Child and Adolescent Psychiatry and Mental Health

RESEARCH

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Psychotic experiences and psychological distress in adolescents: an examination of longitudinal bidirectional effects across sex

(2024) 18.124



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Abstract

Background Although the co-occurrence of psychotic experiences (PEs) and psychological distress symptoms is growingly recognized in several previous studies, there is still a lack of literature, which clearly outlines how these two psychopathological entities affect each other over time. This study is intended to add to the literature by examining: (a) the longitudinal, bidirectional associations between PEs and psychological distress in a sample of Tunisian adolescents, and (b) whether these associations are moderated by sex.

Methods 510 adolescent students (mean age of 16.05 ± 1.01 years, 61.2% females) took part in a prospective longitudinal study. PEs and psychological distress were measured at three occasions over a one-year period. The cross-lagged panel modeling approach was adopted.

Results The current results showed that the experience of PEs at baseline tended to temporally precede prospective increases in psychological distress (at 6 months), which had in turn led to further exacerbation of psychological distress at 12 months of follow-up. Temporally primary psychological distress symptoms were not a significant predictor for the development of later psychotic symptoms in the whole sample. However, temporal patterns between adolescent distress and psychotic symptoms differed for girls and boys. Endorsing PEs at baseline was followed by greater psychological distress at 6 months, which was in turn associated with a significant increased risk of subsequent exacerbation of PEs at 12 months in boys, whereas psychological distress at 12-month follow-up was significantly predicted by pre-existing PEs in girls.

Conclusion These findings suggest that clinicians and support workers are recommended to take into account different social risk profiles for boys and girls when considering interventions to address PEs and distress in adolescents.

Keywords Psychotic experiences, Adolescents, Cross-lagged panel models, Sex moderation

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Introduction

Psychotic experiences (PEs) encompass beliefs, thoughts, or perceptions that are considered unreal, unusual or odd (i.e. sub-threshold hallucinations and delusions) occurring with less frequency and milder severity, in the absence of a psychotic disorder [1, 2]. They are positioned at the lower end of the extended psychosis continuum, and are frequently experienced by people from the general non-help-seeking population [2]. Prevalence estimates of PEs were reported to be 7.5% [3], varying substantially across cohorts and studies [4, 5]. They are most prevalent among children and adolescents [3]. Although some individuals who experience PEs are unlikely to ever convert to psychosis in their life, PEs were found to be significant predictors and risk markers for the development of later psychotic disorders [1, 6], and have therefore attracted considerable research interest in terms of their potential in detecting and predicting psychosis risk status in the general population [7]. In addition, PEs may denote young people's vulnerability to broader psychopathology, and qualifies as a transdiagnostic marker for a range of mental disorders [8].

Indeed, young people who endorse transitory PEs showed an increased likelihood to express both externalizing and internalizing symptoms [9-11], and those who experience persistent PEs were found to be more likely to suffer from adverse mental health outcomes and to perceive a need for care [1, 12]. According to a meta-analysis, the occurrence of PEs is associated with greater than twofold increase in the likelihood of reporting mental health service use [13]. The possible negative outcomes of PEs include chronic physical [14] and mental [15] health conditions, behavioral and addictive problems [16-20], suicidality [21, 22], reduced lifespan [23], disability [24], as well as psychological distress [25]. Therefore, it appears that PEs can potentially play a pivotal role in early detection, prevention, and intervention for a broad array of psychopathology, provided that pathways linking PEs and psychopathology are clearly understood. The present research proposes to shed light on these pathways by exploring the reciprocal relationships between PEs and psychological distress.

Bidirectional associations between PEs and psychological distress

Psychological distress is generally expressed as a state of emotional suffering characterized by negative emotions such as anxiety (e.g., feeling tense, restlessness), depression (e.g., sadness, hopelessness, lost interest) and stress [26]. A compelling and growing body of literature indicated that PEs can induce psychological distress, and conversely, psychological distress can contribute to the development or worsening of PEs. For instance, a metaanalysis by Healy et al. [27] revealed that children and adolescents who report PEs have a 3-fold greater risk of developing any mental disorder. A 10-year longitudinal study demonstrated that young people who reported PE are at a significantly increased risk for both a single lifetime mental disorder and multi-morbid disorders, with the most commonly diagnosis met being lifetime major depression followed by anxiety disorders [28]. The results of the Australian Young Minds Matter Survey showed that hallucinatory and delusional experiences endorsed by adolescents aged 14- to 17-year-olds were significantly related to an elevated likelihood of psychological distress [29]. Findings from World Health Organization World Mental Health (WMH) Surveys showed that temporally primary PEs were linked to the subsequent development of a range of mental health problems, including depression, anxiety and post-traumatic stress, and that, in contrast, numerous mental health problems were significant predictors of first-onset of PEs [30]. In contrast, Stochl et al. [31] argued against the concept of co-morbidity between psychotic phenomena and depression/anxiety symptoms in the general population. They rather suggest that, at a subclinical level, these manifestations reflect the same psychopathological concept and represent a unitary underlying continuum of "common mental distress", with PEs being at the upper end of severity [31].

