

The datasets can be found at
<http://www.biostat.jhsph.edu/~fdominic/teaching/LDA/lda.html> under "DATA SETS";

```
data hiv;  
infile "C:\...\hivstudy.txt" dlm=tab;  
input ID Month CD4 Group;  
run;
```

```
ods rtf file="C:\...\Examples\09 - Profile and Parametric examples2.rtf";
```

```
proc print data = hiv (obs=10);  
run;
```

Obs	ID	Month	CD4	Group
1	1	0	658	1
2	1	2	543	1
3	1	4	520	1
4	1	6	563	1
5	1	8	389	1
6	1	10	371	1
7	2	0	500	1
8	2	2	419	1
9	2	4	431	1
10	2	6	285	1

*Now we will include the mean line on the graph by TRT;

```
proc sort data=hiv;  
by Group Month;
```

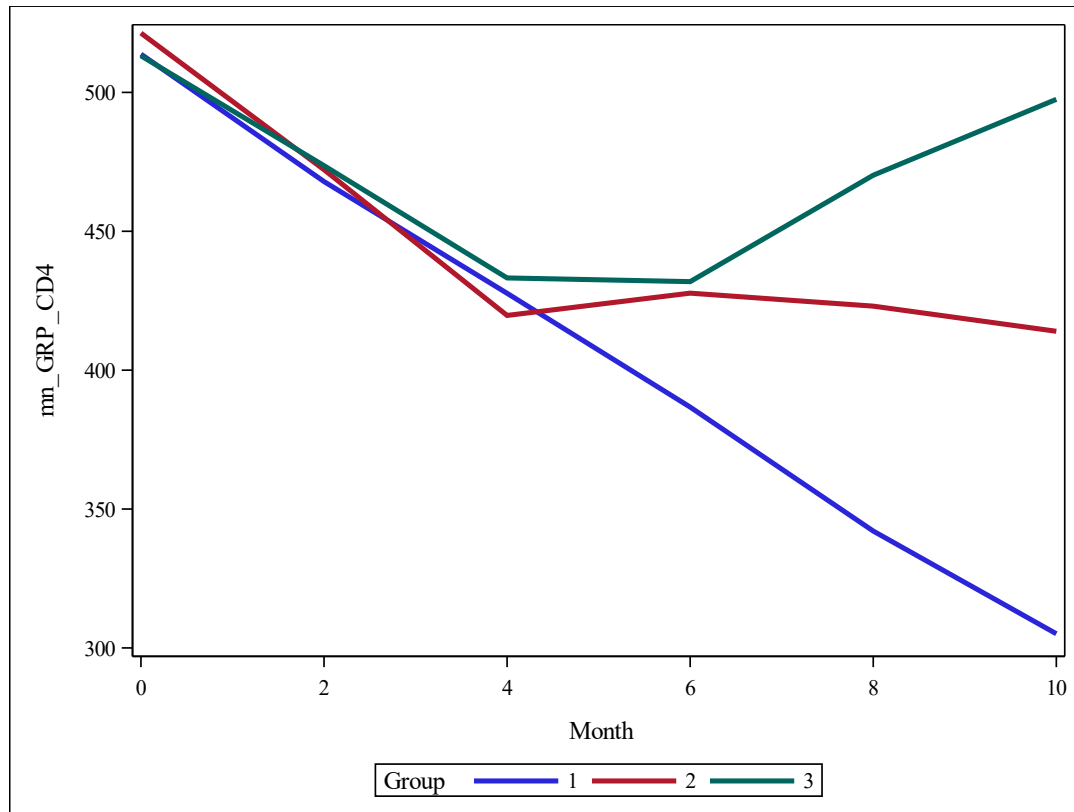
*Calculate the mean by week;

```
proc means mean data=hiv noprint;  
by Group Month;  
var CD4;  
output out = MN_GRP_dat mean = mn_GRP_CD4;  
run;
```

