Supplementary Material for Associations of polygenic risk scores with posttraumatic stress symptom trajectories following combat deployment

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suggested by reviewers

Phenotype	Summary Statistics Description & Sample Size Summary Statistics Sc		
PTSD (Stein et al., 2021)	Continuous severity based on PCL total scores. N = 186689	https://www.ncbi.nlm.nih.g ov/projects/gap/cgi- bin/study.cgi?study_id=phs 001672.v1.p1.	
MDD (Howard et al., 2019)	Diagnosis based on clinician diagnosis, chart review, self-reported symptoms. N = 500199	https://datashare.ed.ac.uk/h andle/10283/3203	
Suicide attempt (Mullins et al., 2022)	Incidence based on structured clinical interviews, self-report questionnaires, and hospital records or ICD codes. N = 538436 (STARRS sample removed)	https://doi.org/10.1016/j.bi opsych.2021.05.029	
Schizophrenia (Ripke et al., 2020)	Diagnosis based on clinician and/or research assessment. N = 306011	https://figshare.com/article s/dataset/scz2021/1467217 8	
Alcohol Use Disorder (Walters et al., 2018)	Diagnosis based on clinician ratings or semi- structured interviews. N = 38686	https://figshare.com/article s/dataset/sud2018- alc/14672187	
Neuroticism (Nagel et al., 2018)	Continuous measure based on 12 items of the Eysenck Personality Questionnaire Revised Short Form. N = 390278	https://ctg.cncr.nl/software/ summary_statistics	

Supplementary Table 1. Data sources for polygenic risk scores

Supplementary Table 2. Correlations between scores calculated using the full and reduced versions of the PCL at the follow-ups

	6-item score and 17-item score	5-item score and 17-item score		
	R [95% CI], p-value	<i>R</i> [95% CI], <i>p-value</i>		
Follow-up 1	.96 [.96, .96], <i>p</i> < .001	.95 [.95, .96], <i>p</i> < .001		
Follow-up 2	.95 [.95, .96], <i>p</i> < .001	.95 [.95, .96], <i>p</i> < .001		

*Note.* The 6-item score reflects the sum of the six PCL items collected at pre-deployment, the 5item score reflects the sum of the five PCL items collected at post-deployment, and the 17-item score reflects the sum of the 17 PCL items (i.e., 17 DSM-IV PTSD symptoms). Supplementary Table 3. Guidelines for reporting on latent trajectory studies (checklist)

Checklist Item	Section Where Item is Reported
<b>1. Is the metric of time used in the statistical model reported?</b>	Method\Data
<i>Yes: "The post-deployment assessment timepoint was coded as 0, and each of the two subsequent follow-up assessments were coded as the number of months after the post-deployment assessment."</i>	analysis
<b>2. Is information presented about the mean and variance of time within a wave?</b>	Method\Data
<i>Yes: "The mean number of months between the first post-deployment assessment and each subsequent assessment were <math>2.38</math> (SD = <math>0.62</math>) for the first follow-up, and <math>11.76</math> (SD = <math>1.74</math>) for the final follow-up."</i>	analysis
<b>3a. Is the missing data mechanism reported?</b>	Method\Data
<i>Yes (missing at random): "Trajectory analyses were conducted in R (R Core Team, 2022) with the lcmm package (version 1.9. 4; Proust-Lima et al., 2017), which uses maximum likelihood estimation to include participants with data assumed to be missing at random."</i>	analysis
<b>3b.</b> Is a description provided of what variables are related to attrition/missing data?	Method\Data
Yes: "A non-monotonic missing data pattern was observed such that only a minority of participants (3.5%) completed the post-deployment assessment but did not return for any follow-ups. Pre-deployment posttraumatic stress symptom severity was significantly higher among participants with missing data at post-deployment (M difference = 3.50, $p < .001$ ) and at the first follow-up (M difference = 3.03, $p < .001$ ), but not the last follow-up (M difference = -0.48, $p = .37$ )."	analysis
<b>3c. Is a description provided of how missing data in the analyses were dealt with?</b>	Method\Data
<i>Yes (maximum likelihood estimation): "Trajectory analyses were conducted in R (R Core Team, 2022) with the lcmm package (version 1.9.4; Proust-Lima et al., 2017), which uses maximum likelihood estimation to include participants with data assumed to be missing at random."</i>	analysis
<b>4.</b> Is information about the distribution of the observed variables included?	Results\Sample
Yes: "Among participants included in the trajectory analyses, the average harmonized PCL score (range 0-100% of total) was $16.84\%$ (SD = $18.79$ , median = $10\%$ , $IQR = 25$ ; $n = 3897$ ) at post-deployment, $11.84\%$ (SD = $16.31$ , median = $5.88\%$ , $IQR = 16.18$ ; $n = 3566$ ) at the first follow-up, and $14.14\%$ (SD = $19.39$ , median = $4.41\%$ , $IQR = 22.06$ ; $n = 3351$ ) at the second follow-up."	Characteristics

<b>5. Is the software mentioned?</b> <i>Yes: "Trajectory analyses were conducted using the open source software R</i> <i>(R Core Team, 2022) with the lcmm package (version 1.9. 4; Proust-Lima et al., 2017"</i>	Method\Analysis
6a. Are alternative specifications of within-class heterogeneity considered (e.g., LGCA vs. LGMM) and clearly documented? If not, was sufficient justification provided as to eliminate certain specifications from consideration? Yes: "We examined modeling approaches that varied in their specifications of within-class heterogeneity: latent growth mixture models (LGMM; within-trajectory variation in the intercepts and/or slopes is allowed) and latent class growth analysis (LCGA; within-trajectory variance of the intercepts and slopes are fixed to zero)"	Method\Data analysis
<b>6b.</b> Are alternative specifications of the between-class differences in variance–covariance matrix structure considered and clearly documented? If not, was sufficient justification provided as to eliminate certain specifications from consideration? <i>Justification provided: "The variance-covariance matrix of the random effects was constrained to be equal across trajectories to facilitate model convergence."</i>	Method\Data analysis
<b>7. Are alternative shape/functional forms of the trajectories described?</b> <i>Yes: "We selected the post-deployment assessment as the starting point of our trajectory analyses (described in Data Analysis). This allowed us to model the course of posttraumatic stress symptom trajectories after exposure to potentially traumatic experiences during deployment. Additionally, length of deployment differed across the three BCTs, making the time from pre- to post-deployment assessments highly variable, which would have limited the interpretability of the trajectories."</i>	Method\Data analysis
<b>8. If covariates have been used, can analyses still be replicated?</b> <i>Yes: "Associations between each PRS and trajectory membership were examined in multinomial logistic regression analyses with the nnet package (version 7.3-14; Venables and Ripley, 2002)"</i>	Method\Data analysis
<b>9. Is information reported about the number of random start values and final iterations included?</b> Yes: "Random starting values for the multi-trajectory models used information from the single-trajectory model; values were generated using grid-search with 100 repetitions and a maximum of 100 iterations in the optimization algorithm. Analysis outputs with precise information about the number of random start values and final iterations included are provided in the Supplementary Material."	Method\Data analysis & Supplementary Material