At the same time, some controversial findings have been reported. For instance, Wigman et al. [32] evaluated 138 adolescent help-seekers treated for depression at 4 time-points over a two-year period, and found that PEs and depressive symptoms did not predict each other longitudinally. Nonetheless, the statistical power of these findings might have been limited by sample size [32]. To sum up, there is longitudinal evidence in diverse clinical and non-clinical populations showing that PEs are related to subsequent depression (e.g [33-37]), and that depression is related to later psychotic symptoms (e.g [38, 39]). These previous findings suggest that PEs and psychological distress are interrelated, and that arrows of causality could possibly run in both directions. However, this topic and field of study has paid relatively little attention to understanding temporal patterns of co-occurrence between PEs and psychological distress in non-clinical samples with milder forms of distress, which is determinant for evidence-based prevention and intervention practices. At last, previous evidence suggests that sex is an important consideration when examining the relationship between PEs and distress.

Moderating effect of sex

There are multiple plausible theoretical and empirical reasons why sex moderation is expected in this study. The sex disparity in rates of psychological distress, such as depression where girls predominate in a 1.95 ratio over boys, emerges during the adolescent years and persists into adulthood [40, 41]. The observed sex differences in psychological distress are due to the joint influence of biological, cognitive, affective, and sociocultural determinants [41]. Prior research has also pointed to differences in how males and females express and experience their psychotic symptoms that extend across the psychosis continuum [42]. Females tend to endorse more PEs [43]; however, opposite findings have also been reported [44, 45]. In addition, sex differences were noticed regarding aspects of psychotic symptoms, with female adolescents being more likely to experience persecutory ideation and subclinical hallucinations compared to males [46, 47]. Altogether, these observations advocate that sex differences in interrelationships between PEs and psychological distress are plausible. However, there is only a paucity of studies specifically focusing on how the associations between PEs and distress may differ across sex, and this research question is yet to be addressed. To fill this gap in the literature, the present study included sex as a moderating variable.

Rationale and purpose of the present study

Adolescence represents the peak age of emergence of mental illness [48], as well of PEs, and is therefore a critical period for inquiry into factors that pose a risk to mental health. The reciprocal time-lagged associations between PEs and psychological distress are still poorly understood and under-researched, as most prior research has examined unidirectional effects. Elucidating these associations in a community sample of adolescents offers an opportunity to gauge the extent to which the occurrence of PEs leads to psychological distress, and whether temporally primary distress predicts the subsequent first onset of PEs. This might provide significant clinical implications for prevention and early intervention, as research showed that the intersection between PEs and nonpsychotic symptoms/disorders confers substantial public mental health burden. Indeed, the co-occurrence of the two entities was found to be associated with great functional deficits (e.g [49]), higher proportions of lifetime suicide attempt reporting, more use of psychological treatment, and more elevated odds of mental health services use [50]. Furthermore, large crosscountry and cross-cultural differences have been seen in the way distress [51], PEs [52], and sex gaps in mental health [53] are expressed. Although cross-sectional associations between hallucinatory/delusional symptoms and increased anxiety, depression and stress have been documented in samples from Arab countries [18, 54-58], the directionality and nature of these associations remain to be disentangled, and particularly in the context of a longitudinal design. This study is intended to add to the literature by examining: (a) the longitudinal, bidirectional associations between PEs and psychological distress using cross-lagged panel modeling approach in a sample of Tunisian high-school adolescent students, and (b) whether these associations are moderated by sex. Guided by previous research, it is expected that there will be a bidirectional association between PEs and distress symptoms, such that PEs will act as contributor to distress, and that distress will be a significant predictor of later PEs. In addition, it is anticipated that positive associations between PEs and distress will be stronger in both directions for girls as compared to boys.