*First, let's look at the mean by TRT group;

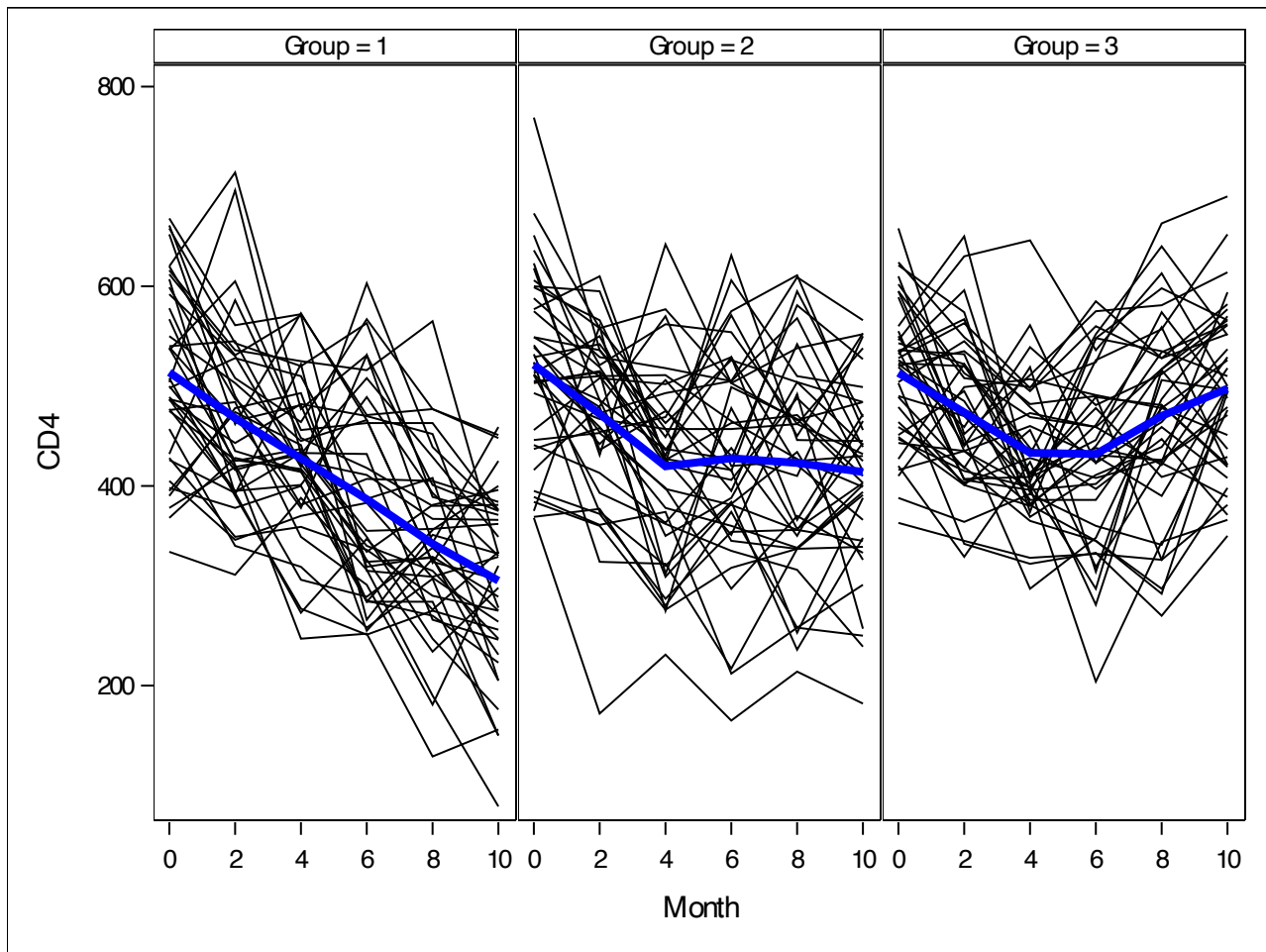
```
Proc SGplot data = MN_GRP_dat;  
series x=Month y=mn_GRP_CD4 / group =Group LineAttrs= (pattern=1  
thickness=3);
```

```
run;
```



```
*We'll look at the means by TRT group with the rest of the data;  
data stacked_hiv;  
set hiv MN_GRP_dat;  
run;
```

```
* Now we'll combine them onto one plot with two panels;  
Proc SGpanel data = stacked_hiv;  
PanelBy Group / columns=3;  
series x=Month y=CD4 / group =ID LineAttrs= (pattern=1 color="black");  
series x=Month y=mn_GRP_CD4 / LineAttrs= (pattern=1 color="blue"  
thickness=4);  
run;
```



```
data hiv_spline;
set hiv;
sp_mn1 = min(month,4);
sp_mn2 = max(0,month-4);
run;
```

```
proc mixed data=hiv_spline;
class ID group(ref='1');
model CD4 = group month sp_mn2 group*month group*sp_mn2/solution;
random intercept/ subject=ID type=UN g gcorr;
run;
```