<b>10.</b> Are the model comparison (and selection) tools described from a statistical perspective?	Method\Data
Yes: "We compared models with one to five trajectories and selected a final model based on sample size adjusted Bayesian information criteria (lower is better), entropy (reflects classification accuracy; values closer to 1 are better), and the Lo-Mendell-Rubin likelihood ratio test (significant value indicates better fit). Parsimonious models were favored over models that improved fit by adding an insufficiently distinct trajectory that included a low proportion of participants."	analysis
<b>11. Are the total number of fitted models reported, including a one-class solution?</b>	Supplementary
<i>Yes, Supplementary Table S2 shows the fit indices of the LGMM results with 1-5 trajectories.</i>	Table 2
<b>12.</b> Are the number of cases per class reported for each model (absolute sample size, or proportion)? <i>Yes, reported in Table 1.</i>	Table 1
<b>13. If classification of cases in a trajectory is the goal, is entropy reported?</b>	Supplementary
<i>Yes, Supplementary Table 2 reports entropy of the LGMM results with 1-5 trajectories.</i>	Table 2
<b>14a. Is a plot included with the estimated mean trajectories of the final solution?</b> <i>Yes: Figure 1.</i>	Figure 1
<b>14b. Are plots included with the estimated mean trajectories for each model?</b>	Supplementary
<i>Yes: Supplementary Figure 2.</i>	Figure 2
<b>14c. Is a plot included of the combination of estimated means of the final model and the observed individual trajectories split out for each latent class?</b>	Supplementary
<i>Yes: Supplementary Figure 1.</i>	Figure 1
<b>15. Are characteristics of the final class solution numerically described</b> (i.e., means, SD/SE, n, CI, etc.)? <i>Reported in Table 1.</i>	Table 1
<b>16.</b> Are the syntax files available (either in the appendix, supplementary materials, or from the authors)? <i>Provided in Appendix 1-3 of the Supplementary Materials.</i>	Supplementary Materials

						% in	
	Log					Smallest	LMR-LRT
Model	Likelihood	AIC	BIC	SSA-BIC	Entropy	Trajectory	(p-value)
LGMM							
1 trajectory	-45504.87	91021.75	91060.02	91040.96	1	100	NA
2 trajectories	-44550.01	89120.02	89183.81	89152.03	0.90	13.50	1836.7 (<.001)
3 trajectories	-44243.42	88514.84	88604.15	88559.66	0.85	6.64	589.7 (<.001)
4 trajectories	-43892.01	87820.01	87934.83	87877.64	0.86	4.32	675.9 (<.001)
5 trajectories	-43756.79	87557.58	87697.91	87628.01	0.85	2.66	260.1 (<.001)
LGMM with							
fixed slopes							
1 trajectory	-45585.63	91179.26	91204.78	91192.07	1	100	NA
2 trajectories	-44640.54	89297.08	89348.11	89322.69	0.90	14.10	1817.9 (<.001)
3 trajectories	-44193.72	88411.44	88487.99	88449.86	0.89	7.56	859.4 (<.001)
4 trajectories	-43897.56	87827.13	87929.19	87878.35	0.86	4.25	569.7 (<.001)
5 trajectories	-43772.34	87584.68	87712.25	87648.7	0.85	2.07	240.9 (<.001)
LCGA							
1 trajectory	-46794.13	93594.25	93613.39	93603.85	1	100	NA
2 trajectories	-44745.41	89504.83	89549.48	89527.24	0.91	16.72	3940.7 (<.001)
3 trajectories	-44321.82	88665.64	88735.8	88700.85	0.87	6.01	814.8 (<.001)
4 trajectories	-43929.83	87889.65	87985.34	87937.67	0.86	5.26	754.0 (<.001)
5 trajectories	-43772.34	87582.68	87703.87	87643.5	0.85	2.07	302.9 (<.001)

Supplementary Table 4. Model fit indices of the different models.

*Note*. SSA-BIC = Sample Size Adjusted Bayesian Information Criteria; LMR-LRT = Lo-

Mendell-Rubin Likelihood Ratio Test for all of the estimated models.



Supplementary Figure 1. PTSD trajectories from each of the estimated models



Supplementary Figure 2. Individual observed trajectories for each LGMM trajectory.

Appendix 1. Analysis code and output of the LGMM models (random intercepts and slopes)

```
#Svntax
library(lcmm)
#1 trajectory
lgmm1 <- hlme(pcl ~ months, subject = "id", ng = 1, data = df,</pre>
verbose = F, returndata = TRUE, random = \sim 1 + months)
summary(lqmm1)
saveRDS(lgmm1, "lgmm1.prs.rds")
#2 trajectories
lgmm2 <- gridsearch(rep = 100, maxiter = 100, minit = lgmm1, cl = 16,
                     hlme(pcl ~ months, subject = "id", ng = 2, data =
df, mixture = ~1 + months, verbose = F, returndata = TRUE, classmb =
\simpcl.bl, random = \sim1 + months))
summary(lqmm2)
saveRDS(lgmm2, "lgmm2.prs.rds")
#3 trajectories
lgmm3 <- gridsearch(rep = 100, maxiter = 100, minit = lgmm1, cl = 16,</pre>
                     hlme(pcl ~ months, subject = "id", ng = 3, data =
df, mixture = ~1 + months, verbose = F, returndata = TRUE, classmb =
\simpcl.bl, random = \sim1 + months))
summary(lqmm3)
saveRDS(lgmm3, "lgmm3.prs.rds")
#4 trajectories
lgmm4 <- gridsearch(rep = 100, maxiter = 100, minit = lgmm1, cl = 16,</pre>
                     hlme(pcl ~ months, subject = "id", ng = 4, data =
df, mixture = ~1 + months, verbose = F, returndata = TRUE, classmb =
~pcl.bl, random = ~1 + months))
summary(lgmm4)
saveRDS(lgmm4, "lgmm4.prs.rds")
#5 trajectories
lgmm5 <- gridsearch(rep = 100, maxiter = 100, minit = lgmm1, cl = 16,
hlme(pcl ~ months, subject = "id", ng = 5, data =
df, mixture = ~1 + months, verbose = F, returndata = TRUE, classmb =
\simpcl.bl, random = \sim1 + months))
summary(lgmm5)
saveRDS(lgmm5, "lgmm5.prs.rds")
sessionInfo()
#Output
Heterogenous linear mixed model
     fitted by maximum likelihood method
hlme(fixed = pcl ~ months, random = ~1 + months, subject = "id",
    ng = 1, data = df, verbose = F, returndata = TRUE)
Statistical Model:
     Dataset: df
     Number of subjects: 4355
     Number of observations: 10815
     Number of observations deleted: 3885
```