Methods

Sample and procedure

A prospective longitudinal study was performed over a 12-month period (April 2022-April 2023), by adopting a non-probability convenience sampling technique. The target population was high-school students from four high schools in Tunis, Tunisia. Students were approached in their classrooms after lectures. They were eligible to participate if they: (1) were aged 13–18 years, (2) had no personal history of psychosis or antipsychotic medication intake, (3) had a French language proficiency, and (4) have completed the three scheduled assessments. Of the 1692 students initially approached to participate, 510 were included in the final analyses (Further details on the study sample are provided in [59]).

Ethics

The study was performed following the Declaration of Helsinki for human research. Ethical approval was provided by the ethics committee of the Razi psychiatric hospital, Manouba, Tunisia. Before approaching students, approval was obtained from school authorities and the parent or legal guardian after a first contact held to introduce the research project. In addition, each participating student provided a voluntary oral informed assent before beginning the survey. All participants and their parent(s)/legal guardian(s) were informed that the study necessitates a three-point assessment procedure.

Measures

Participants took part of three-time-point assessments: at baseline (T0), 6 months (T1) and 12 months (T2). They were invited to complete a self-report paper-andpencil questionnaire in the French language composed of two sections. The language of the questionnaire was not considered as a barrier to participants' completion of the survey, as it represents the language of high-school studies in the country. The first section contained demographic information, while the second section contained two scales: The "Community Assessment of Psychic Experiences" (CAPE-42) and the Depression, Anxiety, Stress Scales-21 (DASS-21). Sociodemographic data collected consisted of age, sex (male, female), and residency (rural, urban). In addition, the socioeconomic status was reflected through the participants' monthly family income in Tunisia Dinars (i.e. <1000, 1000-2000, 2000-3000, >3000) as done in previous studies among Tunisian students (e.g [60-62]).

The CAPE-42

This is self-administered scale composed of 42 items that assess positive, negative and depressive symptoms through two dimensions (i.e., frequency of symptoms and related distress) [63]. The frequency dimension is scored on a four-point scale from 1 (never) to 4 (nearly always), while the distress dimension is rated from 1 (not distressed) to 4 (very distressed). Only the positive CAPE subscale, which is composed of 20 items, was used. Total scores for the CAPE positive subscale range from 20 to 80 on both dimensions. Higher scores reflect greater PEs. The French version adopted demonstrated excellent psychometric qualities [64], and was previously used in non-clinical Tunisian populations [65-67]. The present sample yielded Cronbach alpha values for the positive CAPE dimension ranging from 0.817 to 0.849 for the three time points.

The DASS-21

This is a 4-point Likert-type measure composed of 21 items, which are divided into three subscales of seven items each, reflecting the severity of anxiety, depression and stress symptoms [68]. Scores range from 0 to 21 for each dimension, and from 0 to 63 for the DASS-21 total score. The DASS-21 has showed good internal consistency in different non-clinical and clinical samples [69]. The DASS-21 also showed good psychometric qualities among adolescents across different countries [70]. For the current sample, the reliability of the DASS-21 total score was good (Cronbach's alpha ranging from 0.897 to 0.917 for the three time points).

Data analysis

We used SPSS AMOS v.28 to examine the bidirectional relationship between psychological distress and CAPE positive dimension over time and to assess model fit. The scores were considered normally distributed as their skewness and kurtosis values varied between +1.96, -1.96 [71]. We employed the structural equation modeling approach to test the hypothesized model. To address the primary purpose of the study, we tested a cross-lagged model, including both the autoregressive effects to examine the temporal stability of psychological distress and CAPE positive dimension across time, and the regression effects to examine the bidirectional relationship between psychological distress and CAPE positive effects included a path from psychological cal distress at Time 1 to Time 2 to Time 3, and a second