Estimated V Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	7932.96	4252.14	4252.14	4252.14	4252.14	4252.14
2	4252.14	7932.96	4252.14	4252.14	4252.14	4252.14
3	4252.14	4252.14	7932.96	4252.14	4252.14	4252.14
4	4252.14	4252.14	4252.14	7932.96	4252.14	4252.14

Estimated V Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
5	4252.14	4252.14	4252.14	4252.14	7932.96	4252.14
6	4252.14	4252.14	4252.14	4252.14	4252.14	7932.96

Estimated V Correlation Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	1.0000	0.5360	0.5360	0.5360	0.5360	0.5360
2	0.5360	1.0000	0.5360	0.5360	0.5360	0.5360
3	0.5360	0.5360	1.0000	0.5360	0.5360	0.5360
4	0.5360	0.5360	0.5360	1.0000	0.5360	0.5360
5	0.5360	0.5360	0.5360	0.5360	1.0000	0.5360
6	0.5360	0.5360	0.5360	0.5360	0.5360	1.0000

Covariance Parameter Estimates					
Cov Parm	Subject	Estimate	Standard Error	Z Value	Pr > Z
UN(1,1)	ID	4252.14	637.15	6.67	<.0001
Residual		3680.82	213.58	17.23	<.0001

Fit Statistics	
-2 Res Log Likelihood	8153.7
AIC (Smaller is Better)	8157.7
AICC (Smaller is Better)	8157.7
BIC (Smaller is Better)	8163.3

```
proc mixed data=hiv_spline;
class ID group(ref='1');
model CD4 = group month sp_mn2 group*month group*sp_mn2/solution;
random intercept month/ subject=ID type=UN g gcorr;
run;
```

Estimated G Matrix				
Row	Effect	ID	Col1	Col2
1	Intercept	1	3315.86	85.1966
2	Month	1	85.1966	3.7143

Estimated G Correlation Matrix				
Row	Effect	ID	Col1	Col2
1	Intercept	1	1.0000	0.7677
2	Month	1	0.7677	1.0000

Estimated V Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	6945.46	3486.25	3656.64	3827.04	3997.43	4167.82
2	3486.25	7301.10	3856.75	4042.00	4227.25	4412.50
3	3656.64	3856.75	7686.46	4256.97	4457.07	4657.18
4	3827.04	4042.00	4256.97	8101.53	4686.89	4901.86
5	3997.43	4227.25	4457.07	4686.89	8546.32	5146.54
6	4167.82	4412.50	4657.18	4901.86	5146.54	9020.82

