

Examining Lagged Effects of Stress in Predicting Psoriasis Severity

These data come from a study of weekly fluctuation in psoriasis severity, as also featured in Example 4 of CLDP 944. The goal of these analyses is to examine how weekly fluctuations in psoriasis severity may be predicted by weekly fluctuation in stress, and at what time lag these effects might be observed.

SAS Data Setup for Lags:

```
* Location for original SPSS portable multivariate file for these models;
LIBNAME import SPSS "C:\Dropbox\_Archive\Example Data\Psoriasis\p2stress.por";
DATA work.p2stack; SET import.p2stress;
  * Calculating person mean for time-varying predictor;
  pmStress = MEAN(of lesneg1-lesneg8); LABEL pmStress = "Person-Mean Stress";
  * Stacking data;
  Week=1; severity=sapasi1; stress=lesneg1; OUTPUT;
  Week=2; severity=sapasi2; stress=lesneg2; OUTPUT;
  Week=3; severity=sapasi3; stress=lesneg3; OUTPUT;
  Week=4; severity=sapasi4; stress=lesneg4; OUTPUT;
  Week=5; severity=sapasi5; stress=lesneg5; OUTPUT;
  Week=6; severity=sapasi6; stress=lesneg6; OUTPUT;
  Week=7; severity=sapasi7; stress=lesneg7; OUTPUT;
  Week=8; severity=sapasi8; stress=lesneg8; OUTPUT;
LABEL week = "Week of Study" severity = "Psoriasis Severity"
      stress = "Weekly Stress"; RUN;

* Creating BP/WP predictors for analysis;
DATA work.p2stack; SET work.p2stack;
* Selecting complete cases;
  WHERE NMISS(severity, stress)=0;
* Creating between-person indicator for LES;
  BPstress2 = pmStress - 2; LABEL BPstress2 = "Between-Person Stress (0=2)";
* Person-centering stress to be a level 1 predictor;
  WPstressLag0 = stress - pmStress; LABEL WPstressLag0 = "Within-Person Stress Lag=0";
RUN;
* Creating lagged variables;
PROC EXPAND DATA=work.p2stack OUT=work.p2lag;
  BY subid;
  ID week;          * Old = new name;
  CONVERT WPstressLag0=WPstressLag1 / METHOD=NONE TRANSFORMOUT=(LAG 1);
  CONVERT WPstressLag0=WPstressLag2 / METHOD=NONE TRANSFORMOUT=(LAG 2); RUN;
```

Example of Data File after Lagging:

subid	Week	severity	pmStress	PMstress2	stress	WPstressLag0	WPstressLag1	WPstressLag2
100	1	1.93	0.60	-1.40	0.00	-0.60	.	.
100	2	2.00	0.60	-1.40	0.00	-0.60	-0.60	.
100	3	1.85	0.60	-1.40	1.10	0.50	-0.60	-0.60
100	4	1.68	0.60	-1.40	0.69	0.09	0.50	-0.60
100	5	1.82	0.60	-1.40	0.00	-0.60	0.09	0.50
100	6	0.00	0.60	-1.40	1.39	0.79	-0.60	0.09
100	7	1.47	0.60	-1.40	1.61	1.01	0.79	-0.60
100	8	0.66	0.60	-1.40	0.00	-0.60	1.01	0.79
101	1	2.10	0.45	-1.55	0.69	0.24	.	.
101	2	1.70	0.45	-1.55	1.10	0.64	0.24	.
101	3	1.50	0.45	-1.55	0.00	-0.45	0.64	0.24
101	4	2.00	0.45	-1.55	0.00	-0.45	-0.45	0.64
101	5	1.42	0.45	-1.55	0.00	-0.45	-0.45	-0.45
101	6	1.26	0.45	-1.55	0.00	-0.45	-0.45	-0.45
101	7	0.98	0.45	-1.55	1.39	0.93	-0.45	-0.45
101	8	.	0.45	-1.55	.	.	0.93	-0.45