path from CAPE positive dimension at Time 1 to Time 2 to Time 3. The regression effects included a path from psychological distress at Time 1 to CAPE positive dimension at Time 2 to psychological distress at Time 3, and a path from CAPE positive dimension at Time 1 to psychological distress at Time 2 to CAPE positive dimensions at Time 3. For standardized Beta coefficients of the cross-lagged model, the values of 0.03, 0.07, and 0.12 correspond to small, moderate and high effect sizes respectively [72]. The results were adjusted over age and sex; age was a continuous variable, with higher values indicating older age in years; sex was coded as 0=male and 1=female. The model also examined synchronous correlations by allowing psychological distress and CAPE positive dimension to intercorrelate within each time point (represented by the curved, double-headed arrows in Fig. 1). We examined the model fit of each of these models and interpreted the significant parameter estimates. To address the secondary purpose of the study, we conducted multiple group analyses to determine whether the relationships between psychological distress and positive dimension differed by sex. We tested the cross-lagged models in each of the prior analyses, while controlling for age. In this set of analyses, the relative fit of an unrestricted model, which allowed parameters to be estimated freely across the two groups (boys versus girls), was compared to the fit of a constrained model that held the paths constant (fixed paths to be the same for both boys and girls). The fit of the models was assessed with the lack of significance (p < 0.05) of the chi-square test (χ^2), the comparative fit index (CFI), the root mean square error of approximation (RMSEA), and the standardized root mean square residual (SRMR). We reported SRMR because of the relatively small sample sizes (N < 250). χ^2 df \leq 5, RMSEA values \leq 0.08, CFI and TLI values \geq 0.90, and SRMR values less than 0.05 were used as cut-off points for acceptable model fit [72]. P < 0.05 was deemed statistically significant.

Results

Descriptive statistics

Five hundred ten participants took part of the study. The mean age of the sample was 16.05 ± 1.01 , with 61.2% females. Other characteristics of the sample are shown in Table 1. Table 2 shows the correlation matrix between the scores of positive PEs and psychological distress.

Comparison of variables between sexes

Higher means psychological distress and CAPE positive dimension scores at baseline, 6 and 12 months were significantly found in females compared to males (Table 3).

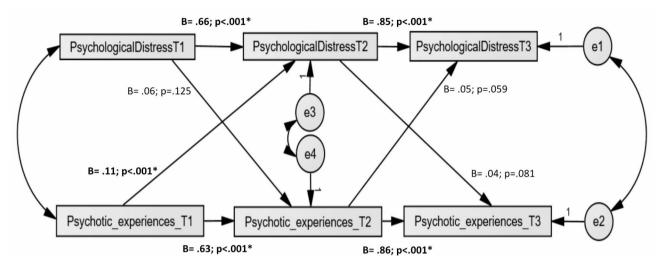


Fig. 1 Structural equation model examining the associations between psychological distress and positive dimensions in the total sample. Rectangles indicate observed variables. Small circles reflect residuals (e); Unidirectional arrows depict hypothesized directional associations. Models were adjusted over age and sex but are not presented visually for parsimony. (*Standardized* Beta = 0.06, p = 0.125)

 Table 1
 Sociodemographic and other characteristics of the participants (n = 510)

Sex	
Male	198 (38.8%)
Female	312 (61.2%)
Residency	
Rural	11 (2.2%)
Urban	499 (97.8%)
Family income (in Tunisian Dinars)	
< 1000	21 (4.1%)
1000–2000	132 (25.9%)
2000–3000	164 (32.2%)
> 3000	193 (37.8%)

Table 2 Correlation matrix

	$Mean \pm SD$	1	2	3	4	5
1. Psychological distress T1	21.32±11.86	1				
2. Psychological distress T2	20.83 ± 12.48	0.75***	1			
3. Psychological distress T3	20.11 ± 11.97	0.71***	0.88***	1		
4. Psychotic experiences T1	39.60 ± 8.74	0.51***	0.47***	0.43***	1	
5. Psychotic experiences T2	39.14 ± 9.07	0.40***	0.53***	0.50***	0.68***	1
6. Psychotic experiences T3	39.71 ± 9.48	0.38***	0.50***	0.52***	0.62***	0.88***

CAPE: Community Assessment of Psychic Experiences

*p<0.05; **p<0.01; ***p<0.001

Table 3 Comparison of variables between sexes

	Males	Females	t	df	p	95% CI of the difference
	(Mean±SD)	(Mean±SD)				
Age	16.04 ± 0.96	16.05 ± 1.05	- 0.09	508	0.932	-0.19; 0.17
Psychological distress T1	16.23 ± 10.44	24.55 ± 11.57	- 8.22	508	< 0.001	- 10.32; - 6.34
Psychological distress T2	15.42 ± 10.82	24.25 ± 12.26	- 8.52	508	< 0.001	- 10.86; - 6.79
Psychological distress T3	14.57 ± 10.23	23.62 ± 11.67	- 9.22	508	< 0.001	- 10.99; - 7.13
Psychotic experiences T1	37.51 ± 8.49	40.92 ± 8.65	- 4.38	508	< 0.001	- 4.95; - 1.89
Psychotic experiences T2	36.81 ± 9.06	40.61 ± 8.77	- 4.70	508	< 0.001	- 5.38; - 2.21
Psychotic experiences T3	37.31±9.60	41.23±9.10	- 4.64	508	< 0.001	- 5.58; - 2.26