Number of latent classes: 1 Number of parameters: 6 Iteration process: Convergence criteria satisfied Number of iterations: 15 Convergence criteria: parameters= 2.8e-11 : likelihood= 1.5e-09 : second derivatives= 7.1e-14 Goodness-of-fit statistics: maximum log-likelihood: -45504.87 AIC: 91021.75 BIC: 91060.02 Maximum Likelihood Estimates: Fixed effects in the longitudinal model: coef Se Wald p-value intercept 15.30635 0.26667 57.397 0.00000 -0.15123 0.02739 -5.521 0.00000 months Variance-covariance matrix of the random-effects: intercept months intercept 208.44789 -3.12414 0.83152 months coef Se Residual standard error: 11.13530 0.13596 Be patient, grid search is running ... Search completed, performing final estimation Heterogenous linear mixed model fitted by maximum likelihood method hlme(fixed = pcl ~ months, mixture =  $\sim 1$  + months, random =  $\sim 1$  + months, subject = "id", classmb = ~pcl.bl, ng = 2, data = df, verbose = F, returndata = TRUE) Statistical Model: Dataset: df Number of subjects: 4354 Number of observations: 10814 Number of observations deleted: 3886 Number of latent classes: 2 Number of parameters: 10 Iteration process:

Convergence criteria satisfied Number of iterations: 1 Convergence criteria: parameters= 2.4e-13 : likelihood= 4.4e-11 : second derivatives= 3.5e-13 Goodness-of-fit statistics: maximum log-likelihood: -44550.01 AIC: 89120.02 BIC: 89183.81 Maximum Likelihood Estimates: Fixed effects in the class-membership model: (the class of reference is the last class) coef Se Wald p-value 2.43690 0.07077 34.435 0.00000 intercept class1 pcl.bl class1 -0.05377 0.00301 -17.862 0.00000 Fixed effects in the longitudinal model: coef Se Wald p-value intercept class1 11.16041 0.25299 44.115 0.00000 intercept class2 41.07676 0.88145 46.601 0.00000 months class1 -0.31136 0.02855 -10.905 0.00000 months class2 0.84421 0.09490 8.896 0.00000 Variance-covariance matrix of the random-effects: intercept months intercept 103.38601 -6.58259 0.5267 months coef Se Residual standard error: 11.09340 0.13379 Be patient, grid search is running ... Search completed, performing final estimation Heterogenous linear mixed model fitted by maximum likelihood method hlme(fixed = pcl ~ months, mixture =  $\sim 1$  + months, random =  $\sim 1$  + months, subject = "id", classmb = ~pcl.bl, ng = 3, data = df, verbose = F, returndata = TRUE) Statistical Model: Dataset: df Number of subjects: 4354 Number of observations: 10814 Number of observations deleted: 3886 Number of latent classes: 3

Number of parameters: 14 Iteration process: Convergence criteria satisfied Number of iterations: 7 Convergence criteria: parameters= 4.2e-13 : likelihood= 3.3e-10 : second derivatives= 3.6e-10 Goodness-of-fit statistics: maximum log-likelihood: -44243.42 AIC: 88514.84 BIC: 88604.15 Maximum Likelihood Estimates: Fixed effects in the class-membership model: (the class of reference is the last class) Wald p-value coef Se intercept class1 -1.27708 0.12128 -10.530 0.00000 intercept class2 1.96632 0.06885 28.560 0.00000 0.01612 0.00334 pcl.bl class1 4.827 0.00000 pcl.bl class2 -0.05725 0.00385 -14.884 0.00000 Fixed effects in the longitudinal model: coef Se Wald p-value intercept class1 51.09980 1.23129 41.501 0.00000 intercept class2 9.20397 0.26353 34.926 0.00000 intercept class3 28.40806 0.78870 36.019 0.00000 months class1 1.03706 0.13486 7.690 0.00000 months class2 -0.34982 0.03186 -10.980 0.00000 months class3 0.23262 0.08764 2.654 0.00795 Variance-covariance matrix of the random-effects: intercept months intercept 75.99035 months -7.51830 0.74384 coef Se Residual standard error: 10.52933 0.12642 Be patient, grid search is running ... Search completed, performing final estimation Heterogenous linear mixed model fitted by maximum likelihood method hlme(fixed = pcl ~ months, mixture =  $\sim 1$  + months, random =  $\sim 1$  + months, subject = "id", classmb = ~pcl.bl, ng = 4, data = df, verbose = F, returndata = TRUE)