1a) Empty Means, Random Intercept Model for Severity (DV)

```
TITLE 'Empty Means, Random Intercept Model for Severity Outcome';
PROC MIXED DATA=work.p2lag COVTEST NOCLPRINT NAMELEN=100 METHOD=REML;
  CLASS subid week;
  MODEL severity = / SOLUTION DDFM=Satterthwaite;
  RANDOM INTERCEPT / VCORR TYPE=UN SUBJECT=subid;
  REPEATED week / TYPE=VC SUBJECT=subid; RUN;
```

$$\begin{aligned} \text{Level 1: } \text{Severity}_{ti} &= \beta_{0i} + e_i \\ \text{Level 2: } \beta_{0i} &= \gamma_{00} + U_{0i} \end{aligned}$$

Covariance Parameter Estimates					
Cov Parm	Subject	Estimate	Standard Error	Z Value	Pr Z
UN(1,1)	SUBID	0.6667	0.08782	7.59	<.0001
week	SUBID	0.1318	0.006780	19.44	<.0001

ICC for Severity:
 $.6667 / (.6667 + .1318) = .83$

Houston, we may have a problem

1b) Empty Model for Stress (Time-Varying Predictor)

```
TITLE 'Empty Means, Random Intercept Model for Stress TV Predictor';
PROC MIXED DATA= work.p2lag COVTEST NOCLPRINT NAMELEN=100 METHOD=REML;
  CLASS subid week;
  MODEL stress = / SOLUTION DDFM=Satterthwaite;
  RANDOM INTERCEPT / VCORR TYPE=UN SUBJECT=subid;
  REPEATED week / TYPE=VC SUBJECT=subid; RUN;
```

$$\begin{aligned} \text{Level 1: } \text{Stress}_{ti} &= \beta_{0i} + e_i \\ \text{Level 2: } \beta_{0i} &= \gamma_{00} + U_{0i} \end{aligned}$$

Covariance Parameter Estimates					
Cov Parm	Subject	Estimate	Standard Error	Z Value	Pr Z
UN(1,1)	SUBID	0.3562	0.05102	6.98	<.0001
week	SUBID	0.2519	0.01298	19.41	<.0001

ICC for Stress:
 $.3562 / (.3562 + .2519) = .56$

At least stress is time-varying!

Our best-fitting unconditional model for time (as found in Example 4 of CLDP 944) included a random intercept variance in the **G** matrix and a lag-3 correlation with heterogeneous residual variances in the **R** matrix. We now add fixed effects of predictors to that model.

2) Predicting Severity from Between-Person Effect of Stress

```
TITLE "SAS: BP Effect of Stress";
PROC MIXED DATA= work.p2lag COVTEST NOCLPRINT NAMELEN=100 METHOD=REML;
  CLASS subid week;
  MODEL severity = PMstress2 / SOLUTION DDFM=Satterthwaite;
  RANDOM INTERCEPT / TYPE=UN SUBJECT=subid;
  REPEATED week / TYPE=TOEPH(4) SUBJECT=subid; RUN;
```

$$\begin{aligned} \text{Level 1: } \text{Severity}_{ti} &= \beta_{0i} + e_{ti} \\ \text{Level 2: } \beta_{0i} &= \gamma_{00} + \gamma_{01} (\bar{\text{Stress}}_i - 2) + U_{0i} \end{aligned}$$

Dimensions	
Subjects	124
Max Obs Per Subject	8

Number of Observations	
Number of Observations Read	880
Number of Observations Used	880
Number of Observations Not Used	0

Interpret each fixed effect:

Solution for Fixed Effects					
Effect	Estimate	Error	DF	t Value	Pr > t
Intercept	1.9464	0.1272	123	15.31	<.0001
PMstress2	0.4572	0.1097	123	4.17	<.0001