Numbers in bold indicate significant \boldsymbol{p} values

Bidirectional associations between psychological distress and PEs (CAPE positive dimension)

Figure 1 presents the results of the autoregressive effects and cross-lagged associations between psychological distress and CAPE positive dimension (PEs), adjusted over age and sex. This model was adequate as shown by the following fit indices: $\chi^2/df = 33.08/9 = 3.68$, *p*<0.001; RMSEA=0.073 [90% CI 0.047, 0.100]; CFI=0.991; TLI=0.971 and SRMR=0.016. Autoregressive paths were significant; Time 1 psychological distress was positively and strongly associated with Time 2 psychological distress, Standardized Beta=0.66, p<0.001, and Time 2 psychological distress with Time 3 psychological distress, Standardized Beta=0.85, p < 0.001. In addition, PEs at Time 1 strongly predicted higher PEs at Time 2, Stan*dardized Beta* = 0.63, p < 0.001, which strongly predicted higher PEs at Time 3, *Standardized Beta*=0.85, *p*<0.001. Moreover, PEs at Time 1 significantly but weakly increased psychological distress at Time 2 (Standardized Beta=0.11, p < 0.001), whereas psychological distress at Time 1 was not significantly associated with PEs at Time 2 (Standardized Beta=0.06, p=0.125). Finally, psychological distress at Time 2 was not significantly associated with PEs at Time 3 (*Standardized Beta*=0.04, p=0.081), whereas PEs at T2 was not significantly associated with psychological distress at Time 3 (Standardized Beta = 0.05, p = 0.059).

Moreover, three significant indirect effects were observed in the total sample: (1) psychological distress at Time 2 weakly mediated the association between PEs at Time 1 and psychological distress at Time 3 (Beta=0.13; 95% CI 0.06, 0.19, p=0.004, (2) PEs at Time T2 strongly mediated the association between positive dimensions at Time 1 and Time 3 (Beta=0.55, 95% CI 0.46, 0.61, p=0.007, (3) psychological distress at Time 2 strongly mediated the association between psychological distress at Time 1 and Time 3 (Beta=0.56, 95% CI 0.49, 0.64, p=0.003).

Moderation analysis by sex

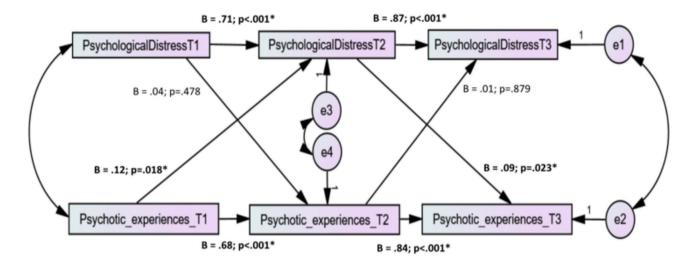
Figure 2 presents the results of the autoregressive effects and cross-lagged associations between psychological distress and CAPE positive dimension (PEs), stratified by sex. In the model related to males (Fig. 2A), the fit indices were adequate: $\chi^2/df=16.95/4=4.24$, p=0.002; RMSEA=0.128 [90% CI 0.069, 0.194]; CFI=0.988; TLI=0.953 and SRMR=0.019. Autoregressive paths were significant; Time 1 psychological distress was positively and strongly associated with Time 2 psychological distress, *Standardized Beta*=0.71, p<0.001, and Time 2 psychological distress with Time 3 psychological distress, *Standardized Beta*=0.87, p<0.001. In addition, PEs at Time 1 strongly predicted higher PEs at Time 2, *Standardized Beta*=0.68, p<0.001, which strongly predicted higher PEs at Time 3, *Standardized Beta*=0.84, p < 0.001. Moreover, PEs at Time 1 significantly but weakly increased psychological distress at Time 2 (*Standardized Beta*=0.12, p=0.018), whereas psychological distress at Time 1 was not significantly associated with PEs at Time 2 (*Standardized* Beta=0.04, p=0.478). Finally, psychological distress at Time 2 was significantly associated with PEs at Time 3 (*Standardized Beta*=0.09, p=0.023), whereas PEs at T2 was not significantly associated with psychological distress at Time 3 (*Standardized Beta*=0.09, p=0.023), whereas PEs at T2 was not significantly associated with psychological distress at Time 3 (*Standardized Beta*=0.09, p=0.023).