Statistical Model: Dataset: df Number of subjects: 4354 Number of observations: 10814 Number of observations deleted: 3886 Number of latent classes: 4 Number of parameters: 18 Iteration process: Convergence criteria satisfied Number of iterations: 1 Convergence criteria: parameters= 5.1e-10 : likelihood= 2.9e-09 : second derivatives= 4.1e-10 Goodness-of-fit statistics: maximum log-likelihood: -43892.01 AIC: 87820.01 BIC: 87934.83 Maximum Likelihood Estimates: Fixed effects in the class-membership model: (the class of reference is the last class) Wald p-value coef Se intercept class1 1.43755 0.15816 9.089 0.00000 intercept class2 3.80563 0.14248 26.711 0.00000 intercept class3 1.10883 0.16564 6.694 0.00000 pcl.bl class1 -0.02297 0.00411 -5.586 0.00000 pcl.bl class2 -0.07965 0.00454 -17.548 0.00000 pcl.bl class3 -0.01937 0.00430 -4.505 0.00001 Fixed effects in the longitudinal model: coef Se Wald p-value intercept class1 20.80435 0.83530 24.906 0.00000 intercept class2 8.56505 0.22412 38.216 0.00000 intercept class3 43.76547 1.22642 35.685 0.00000 intercept class4 61.86607 1.29297 47.848 0.00000 months class1 1.96858 0.09538 20.640 0.00000 months class2 -0.23333 0.02769 -8.427 0.00000 months class3 -2.40431 0.11943 -20.132 0.00000 months class4 0.06508 0.14336 0.454 0.64987 Variance-covariance matrix of the random-effects: intercept months intercept 22.84574 months -0.77236 0.02611

```
coef Se
```

Residual standard error: 10.55853 0.10292 Be patient, grid search is running ... Search completed, performing final estimation Heterogenous linear mixed model fitted by maximum likelihood method hlme(fixed = pcl ~ months, mixture =  $\sim 1$  + months, random =  $\sim 1$  + months, subject = "id", classmb = ~pcl.bl, ng = 5, data = df, verbose = F, returndata = TRUE) Statistical Model: Dataset: df Number of subjects: 4354 Number of observations: 10814 Number of observations deleted: 3886 Number of latent classes: 5 Number of parameters: 22 Iteration process: Convergence criteria satisfied Number of iterations: 4 Convergence criteria: parameters= 3.4e-10 : likelihood= 1.3e-06 : second derivatives= 5e-07 Goodness-of-fit statistics: maximum log-likelihood: -43756.79 AIC: 87557.58 BIC: 87697.91 Maximum Likelihood Estimates: Fixed effects in the class-membership model: (the class of reference is the last class) Wald p-value coef Se intercept class1 -3.65972 0.19682 -18.594 0.00000 intercept class2 -2.78195 0.11091 -25.084 0.00000 intercept class3 -2.19165 0.08185 -26.776 0.00000 intercept class4 -3.83536 0.15103 -25.395 0.00000 pcl.bl class1 0.06506 0.00615 10.588 0.00000 pcl.bl class2 pcl.bl class3 0.06279 0.00477 13.161 0.00000 0.05881 0.00460 12.774 0.00000 pcl.bl class4 0.08518 0.00504 16.895 0.00000 Fixed effects in the longitudinal model: coef Se Wald p-value intercept class1 25.87088 2.03726 12.699 0.00000 intercept class2 44.31612 1.21380 36.510 0.00000

intercept class3 19.41817 0.83901 23.144 0.00000 intercept class4 63.39375 1.41826 44.698 0.00000 intercept class5 8.23760 0.22381 36.807 0.00000 months class13.580530.2950512.1350.00000months class2-2.671920.11874-22.5030.00000months class31.238780.0902313.7290.00000months class4-0.603820.17385-3.4730.00051months class5-0.283940.02658-10.6810.00000 Variance-covariance matrix of the random-effects: intercept months intercept 26.62199 months -1.55051 0.0903 coef Se Residual standard error: 10.05975 0.10128 R version 4.1.0 (2021-05-18) Platform: x86 64-pc-linux-gnu (64-bit) Running under: CentOS Linux 7 (Core) Matrix products: default BLAS: /sw/arcts/centos7/stacks/gcc/8.2.0/R/4.1.0/lib64/R/lib/libRbla s.so LAPACK: /sw/arcts/centos7/stacks/gcc/8.2.0/R/4.1.0/lib64/R/lib/libRlapack.so Random number generation: RNG: L'Ecuyer-CMRG Normal: Inversion Sample: Rejection locale: [1] LC\_CTYPE=en\_US.UTF-8LC\_NUMERIC=C[3] LC\_TIME=en\_US.UTF-8LC\_COLLATE=en\_US.UTF-8[5] LC\_MONETARY=en\_US.UTF-8LC\_MESSAGES=en\_US.UTF-8[7] LC\_PAPER=en\_US.UTF-8LC\_NAME=C LC TELEPHONE=C [9] LC ADDRESS=C [11] LC MEASUREMENT=en US.UTF-8 LC IDENTIFICATION=C attached base packages: [1] parallel stats graphics grDevices utils datasets methods [8] base other attached packages: [1] lcmm 1.9.3 mvtnorm 1.1-3 survival 3.2-11 loaded via a namespace (and not attached): [1] compiler 4.1.0 Matrix 1.3-3 splines 4.1.0 grid 4.1.0 [5] lattice 0.20-44

Appendix 2. Analysis code and output of the LGMM models (random intercepts, fixed slopes) #Syntax library(lcmm) #1 trajectory lqmmfs1 <- hlme(pcl ~ months, subject = "id", nq = 1, data = df, verbose = F, returndata = TRUE, random =  $\sim$ 1) summary(lgmmfs1) saveRDS(lgmmfs1, "lgmmfs1.prs.rds") #2 trajectories lgmmfs2 <- gridsearch(rep = 100, maxiter = 100, minit = lgmmfs1, cl = 16, hlme(pcl ~ months, subject = "id", ng = 2, data = df, mixture = ~1 + months, verbose = F, returndata = TRUE, classmb = ~pcl.bl, random = ~1)) summary(lgmmfs2) saveRDS(lgmmfs2, "lgmmfs2.prs.rds") #3 trajectories lgmmfs3 <- gridsearch(rep = 100, maxiter = 100, minit = lgmmfs1, cl = 16, hlme(pcl ~ months, subject = "id", ng = 3, data = df, mixture = ~1 + months, verbose = F, returndata = TRUE, classmb = ~pcl.bl, random = ~1)) summary(lgmmfs3) saveRDS(lgmmfs3, "lgmmfs3.prs.rds") #4 trajectories lgmmfs4 <- gridsearch(rep = 100, maxiter = 100, minit = lgmmfs1, cl =</pre> 16, hlme(pcl ~ months, subject = "id", ng = 4, data = df, mixture = ~1 + months, verbose = F, returndata = TRUE, classmb = ~pcl.bl, random = ~1)) summary(lgmmfs4) saveRDS(lgmmfs4, "lgmmfs4.prs.rds") #5 trajectories lgmmfs5 <- gridsearch(rep = 100, maxiter = 100, minit = lgmmfs1, cl = 16, hlme(pcl ~ months, subject = "id", ng = 5, data = df, mixture = ~1 + months, verbose = F, returndata = TRUE, classmb = ~pcl.bl, random = ~1)) summary(lgmmfs5) saveRDS(lgmmfs5, "lgmmfs5.prs.rds") sessionInfo() #Output Heterogenous linear mixed model fitted by maximum likelihood method hlme(fixed = pcl ~ months, random = ~1, subject = "id", ng = 1, data = df, verbose = F, returndata = TRUE) Statistical Model: Dataset: df