3a) Predicting Severity from Between- and Within-Person (Simultaneous) Effects of Stress

$$\begin{aligned} \text{Level 1: Severity}_{ti} &= \beta_{0i} + \beta_{1i} (\text{Lag0Stress}_{ti} - \bar{\text{Stress}}_i) + e_{ti} \\ \text{Level 2: Intercept: } \beta_{0i} &= \gamma_{00} + \gamma_{01} (\bar{\text{Stress}}_i - 2) + U_{0i} \\ \text{Lag-0 WP Stress: } \beta_{1i} &= \gamma_{10} \end{aligned}$$

```

TITLE "Add WP Effect of Stress at 0-Week Lag";
PROC MIXED DATA= work.p2lag COVTEST NOCLPRINT NAMELEN=100 METHOD=REML;
  CLASS subid week;
  MODEL severity = PMstress2 WPstressLag0 / SOLUTION DDFM=Satterthwaite;
  RANDOM INTERCEPT / TYPE=UN SUBJECT=subid;
  REPEATED week / TYPE=TOEPH(4) SUBJECT=subid; RUN;

```

Dimensions

Subjects 124
Max Obs Per Subject 8

Note: 8 observations per person are used here.

Number of Observations

Number of Observations Used 880
Number of Observations Not Used 0

Solution for Fixed Effects Standard

Effect	Estimate	Error	DF	t Value	Pr > t
Intercept	1.9463	0.1272	123	15.30	<.0001
PMstress2	0.4572	0.1097	123	4.17	<.0001
WPstressLag0	-0.00720	0.02348	680	-0.31	0.7591

Interpret the effect of WPstressLag0:

3b) Predicting Severity from Between- and Within-Person (+1-Week Lag) Effects of Stress

$$\begin{aligned}
\text{Level 1: Severity}_{ti} &= \beta_{0i} + \beta_{1i} (\text{Lag0Stress}_{ti} - \bar{\text{Stress}}_i) + \beta_{2i} (\text{Lag1Stress}_{ti} - \bar{\text{Stress}}_i) + e_{ti} \\
\text{Level 2: Intercept: } \beta_{0i} &= \gamma_{00} + \gamma_{01} (\bar{\text{Stress}}_i - 2) + U_{0i} \\
\text{Lag-0 WP Stress: } \beta_{1i} &= \gamma_{10} \\
\text{Lag-1 WP Stress: } \beta_{2i} &= \gamma_{20}
\end{aligned}$$

```

TITLE "Add WP Effect of Stress at 1-Week Lag";
PROC MIXED DATA= work.p2lag COVTEST NOCLPRINT NAMELEN=100 METHOD=REML;
  CLASS subid week;
  MODEL severity = PMstress2 WPstressLag0 WPstressLag1
    / SOLUTION DDFM=Satterthwaite;
  RANDOM INTERCEPT / TYPE=UN SUBJECT=subid;
  REPEATED week / TYPE=TOEPH(4) SUBJECT=subid: RUN;