The total indirect effect from PEs at T1 and T3 (Standardized Beta=0.58; 95% CI 0.48, 0.68; p=0.005) was significant, with pathways PEs at T1 \rightarrow PEs at T2 \rightarrow PEs at T3 and PEs at T1 \rightarrow psychological distress at T2 \rightarrow PEs at T3 being both significant. Furthermore, the total indirect effect from psychological distress at T1 and T3 (Standardized Beta=0.61; 95% CI 0.50, 0.73; p=0.003) was significant, with pathway psychological distress at T1 \rightarrow psychological distress at T2 \rightarrow psychological distress at T3 being significant, while pathway psychological distress at T1 \rightarrow PEs at T2 \rightarrow psychological distress at T1 was not significant.

In the model related to females (Fig. 2B), the fit indices were adequate: $\chi^2/df = 10.20/4 = 2.55$, p = 0.037; RMSEA=0.071 [90% CI 0.016, 0.126]; CFI=0.995; TLI=0.982 and SRMR=0.013. Autoregressive paths were significant; Time 1 psychological distress was positively and strongly associated with Time 2 psychological distress, *Standardized Beta*=0.63, p<0.001, and Time 2 psychological distress with Time 3 psychological distress, Standardized Beta=0.82, p < 0.001. In addition, PEs at Time 1 strongly predicted higher PEs at Time 2, Stan*dardized Beta* = 0.61, p < 0.001, which strongly predicted higher PEs at Time 3, *Standardized Beta*=0.86, *p*<0.001. Moreover, PEs at Time 1 significantly but weakly increased psychological distress at Time 2 (Standardized Beta=0.11, p=0.014), whereas psychological distress at Time 1 was not significantly associated with PEs at Time 2 (Standardized Beta=0.07, p=0.177). Finally, psychological distress at Time 2 was not significantly associated with PEs at Time 3 (*Standardized Beta*=0.01, p=0.720), whereas PEs at T2 was significantly associated with psychological distress at Time 3 (Standardized Beta=0.07, p = 0.041).

The total indirect effect from PEs at T1 and T3 (Standardized Beta=0.52; 95% CI 0.43, 0.61; p=0.004) was significant, with pathways PEs at T1 \rightarrow PEs at T2 \rightarrow PEs at T3 being significant while pathway PEs at T1 \rightarrow psychological distress at T2 \rightarrow PEs T3 was not significant. Furthermore, the total indirect effect from psychological distress at T1 and T3 (Standardized Beta=0.52; 95% CI 0.44, 0.62; p=0.003) was significant, with pathway psychological distress at T1 \rightarrow psychological distress at T2

A) Males



B) Females

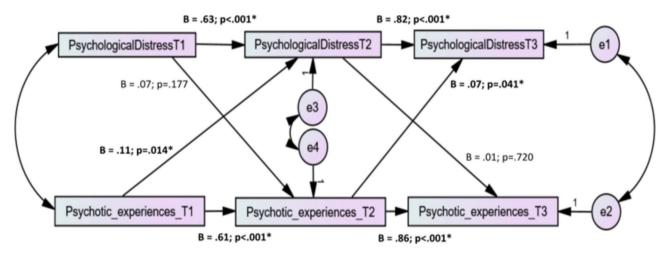


Fig. 2 Structural equation model examining the associations between psychological distress and positive dimensions in males (A) and females (B). Rectangles indicate observed variables. Small circles reflect residuals (e); Unidirectional arrows depict hypothesized directional associations. Models were adjusted over age and sex but are not presented visually for parsimony

→ psychological distress at T3 being significant, while pathway psychological distress at T1 → PEs at T2 → psychological distress at T3 was not significant. Finally, the total indirect effect of PEs at T1 and psychological distress at T3 (standardized Beta=0.14; 95% CI 0.05, 0.21; p=0.007) was significant, with pathway PEs at T1 → psychological distress at T2 → psychological distress at T3 being significant, while pathway PEs at T1 → PEs at T2 → psychological distress at T3 was not significant.