Number of subjects: 4355 Number of observations: 10815 Number of observations deleted: 3885 Number of latent classes: 1 Number of parameters: 4 Iteration process: Convergence criteria satisfied Number of iterations: 15 Convergence criteria: parameters= 1.2e-06 : likelihood= 2.9e-05 : second derivatives= 9.3e-13 Goodness-of-fit statistics: maximum log-likelihood: -45585.63 AIC: 91179.26 BIC: 91204.78 Maximum Likelihood Estimates: Fixed effects in the longitudinal model: coef Se Wald p-value intercept 15.28150 0.26600 57.450 0.00000 months -0.13672 0.02460 -5.557 0.00000 Variance-covariance matrix of the random-effects: intercept intercept 188.6323 coef Se Residual standard error: 12.44031 0.11078 Be patient, grid search is running ... Search completed, performing final estimation Heterogenous linear mixed model fitted by maximum likelihood method hlme(fixed = pcl ~ months, mixture =  $\sim 1 + months$ , random =  $\sim 1$ , subject = "id", classmb = ~pcl.bl, ng = 2, data = df, verbose = F, returndata = TRUE) Statistical Model: Dataset: df Number of subjects: 4354 Number of observations: 10814 Number of observations deleted: 3886 Number of latent classes: 2 Number of parameters: 8 Iteration process:

Convergence criteria satisfied Number of iterations: 1 Convergence criteria: parameters= 1.5e-12 : likelihood= 1.4e-10 : second derivatives= 2.8e-12 Goodness-of-fit statistics: maximum log-likelihood: -44640.54 AIC: 89297.08 BIC: 89348.11 Maximum Likelihood Estimates: Fixed effects in the class-membership model: (the class of reference is the last class) coef Se Wald p-value intercept class1 2.43165 0.07182 33.858 0.00000 pcl.bl class1 -0.05644 0.00311 -18.145 0.00000 Fixed effects in the longitudinal model: coef Se Wald p-value intercept class1 10.34588 0.21786 47.490 0.00000 intercept class2 45.01607 0.72126 62.413 0.00000 months class1 -0.19437 0.02717 -7.155 0.00000 1.890 0.05878 months class2 0.15521 0.08213 Variance-covariance matrix of the random-effects: intercept intercept 31.97744 coef Se Residual standard error: 12.31836 0.10784 Be patient, grid search is running ... Search completed, performing final estimation Heterogenous linear mixed model fitted by maximum likelihood method  $hlme(fixed = pcl \sim months, mixture = ~1 + months, random = ~1,$ subject = "id", classmb = ~pcl.bl, ng = 3, data = df, verbose = F, returndata = TRUE)Statistical Model: Dataset: df Number of subjects: 4354 Number of observations: 10814 Number of observations deleted: 3886 Number of latent classes: 3

Number of parameters: 12 Iteration process: Convergence criteria satisfied Number of iterations: 1 Convergence criteria: parameters= 2e-11 : likelihood= 5.8e-10 : second derivatives= 3.4e-11 Goodness-of-fit statistics: maximum log-likelihood: -44193.72 AIC: 88411.44 BIC: 88487.99 Maximum Likelihood Estimates: Fixed effects in the class-membership model: (the class of reference is the last class) Wald p-value coef Se intercept class1 0.34469 0.13350 2.582 0.00982 intercept class2 3.01771 0.09879 30.546 0.00000 -0.00665 0.00352 -1.891 0.05862 pcl.bl class1 pcl.bl class2 -0.06051 0.00362 -16.732 0.00000 Fixed effects in the longitudinal model: coef Se Wald p-value intercept class1 26.92831 1.08929 24.721 0.00000 intercept class2 9.91954 0.20771 47.757 0.00000 intercept class3 57.27827 0.97959 58.472 0.00000 months class1 2.26532 0.10887 20.808 0.00000 months class2 -0.24000 0.02528 -9.495 0.00000 months class3 -2.33187 0.13207 -17.657 0.00000 Variance-covariance matrix of the random-effects: intercept intercept 39.97696 coef Se Residual standard error: 10.85302 0.10439 Be patient, grid search is running ... Search completed, performing final estimation Heterogenous linear mixed model fitted by maximum likelihood method hlme(fixed = pcl ~ months, mixture =  $\sim 1 + months$ , random =  $\sim 1$ , subject = "id", classmb = ~pcl.bl, ng = 4, data = df, verbose = F, returndata = TRUE)