```

Dimensions
Subjects 122

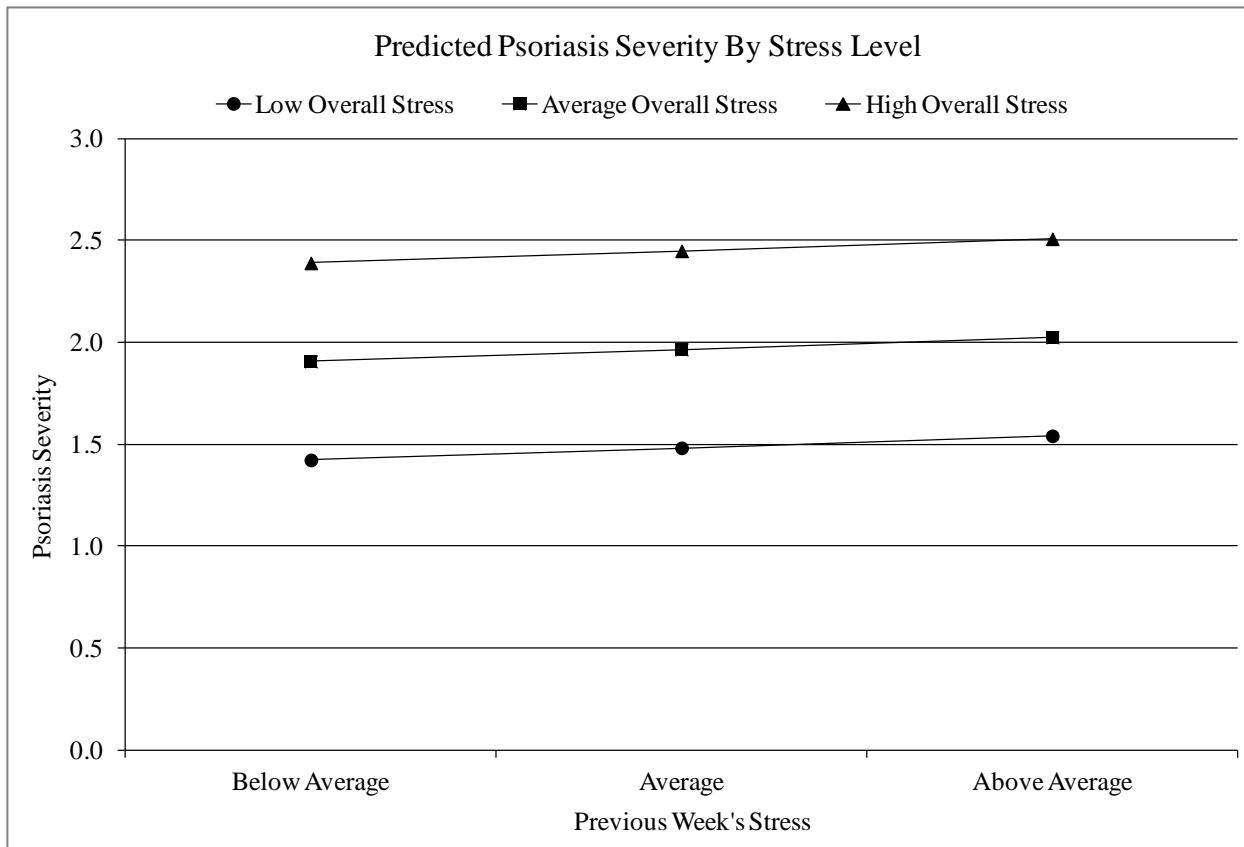
Number of Observations	
Number of Observations Used	756
Number of Observations Not Used	124

Note the change in number of observations used: only 7 occasions per person in adding a 1-week lag within-person predictor. Estimating models on different samples means we cannot compare variance components (e.g., compute pseudo- R^2) or do any kind of LRT for change in model fit.

Effect	Solution for Fixed Effects				
	Estimate	Error	DF	t Value	Pr > t
Intercept	1.9649	0.1307	121	15.04	<.0001
PMstress2	0.4833	0.1132	121	4.27	<.0001
WPstressLag0	0.01976	0.02662	557	0.74	0.4582
WPstressLag1	0.05919	0.02509	545	2.36	0.0187

Interpret the effect of WPstressLag1:

Calculated predicted values in excel worksheet to plot these effects:



3b) Predicting Severity from Between- and Within-Person (+2-Week Lag) Effects of Stress

$$\text{Level 1: } \text{Severity}_{ti} = \beta_{0i} + \beta_{1i} (\text{Lag0Stress}_{ti} - \bar{\text{Stress}}_i) + \beta_{2i} (\text{Lag1Stress}_{ti} - \bar{\text{Stress}}_i) + \beta_{3i} (\text{Lag2Stress}_{ti} - \bar{\text{Stress}}_i) + e_{ti}$$

$$\text{Level 2: Intercept: } \beta_{0i} = \gamma_{00} + \gamma_{01} (\bar{\text{Stress}}_i - 2) + U_{0i}$$

$$\text{Lag-0 WP Stress: } \beta_{1i} = \gamma_{10}$$

$$\text{Lag-1 WP Stress: } \beta_{2i} = \gamma_{20}$$

$$\text{Lag-2 WP Stress: } \beta_{3i} = \gamma_{30}$$

```

TITLE "Add WP Effect of Stress at 2-Week Lag";
PROC MIXED DATA= work.p2lag COVTEST NOCLPRINT NAMELEN=100 METHOD=REML;
  CLASS subid week;
  MODEL severity = PMstress2 WPstressLag0 WPstressLag1 WPstressLag2
    / SOLUTION DDFM=Satterthwaite;
  RANDOM INTERCEPT / TYPE=UN SUBJECT=subid;
  REPEATED week / TYPE=TOEPh(4) SUBJECT=subid; RUN;

```

Dimensions	
Subjects	119
Max Obs Per Subject	6
Number of Observations	
Number of Observations Used	634
Number of Observations Not Used	246

Note the change in number of observations used: only 6 occasions per person in adding a 2-week lag within-person predictor.

Solution for Fixed Effects

Effect	Estimate	Error	DF	t Value	Pr > t
Intercept	1.9295	0.1397	116	13.81	<.0001
PMstress2	0.4771	0.1199	117	3.98	0.0001
WPstressLag0	0.01772	0.03117	475	0.57	0.5699
WPstressLag1	0.08317	0.02946	380	2.82	0.0050
WPstressLag2	0.04163	0.03089	489	1.35	0.1785

4) Examining Fixed Effects under Alternative Models for the Variances (Listed in Order of Fit)

Unstructured R:	Solution for Fixed Effects Standard Effect Estimate Error DF t Value Pr > t Intercept 1.9860 0.1274 120 15.59 <.0001 PMstress2 0.4723 0.1107 122 4.27 <.0001 WPstressLag0 0.02335 0.02603 529 0.90 0.3701 WPstressLag1 0.06751 0.02418 494 2.79 0.0054
Random Intercept + 3-Lag Heterogeneous Toeplitz R:	Solution for Fixed Effects Standard Effect Estimate Error DF t Value Pr > t Intercept 1.9649 0.1307 121 15.04 <.0001 PMstress2 0.4833 0.1132 121 4.27 <.0001 WPstressLag0 0.01976 0.02662 557 0.74 0.4582 WPstressLag1 0.05919 0.02509 545 2.36 0.0187
AR(1) in R only:	Solution for Fixed Effects Standard Effect Estimate Error DF t Value Pr > t Intercept 1.9868 0.1218 131 16.31 <.0001 PMstress2 0.4828 0.1055 132 4.57 <.0001 WPstressLag0 0.03198 0.02866 613 1.12 0.2650 WPstressLag1 0.05776 0.02726 614 2.12 0.0345
Random Intercept Only (CS):	Solution for Fixed Effects Standard Effect Estimate Error DF t Value Pr > t Intercept 1.9522 0.1307 122 14.94 <.0001 PMstress2 0.4872 0.1131 122 4.31 <.0001 WPstressLag0 0.04558 0.02926 633 1.56 0.1199 WPstressLag1 0.05712 0.02831 632 2.02 0.0440
Residual Variance Only (VC):	Solution for Fixed Effects Standard Effect Estimate Error DF t Value Pr > t Intercept 1.9259 0.05754 752 33.47 <.0001 PMstress2 0.4842 0.04999 752 9.69 <.0001 WPstressLag0 0.006840 0.06633 752 0.10 0.9179 WPstressLag1 0.06815 0.06450 752 1.06 0.2911