Discussion

Even though the co-occurrence of PEs and psychological distress symptoms is growingly recognized in several major previous studies [33, 34, 73–75], the temporal sequence and causal relationships between these two psychopathological entities cannot be determined on the basis of the current state of the literature. This underlines the need for longitudinal studies investigating both directions between PEs and psychological distress, which may allow to improve understanding of the challenges confronted by adolescents with both conditions and intervene more effectively to promote their mental health. Our findings revealed that the model predicting general symptoms of psychological distress at 6 months after temporally primary PEs was significant, whereas no statistical significance was found for the reverse path of causality in the total sample. These findings, however, were moderated by sex, where endorsing PEs at baseline was followed by greater psychological distress at 6 months, which was in turn associated with a significant increased risk of subsequent exacerbation of PEs at 12 months in boys, whereas psychological distress at 12-month follow-up was significantly predicted by preexisting PEs in girls.

The current results showed that the experience of PEs at baseline tended to temporally precede prospective increases in psychological distress (at 6 months), which had in turn led to further exacerbation of psychological distress at 12 months of follow-up. This supports previous findings that subclinical psychotic symptoms can provoke negative emotions and distress in non-help-seeking community-based young people [27, 28]. However, temporally primary psychological distress symptoms were not a significant predictor (Standardized Beta=0.06, p=0.125) for the development of later psychotic symptoms in the whole sample, but non-significant effect is plausible to have arisen from a lack of statistical power. Previous findings showed that, in nonclinical adults, stress has a significant effect on the emergence of PEs [76], and dysphoria (a combination of depression and anxiety) is considered a potential contributor to the occurrence of positive psychotic symptoms [77-79]. Concurrent with the present findings, a prospective cohort study in a large community sample of adolescents and young adults with a 6-year follow-up demonstrated that PEs were linked to a significant increase in depressive symptoms at 18 years, but no link was shown between depressive symptoms at 12 years and PEs at 18 years, thus suggesting that depression in youth do not seem to enhance subsequent risk of PEs [73]. It is of note, however, that the moderating effect of sex was not explored in this study. Another study using cross-lagged path modeling was not able to confirm the hypothesized longitudinal bidirectional association between subclinical psychotic symptoms and depression in a clinical sample of adolescents with major depression [32].

Another relevant finding of the current study is that temporal patterns between adolescent distress and psychotic symptoms differed for girls and boys. Adolescent girls who initially presented with PEs had an elevated risk of developing later psychological distress (i.e., depression, anxiety and stress), and adolescent boys who firstly endorsed psychological distress were at higher risk of experiencing new onset psychotic symptoms over time. Yet, little is reported in the literature about whether any sex differences in the longitudinal associations between PEs and distress could prove to exist and be informative for prevention and intervention efforts. There is some evidence that females with psychotic disorders present with worse depressive symptoms [80] and have increased vulnerability to stress levels [81] than males. Inconsistent with our results, a recent study by Stainton et al. [82] demonstrated that depressive, anxiety, and stress symptoms contributed equally and significantly to the prediction of PEs in both male and female adolescents from the general population. In the present sample, PEs preceded subsequent increases in distress during follow-up in both boys and girls; whereas distress was a significant predictor of later PEs only in boys. Previous findings pointed to the complex nature of sex disparities in adolescent mental health, which remains largely poorly understood [53]. The sex-based differences observed in the prospective relationship between distress and PEs can be partly explained by a variety of socioenvironmental and cultural factors. A large national survey among Tunisian high school adolescents aged 16 to 18 years showed that boys reported significantly greater lifetime prevalence of cannabis use than girls (16.1% versus 2.5%) [83]. This might explain our findings, as cannabis is consistently shown to be a strong predictor of psychosis onset in vulnerable individuals [84]. In addition, male adolescents have a significantly higher likelihood of being victims of child abuse in Tunisia [85], which can in turn lead to increased proneness to develop psychotic symptoms in those who are more distressed [67, 86]. Finally, Tunisian males had poorer knowledge about mental illness and were less likely to seek help from professional sources than females, which can be explained by social norms and sex-based stereotypes, such as autonomy and strength, that are associated with negative attitudes toward helpseeking behaviors [87]. Failure to seek help or disclose distress can contribute to negative mental health consequences, including the onset of psychotic symptoms. Overall, the mixed findings call for further investigations of the sex moderation effect on relationships between psychological distress and psychosis in different populations and settings.