Statistical Model: Dataset: df Number of subjects: 4354 Number of observations: 10814 Number of observations deleted: 3886 Number of latent classes: 4 Number of parameters: 16 Iteration process: Convergence criteria satisfied Number of iterations: 1 Convergence criteria: parameters= 7.7e-12 : likelihood= 1e-10 : second derivatives= 7.8e-12 Goodness-of-fit statistics: maximum log-likelihood: -43897.56 AIC: 87827.13 BIC: 87929.19 Maximum Likelihood Estimates: Fixed effects in the class-membership model: (the class of reference is the last class) Wald p-value coef Se intercept class1 -2.59871 0.09657 -26.911 0.00000 intercept class2 -3.80726 0.13953 -27.286 0.00000 intercept class3 -2.42347 0.08550 -28.344 0.00000 0.05874 0.00429 13.679 0.00000 pcl.bl class1 0.08024 0.00453 17.715 0.00000 pcl.bl class2 pcl.bl class3 0.05716 0.00425 13.452 0.00000 Fixed effects in the longitudinal model: coef Se Wald p-value intercept class1 42.51662 1.08083 39.337 0.00000 intercept class2 62.47446 1.21065 51.604 0.00000 intercept class3 21.11928 0.84371 25.031 0.00000 intercept class4 8.38637 0.21113 39.720 0.00000 months class1 -2.35248 0.11440 -20.564 0.00000 months class2 -0.09866 0.13978 -0.706 0.48028 2.02309 0.09769 20.708 0.00000 months class3 months class4 -0.20315 0.02639 -7.698 0.00000 Variance-covariance matrix of the random-effects: intercept intercept 15.15611 coef Se Residual standard error: 10.62151 0.10316

```
Be patient, grid search is running ...
Search completed, performing final estimation
Heterogenous linear mixed model
     fitted by maximum likelihood method
hlme(fixed = pcl ~ months, mixture = \sim 1 + months, random = \sim 1,
    subject = "id", classmb = ~pcl.bl, ng = 5, data = df, verbose = F,
    returndata = TRUE)
Statistical Model:
     Dataset: df
     Number of subjects: 4354
     Number of observations: 10814
     Number of observations deleted: 3886
     Number of latent classes: 5
     Number of parameters: 20
Iteration process:
     Convergence criteria satisfied
     Number of iterations: 4
     Convergence criteria: parameters= 4.2e-13
                         : likelihood= 4.4e-11
                          : second derivatives= 2.6e-11
Goodness-of-fit statistics:
     maximum log-likelihood: -43772.34
     AIC: 87584.68
     BIC: 87712.25
Maximum Likelihood Estimates:
Fixed effects in the class-membership model:
(the class of reference is the last class)
                     coef
                               Se
                                    Wald p-value
intercept class1 -2.49434 0.21686 -11.502 0.00000
intercept class2 -1.09668 0.13985 -7.842 0.00000
intercept class3 2.13552 0.09050 23.597 0.00000
intercept class4 -0.38875 0.13751 -2.827 0.00470
pcl.bl class1
                 0.03217 0.00505 6.373 0.00000
pcl.bl class20.019190.004224.5520.00001pcl.bl class3-0.056330.00446-12.6280.00000
pcl.bl class4
                 0.00396 0.00449 0.883 0.37722
Fixed effects in the longitudinal model:
                     coef
                              Se
                                    Wald p-value
intercept class1 71.07610 2.06095 34.487 0.00000
intercept class2 54.62429 1.07994 50.581 0.00000
intercept class3 7.18775 0.19994 35.950 0.00000
```

intercept class4 20.12952 1.12214 17.938 0.00000 intercept class5 30.56043 1.09067 28.020 0.00000 months class1 0.29383 0.18353 1.601 0.10938 monthsclass10.293030.103331.0010.10930monthsclass2-1.686160.25338-6.6550.00000monthsclass3-0.139990.02705-5.1750.00000monthsclass42.328230.1145020.3330.00000monthsclass5-1.461500.12659-11.5450.00000 Variance-covariance matrix of the random-effects: intercept intercept 0 coef Se Residual standard error: 10.67058 0.08531 R version 4.1.0 (2021-05-18) Platform: x86 64-pc-linux-qnu (64-bit) Running under: CentOS Linux 7 (Core) Matrix products: default BLAS: /sw/arcts/centos7/stacks/gcc/8.2.0/R/4.1.0/lib64/R/lib/libRbla s.so LAPACK: /sw/arcts/centos7/stacks/gcc/8.2.0/R/4.1.0/lib64/R/lib/libRlapack.so Random number generation: RNG: L'Ecuver-CMRG Normal: Inversion Sample: Rejection locale: [1] LC\_CTYPE=en\_US.UTF-8LC\_NUMERIC=C[3] LC\_TIME=en\_US.UTF-8LC\_COLLATE=en\_US.UTF-8[5] LC\_MONETARY=en\_US.UTF-8LC\_MESSAGES=en\_US.UTF-8[7] LC\_PAPER=en\_US.UTF-8LC\_NAME=C [9] LC ADDRESS=C LC TELEPHONE=C [11] LC MEASUREMENT=en US.UTF-8 LC IDENTIFICATION=C attached base packages: [1] parallel stats graphics grDevices datasets methods utils [8] base other attached packages: [1] lcmm 1.9.3 mvtnorm 1.1-3 survival 3.2-11 loaded via a namespace (and not attached): [1] compiler 4.1.0 Matrix 1.3-3 splines 4.1.0 grid 4.1.0 [5] lattice 0.20-44

Appendix 3. Analysis code and output of the LCGA models (fixed intercepts and slopes)

```
#Svntax
library(lcmm)
#1 trajectory
lcga1 <- hlme(pcl ~ months, subject = "id", ng = 1, data = df,</pre>
verbose = F, returndata = TRUE)
summary(lcga1)
saveRDS(lcga1, "lcga1.prs.rds")
#2 trajectories
lcga2 <- gridsearch(rep = 100, maxiter = 100, minit = lcga1, cl = 16,</pre>
hlme(pcl ~ months, subject = "id", ng = 2, data = df, mixture = ~1 +
months, verbose = F, returndata = TRUE, classmb = ~pcl.bl))
summary(lcga2)
saveRDS(lcga2, "lcga2.prs.rds")
#3 trajectories
lcga3 <- gridsearch(rep = 100, maxiter = 100, minit = lcga1, cl = 16,</pre>
hlme(pcl ~ months, subject = "id", ng = 3, data = df, mixture = ~1 +
months, verbose = F, returndata = TRUE, classmb = ~pcl.bl))
summary(lcga3)
saveRDS(lcga3, "lcga3.prs.rds")
#4 trajectories
lcga4 <- gridsearch(rep = 100, maxiter = 100, minit = lcga1, cl = 16,</pre>
hlme(pcl ~ months, subject = "id", ng = 4, data = df, mixture = \sim 1 + 1
months, verbose = F, returndata = TRUE, classmb = ~pcl.bl))
summary(lcga4)
saveRDS(lcga4, "lcga4.prs.rds")
#5 trajectories
lcga5 <- gridsearch(rep = 100, maxiter = 100, minit = lcga1, cl = 16,</pre>
hlme(pcl ~ months, subject = "id", ng = 5, data = df, mixture = \sim 1 + 1
months, verbose = F, returndata = TRUE, classmb = ~pcl.bl))
summary(lcga5)
saveRDS(lcga5, "lcga5.prs.rds")
sessionInfo()
#Output
Heterogenous linear mixed model
     fitted by maximum likelihood method
hlme(fixed = pcl ~ months, subject = "id", ng = 1, data = df,
    verbose = F, returndata = TRUE)
Statistical Model:
     Dataset: df
     Number of subjects: 4355
     Number of observations: 10815
     Number of observations deleted: 3885
```