Estimated V Correlation Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	1.0000	0.4896	0.5005	0.5102	0.5188	0.5265
2	0.4896	1.0000	0.5148	0.5256	0.5351	0.5437
3	0.5005	0.5148	1.0000	0.5395	0.5499	0.5593
4	0.5102	0.5256	0.5395	1.0000	0.5633	0.5734
5	0.5188	0.5351	0.5499	0.5633	1.0000	0.5861
6	0.5265	0.5437	0.5593	0.5734	0.5861	1.0000

Covariance Parameter Estimates					
Cov Parm	Subject	Estimate	Standard Error	Z Value	Pr Z
UN(1,1)	ID	3315.86	693.12	4.78	<.0001
UN(2,1)	ID	85.1966	54.9406	1.55	0.1210
UN(2,2)	ID	3.7143	8.0032	0.46	0.3213
Residual		3629.60	235.03	15.44	<.0001

Fit Statistics	
-2 Res Log Likelihood	8148.7
AIC (Smaller is Better)	8156.7
AICC (Smaller is Better)	8156.8
BIC (Smaller is Better)	8167.9

```

proc mixed data=hiv_spline;
class ID group(ref='1');
model CD4 = group month sp_mn2 group*month group*sp_mn2/solution;
random intercept month sp_mn2/ subject=ID type=UN g gcorr;
run;

```

Estimated G Matrix					
Row	Effect	ID	Col1	Col2	Col3
1	Intercept	1	3450.37	46.6611	71.0062
2	Month	1	46.6611	9.1407	-15.5416
3	sp_mn2	1	71.0062	-15.5416	37.1470

Estimated G Correlation Matrix					
Row	Effect	ID	Col1	Col2	Col3
1	Intercept	1	1.0000	0.2627	0.1983
2	Month	1	0.2627	1.0000	-0.8434
3	sp_mn2	1	0.1983	-0.8434	1.0000

Estimated V Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	7040.40	3543.69	3637.01	3872.35	4107.68	4343.01
2	3543.69	7263.61	3803.46	4013.19	4222.92	4432.65
3	3637.01	3803.46	7559.94	4154.03	4338.16	4522.29
4	3872.35	4013.19	4154.03	7989.01	4643.92	4888.87
5	4107.68	4222.92	4338.16	4643.92	8539.72	5255.45
6	4343.01	4432.65	4522.29	4888.87	5255.45	9212.07

Estimated V Correlation Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	1.0000	0.4955	0.4985	0.5163	0.5298	0.5393
2	0.4955	1.0000	0.5133	0.5268	0.5362	0.5419
3	0.4985	0.5133	1.0000	0.5345	0.5399	0.5419
4	0.5163	0.5268	0.5345	1.0000	0.5622	0.5699
5	0.5298	0.5362	0.5399	0.5622	1.0000	0.5925
6	0.5393	0.5419	0.5419	0.5699	0.5925	1.0000

Covariance Parameter Estimates					
Cov Parm	Subject	Estimate	Standard Error	Z Value	Pr Z
UN(1,1)	ID	3450.37	866.70	3.98	<.0001
UN(2,1)	ID	46.6611	175.33	0.27	0.7901
UN(2,2)	ID	9.1407	57.9141	0.16	0.4373
UN(3,1)	ID	71.0062	247.20	0.29	0.7739
UN(3,2)	ID	-15.5416	82.4936	-0.19	0.8506
UN(3,3)	ID	37.1470	128.65	0.29	0.3864
Residual		3590.03	267.59	13.42	<.0001

Fit Statistics	
-2 Res Log Likelihood	8148.4
AIC (Smaller is Better)	8162.4
AICC (Smaller is Better)	8162.5
BIC (Smaller is Better)	8181.9

```
proc mixed data=hiv_spline method=ml;
class ID group(ref='1');
model CD4 = group month sp_mn2 group*month group*sp_mn2/solution;
random intercept/ subject=ID type=UN g gcorr;
run;
```