Study limitations

Our findings should be considered in light of a number of limitations. Data were gathered using convenience sampling and self-report measures, which may affect the representativeness of the sample and lead to response bias. Only community adolescents aged 13–18 years and from a single low-to-middle income Arab country (i.e. Tunisia) have been involved in the study, which may limit the generalization of findings to help-seeking and clinical adolescents, emerging adults, as well as those from other countries and cultures (e.g., high-income countries [88], non-Arab countries [89]). Further studies should seek to replicate the findings in these different samples. Certain potential confounding factors were not accounted for in this study's analyses, such as genetic load, negative symptoms, cognitive functioning, traumatic experiences, or substance use [1, 67, 90, 91]. Given the sample is skewed towards girls (61%), it may be that boys with greater distress / PEs were more likely to refuse or drop out. Finally, analysis using RI-CLPM methodology was not possible with the present dataset; therefore, between-person confounding could not be controlled for.

Clinical and research implications

This study builds on and expands previous research efforts by showing that adolescents who experience PEs are at a heightened risk for distressing outcomes in the form of depression, anxiety, and stress. Besides, distress seems to be, in itself, a worsening factor for PEs, contributing to their persistence and/or deterioration over time. These patterns of associations suggest that one factor could be an early marker for the other, and warrant routine screening for one outcome when the other is present, specifically in adolescent populations. Interestingly, the two factors were found to predict each other differently across sex, with persistent psychological distress at 6 months temporally preceding an exacerbation of PEs at 12 months in boys, and PEs at 6 months being at the origin of later psychological distress in girls. These findings suggest that clinicians and support workers are recommended to take into account different social risk profiles for boys and girls when considering interventions to address PEs and distress in adolescents. As for research perspectives, long-term follow-up studies (e.g., birth cohorts) still need to study the different possible pathways by which the two constructs could influence each other over longer periods of time. It is also crucial to investigate and better understand factors other than sex, such as a wider age range (adolescence versus early adulthood), that may moderate the relationships between psychosis and distress. Finally, the field may benefit from future experimental research testing whether targeting one of the two conditions (i.e., PEs and psychological distress) could reduce the incidence of the other in adolescents.

Conclusion

Using advanced methodological approaches, this study contributes to the body of knowledge on this topic by (1) exploring possible bidirectional influences of PEs and distress, and (2) examining potential moderating effects of sex in these relationships. Beyond there general significance, these findings also enhance our understanding of these dynamics within cultures that have been significantly underexplored, thereby contributing to the corpus of cross-cultural research in psychiatry. Results showed that the predictive associations between variables varied across sex. Baseline PEs were found to drive psychological distress at 6 months, which in turn has led to a further increase in PEs at 12 months among adolescent girls. The reverse path leading from reported psychological distress at 6 months to later aggravation of PEs was significant among adolescent boys. These findings have important implications for the understanding of temporal relationships between PEs and psychopathology in adolescence, and for effectively guiding the timing and selection of preventive and interventive strategies.

Acknowledgements

The authors would like to thank all participants.

Author contributions

FFR designed the study; LH processed the data; FFR and SH drafted the manuscript; SH carried out the analysis and interpreted the results; AAL and MC reviewed the paper for intellectual content; all authors reviewed the final manuscript and gave their consent.

Funding

None.

Data availability

The datasets generated and/or analyzed during the current study are not publicly available due to restrictions from the ethics committee but are available from the corresponding author on reasonable request.

Declarations

Ethics approval and consent to participate

Before approaching students, approval was obtained from school authorities and the parent or legal guardian after a first contact held to introduce the research project. In addition, each participating student provided a voluntary oral informed assent before beginning the survey. All participants and their parent(s)/legal guardian(s) were informed that the study necessitates a threepoint assessment procedure. The research protocol was approved by the ethics committee of the Razi psychiatric hospital, Manouba, Tunisia. The study was performed following the standards for medical research involving human subjects recommended by the Declaration of Helsinki for human research.

Consent for publication

Not applicable.

Competing interests

The authors have nothing to disclose.

Received: 17 February 2024 / Accepted: 30 September 2024 Published online: 03 October 2024

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