Number of latent classes: 1 Number of parameters: 3 Iteration process: Convergence criteria satisfied Number of iterations: 15 Convergence criteria: parameters= 1.5e-09 : likelihood= 4.5e-08 : second derivatives= 9.6e-14 Goodness-of-fit statistics: maximum log-likelihood: -46794.13 AIC: 93594.25 BIC: 93613.39 Maximum Likelihood Estimates: Fixed effects in the longitudinal model: Se Wald p-value coef intercept 14.71941 0.23248 63.315 0.00000 months -0.08219 0.03425 -2.400 0.01640 coef Se Residual standard error: 18.31722 0.12458 Be patient, grid search is running ... Search completed, performing final estimation Heterogenous linear mixed model fitted by maximum likelihood method  $hlme(fixed = pcl \sim months, mixture = \sim 1 + months, subject = "id",$ classmb = ~pcl.bl, ng = 2, data = df, verbose = F, returndata = TRUE) Statistical Model: Dataset: df Number of subjects: 4354 Number of observations: 10814 Number of observations deleted: 3886 Number of latent classes: 2 Number of parameters: 7 Iteration process: Convergence criteria satisfied Number of iterations: 1 Convergence criteria: parameters= 2.3e-11 : likelihood= 2e-10 : second derivatives= 8.5e-11 Goodness-of-fit statistics:

maximum log-likelihood: -44745.41 AIC: 89504.83 BIC: 89549.48 Maximum Likelihood Estimates: Fixed effects in the class-membership model: (the class of reference is the last class) Wald p-value coef Se 2.28128 0.06422 35.521 0.00000 intercept class1 -0.05555 0.00299 -18.557 0.00000 pcl.bl class1 Fixed effects in the longitudinal model: Se coef Wald p-value intercept class1 9.90164 0.19867 49.840 0.00000 intercept class2 43.59696 0.58927 73.985 0.00000 months class1 -0.18620 0.02841 -6.554 0.00000 months class2 0.10830 0.07249 1.494 0.13520 coef Se Residual standard error: 13.34773 0.09549 Be patient, grid search is running ... Search completed, performing final estimation Heterogenous linear mixed model fitted by maximum likelihood method hlme(fixed = pcl ~ months, mixture = ~1 + months, subject = "id", classmb = ~pcl.bl, ng = 3, data = df, verbose = F, returndata = TRUE) Statistical Model: Dataset: df Number of subjects: 4354 Number of observations: 10814 Number of observations deleted: 3886 Number of latent classes: 3 Number of parameters: 11 Iteration process: Convergence criteria satisfied Number of iterations: 1 Convergence criteria: parameters= 2.2e-11 : likelihood= 2.7e-10 : second derivatives= 3.1e-11 Goodness-of-fit statistics: maximum log-likelihood: -44321.82 AIC: 88665.64

BIC: 88735.8 Maximum Likelihood Estimates: Fixed effects in the class-membership model: (the class of reference is the last class) Wald p-value coef Se intercept class1 -1.95237 0.06682 -29.218 0.00000 intercept class2 -3.47052 0.11942 -29.063 0.00000 0.05582 0.00365 15.280 0.00000 pcl.bl class1 0.07658 0.00410 18.673 0.00000 pcl.bl class2 Fixed effects in the longitudinal model: Wald p-value coef Se intercept class1 29.61326 0.72118 41.062 0.00000 intercept class2 60.91559 1.08631 56.076 0.00000 intercept class3 8.54712 0.21041 40.620 0.00000 months class1 0.29339 0.09744 3.011 0.00260 months class2 -0.45301 0.16443 -2.755 0.00587 months class3 -0.22076 0.02831 -7.797 0.00000 coef Se Residual standard error: 12.13090 0.09173 Be patient, grid search is running ... Search completed, performing final estimation Heterogenous linear mixed model fitted by maximum likelihood method hlme(fixed = pcl ~ months, mixture = ~1 + months, subject = "id", classmb = ~pcl.bl, ng = 4, data = df, verbose = F, returndata = TRUE) Statistical Model: Dataset: df Number of subjects: 4354 Number of observations: 10814 Number of observations deleted: 3886 Number of latent classes: 4 Number of parameters: 15 Iteration process: Convergence criteria satisfied Number of iterations: 1 Convergence criteria: parameters= 2.9e-12 : likelihood= 1.2e-10 : second derivatives= 1.6e-12

Goodness-of-fit statistics:

maximum log-likelihood: -43929.83 AIC: 87889.65 BIC: 87985.34 Maximum Likelihood Estimates: Fixed effects in the class-membership model: (the class of reference is the last class) coef Se Wald p-value intercept class1 -3.60411 0.12491 -28.854 0.00000 intercept class2 -2.38148 0.08906 -26.739 0.00000 intercept class3 -2.43869 0.09612 -25.372 0.00000 pcl.bl class1 0.08074 0.00433 18.636 0.00000 0.05680 0.00427 13.316 0.00000 pcl.bl class2 pcl.bl class3 0.05699 0.00425 13.420 0.00000 Fixed effects in the longitudinal model: Wald p-value coef Se intercept class1 60.81923 0.99746 60.974 0.00000 intercept class2 20.78174 0.82317 25.246 0.00000 intercept class3 39.49918 1.06474 37.097 0.00000 intercept class4 7.91413 0.19767 40.036 0.00000 months class1 -0.18134 0.13544 -1.339 0.18059 months class21.944980.1108317.5490.00000months class3-2.127810.12321-17.2700.00000months class4-0.183650.02736-6.7130.00000 coef Se Residual standard error: 11.13166 0.08784 Be patient, grid search is running ... Search completed, performing final estimation Heterogenous linear mixed model fitted by maximum likelihood method  $hlme(fixed = pcl \sim months, mixture = ~1 + months, subject = "id",$ classmb = ~pcl.bl, ng = 5, data = df, verbose = F, returndata = TRUE) Statistical Model: Dataset: df Number of subjects: 4354 Number of observations: 10814 Number of observations deleted: 3886 Number of latent classes: 5 Number of parameters: 19 Iteration process: Convergence criteria satisfied

