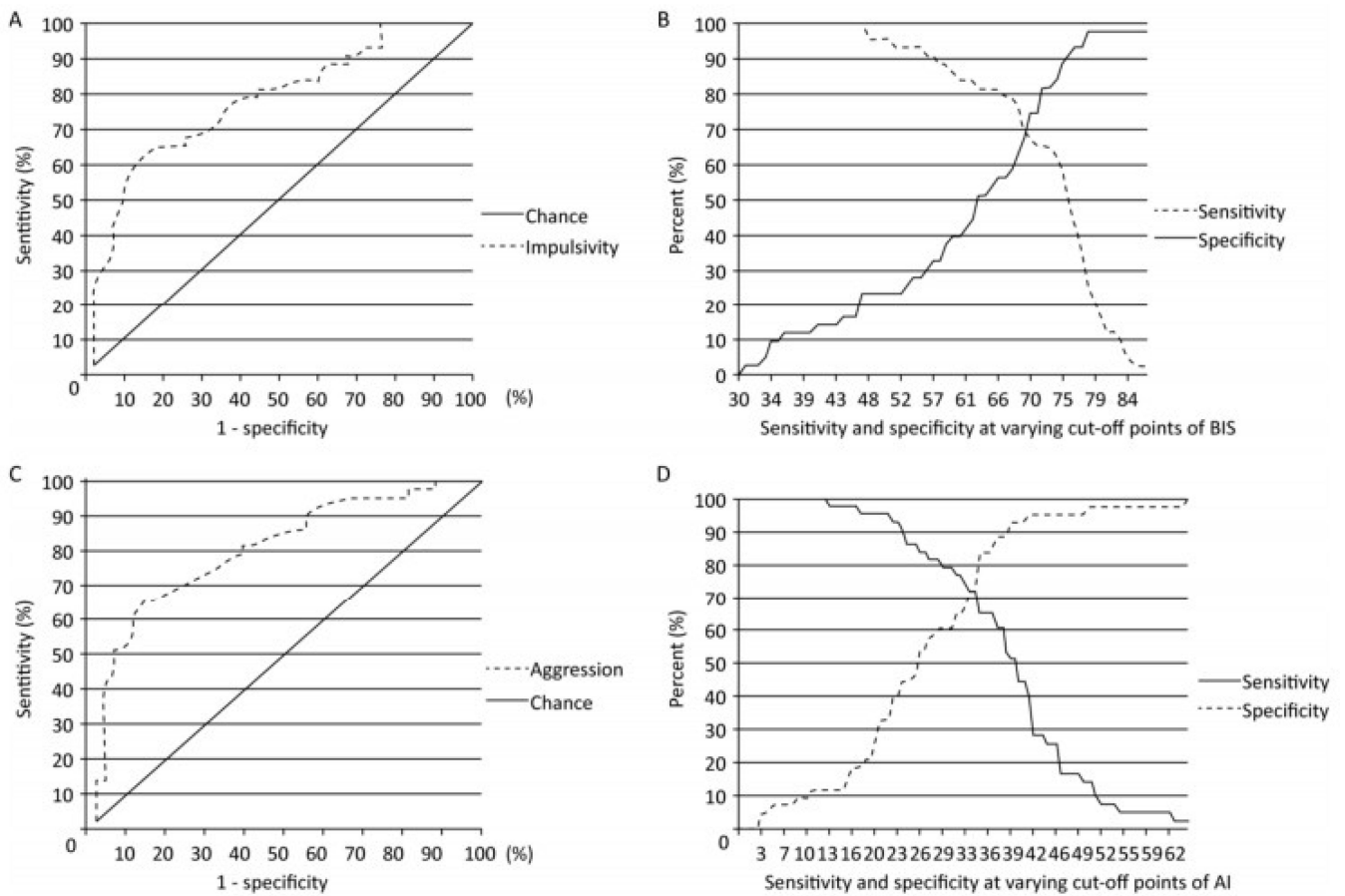


Figure 1. Receiver Operating Characteristic curve and sensitivity and specificity curve for Barrett's Impulsivity Scale and Aggression Inventory as a diagnostic measure of suicide attempt in 43 cases of suicide attempt and 43 controls from a farm resident sample, Jilin Province, China, 2008–2010.

Table 1.

Odds Ratios (*OR*) and 95% Confidence Interval (*CI*) from Logistic Regression Model for the Odds of having had a Suicide Attempt in 43 Cases of Suicide Attempt and 43 Controls

Logistic Regression Model	Unadjusted <i>OR</i> (95% <i>CI</i>)	Adjusted* <i>OR</i> (95% <i>CI</i>)
Number of pesticide poisoning symptoms	1.27 (1.05–1.53)	1.45 (1.14–1.86)
Impulsivity	1.09 (1.04–1.14)	1.07 (1.01–1.12)
Aggression	1.10 (1.05–1.16)	1.08 (1.02–1.14)

**Note.* Adjusted for income and marital status.

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Table 2.

Odds Ratios (*OR*) and 95% Confidence Interval (*CI*) from Logistic Regression Model with Main Effects of Symptoms and Personality Trait in 43 Cases of Suicide Attempt and 43 Controls

Logistic Regression Model	Unadjusted <i>OR</i> (95% <i>CI</i>)	Adjusted [*] <i>OR</i> (95% <i>CI</i>)
Number of pesticide poisoning symptoms	1.23 (1.01–1.50)	1.43 (1.10–1.85)
Impulsivity	1.09 (1.04–1.14)	1.06 (1.00–1.12)
Number of pesticide poisoning symptoms	1.23 (1.01–1.49)	1.45 (1.12–1.89)
Aggression	1.10 (1.05–1.15)	1.08 (1.01–1.15)

^{*}*Note.* Adjusted for income and marital status.