Number of iterations: 1 Convergence criteria: parameters= 3.8e-11 : likelihood= 3.6e-10 : second derivatives= 2.7e-11 Goodness-of-fit statistics: maximum log-likelihood: -43772.34 AIC: 87582.68 BIC: 87703.87 Maximum Likelihood Estimates: Fixed effects in the class-membership model: (the class of reference is the last class) Se Wald p-value coef intercept class1 -2.52427 0.09449 -26.715 0.00000 intercept class2 -2.13552 0.09033 -23.642 0.00000 intercept class3 -3.23220 0.12013 -26.907 0.00000 intercept class4 -4.62986 0.20519 -22.564 0.00000 pcl.bl class1 0.06030 0.00463 13.035 0.00000 0.05633 0.00446 12.636 0.00000 pcl.bl class2 0.07552 0.00451 16.745 0.00000 pcl.bl class3 pcl.bl class4 0.08851 0.00558 15.858 0.00000 Fixed effects in the longitudinal model: Wald p-value coef Se intercept class1 20.12951 1.12198 17.941 0.00000 intercept class2 30.56043 1.09001 28.037 0.00000 intercept class3 54.62429 1.08010 50.574 0.00000 intercept class4 71.07610 2.06283 34.456 0.00000 intercept class5 7.18775 0.19983 35.969 0.00000 2.32823 0.11443 20.346 0.00000 months class1 months class2 -1.46150 0.12645 -11.558 0.00000 months class3 -1.68616 0.25342 -6.654 0.00000 months class4 0.29383 0.18402 1.597 0.11032 -0.13999 0.02702 -5.181 0.00000 months class5 coef Se Residual standard error: 10.67058 0.08531 R version 4.1.0 (2021-05-18) Platform: x86 64-pc-linux-gnu (64-bit) Running under: CentOS Linux 7 (Core) Matrix products: default BLAS: /sw/arcts/centos7/stacks/gcc/8.2.0/R/4.1.0/lib64/R/lib/libRbla s.so

LAPACK: /sw/arcts/centos7/stacks/gcc/8.2.0/R/4.1.0/lib64/R/lib/libRlapack.so

```
Random number generation:
RNG: L'Ecuyer-CMRG
 Normal: Inversion
Sample: Rejection
locale:
[1] LC_CTYPE=en_US.UTF-8LC_NUMERIC=C[3] LC_TIME=en_US.UTF-8LC_COLLATE=en_US.UTF-8[5] LC_MONETARY=en_US.UTF-8LC_MESSAGES=en_US.UTF-8[7] LC_PAPER=en_US.UTF-8LC_NAME=C[9] LC_ADDRESS=CLC_TELEPHONE=C
 [9] LC ADDRESS=C
                                     LC TELEPHONE=C
[11] LC MEASUREMENT=en US.UTF-8 LC IDENTIFICATION=C
attached base packages:
[1] parallel stats graphics grDevices
utils datasets methods
[8] base
other attached packages:
[1] lcmm 1.9.3 mvtnorm 1.1-3 survival 3.2-11
loaded via a namespace (and not attached):
[1] compiler 4.1.0 Matrix 1.3-3 splines 4.1.0 grid 4.1.0
[5] lattice 0.20-44
```

Appendix 4. PRS associations with trajectories after controlling for additional variables suggested by reviewers

In our analyses of PRS associations with trajectories, we controlled for a total of 20 demographic, ancestral, and trauma-related variables (i.e., potentially traumatic experiences; PTEs). Reviewers suggested additional analyses controlling for length of deployment and disaggregating the types of PTEs from the pre-deployment PTE indicators that we used in our analyses. The resulting analyses controlled for 49 variables, including the original control variables, and the addition of length of deployment and 30 types of PTEs measured at predeployment (15 related to prior deployments such as combat, direct/indirect fire, getting wounded by the enemy, having a "close call", member of unit wounded or killed, direct responsibility for death of an enemy combatant, direct responsibility for non-combatant, direct responsibility for death of ally, saving the life of a soldier/civilian, seeing homes/villages destroyed, exposure to sights/sounds/smells of death and dying, witnessing violence toward locals/non-combatants, physical assault, sexual assault, bullied by members of unit; and 15 related to lifetime experiences such as physical assault, sexual assault, serious assault of friend/relative, murder of friend relative, suicide of friend/relative, attempted suicide of friend/relative, combat death of friend/relative, accidental death of friend/relative, witnessing someone being injured or killed, discovering/handling dead body, life threatening illness or injury, natural disaster where death was possible, other life-threatening experience, bullied during childhood/adolescence, and life-threatening experience of friend/relative.)

In the models that controlled for these additional covariates, there were some fluctuations in the *p*-values of PRS associations with trajectories: PTSD p = .029, MDD p = .006, AUD p = .807, neuroticism p = .027, schizophrenia p = .919, suicide attempt p = .817. There were also some fluctuations in the adjusted odds ratios AOR for the PTSD and MDD PRS, though the pattern of results was similar and in some cases, the AOR were nearly identical.

	Original analyses	Additional analyses
	controlling for 20	controlling for 49
	covariates	covariates
PTSD-PRS		
High-Severity vs Increasing-Severity	1.09	1.07
	[0.92, 1.30]	[0.89, 1.28]
High-Severity vs Decreasing-Severity	1.13	1.11
	[0.94, 1.36]	[0.92, 1.33]
High-Severity vs Low-Severity	1.23	1.19
	[1.06, 1.43]	[1.02, 1.40]
Increasing-Severity vs Decreasing-Severity	1.04	1.04
	[0.89, 1.20]	[0.89, 1.21]
Increasing-Severity vs Low-Severity	1.12	1.12
	[1.01, 1.25]	[1.01, 1.25]
Decreasing-Severity vs Low-Severity	1.08	1.08
	[0.96, 1.22]	[0.96, 1.22]
MDD-PRS		
High-Severity vs Increasing-Severity	1.02	1.02
	[0.86, 1.22]	[0.85, 1.22]
High-Severity vs Decreasing-Severity	1.02	0.99
	[0.85, 1.22]	[0.83, 1.19]
High-Severity vs Low-Severity	1.18	1.16
	[1.02, 1.37]	[0.99, 1.35]
Increasing-Severity vs Decreasing-Severity	0.99	0.97
	[0.86, 1.15]	[0.84, 1.13]
Increasing-Severity vs Low-Severity	1.16	1.14
	[1.04, 1.29]	[1.02, 1.27]
Decreasing-Severity vs Low-Severity	1.16	1.17
	[1.03, 1.31]	[1.03, 1.32]