Fit Statistics	
-2 Log Likelihood	8194.5
AIC (Smaller is Better)	8216.5
AICC (Smaller is Better)	8216.9
BIC (Smaller is Better)	8247.2

```
proc mixed data=hiv_spline method=ml;
class ID month group(ref='1');
model CD4 = group month group*month /solution;
random intercept/ subject=ID type=UN g gcorr;
run;
```

Fit Statistics	
-2 Log Likelihood	8189.6
AIC (Smaller is Better)	8229.6
AICC (Smaller is Better)	8230.8
BIC (Smaller is Better)	8285.4

```

proc mixed data=hiv_spline;
class ID month group(ref='1');
model CD4 = group month group*month /solution;
random intercept/ subject=ID type=UN g gcorr;
run;

```

Estimated V Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	7954.81	4247.77	4247.77	4247.77	4247.77	4247.77
2	4247.77	7954.81	4247.77	4247.77	4247.77	4247.77
3	4247.77	4247.77	7954.81	4247.77	4247.77	4247.77
4	4247.77	4247.77	4247.77	7954.81	4247.77	4247.77
5	4247.77	4247.77	4247.77	4247.77	7954.81	4247.77
6	4247.77	4247.77	4247.77	4247.77	4247.77	7954.81

Estimated V Correlation Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	1.0000	0.5340	0.5340	0.5340	0.5340	0.5340
2	0.5340	1.0000	0.5340	0.5340	0.5340	0.5340
3	0.5340	0.5340	1.0000	0.5340	0.5340	0.5340
4	0.5340	0.5340	0.5340	1.0000	0.5340	0.5340
5	0.5340	0.5340	0.5340	0.5340	1.0000	0.5340
6	0.5340	0.5340	0.5340	0.5340	0.5340	1.0000

Covariance Parameter Estimates		
Cov Parm	Subject	Estimate
UN(1,1)	ID	4247.77
Residual		3707.04

Fit Statistics	
-2 Res Log Likelihood	8069.1
AIC (Smaller is Better)	8073.1
AICC (Smaller is Better)	8073.1
BIC (Smaller is Better)	8078.6

```

proc mixed data=hiv_spline;
class ID month group(ref='1');
model CD4 = group month group*month /solution;
repeated month/ subject=ID type=CSH r rcorr;
run;

```


Estimated R Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	7311.02	3829.84	3840.99	4434.63	4322.16	4003.54
2	3829.84	7041.51	3769.53	4352.12	4241.75	3929.05
3	3840.99	3769.53	7082.56	4364.79	4254.09	3940.49
4	4434.63	4352.12	4364.79	9441.01	4911.58	4549.50
5	4322.16	4241.75	4254.09	4911.58	8968.22	4434.12
6	4003.54	3929.05	3940.49	4549.50	4434.12	7694.70

Estimated R Correlation Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	1.0000	0.5338	0.5338	0.5338	0.5338	0.5338
2	0.5338	1.0000	0.5338	0.5338	0.5338	0.5338
3	0.5338	0.5338	1.0000	0.5338	0.5338	0.5338
4	0.5338	0.5338	0.5338	1.0000	0.5338	0.5338
5	0.5338	0.5338	0.5338	0.5338	1.0000	0.5338
6	0.5338	0.5338	0.5338	0.5338	0.5338	1.0000

Covariance Parameter Estimates		
Cov Parm	Subject	Estimate
Var(1)	ID	7311.02
Var(2)	ID	7041.51
Var(3)	ID	7082.56
Var(4)	ID	9441.01
Var(5)	ID	8968.22
Var(6)	ID	7694.70
CSH	ID	0.5338

Fit Statistics	
-2 Res Log Likelihood	8061.8
AIC (Smaller is Better)	8075.8
AICC (Smaller is Better)	8076.0
BIC (Smaller is Better)	8095.3

If we have good motivation for the spline term at 4 months, I would use the spline model. If there wasn't motivation for the spline term, I would use the profile analysis.

```
proc mixed data=hiv_spline;
class ID group(ref='1');
model CD4 = group month sp_mn2 group*month group*sp_mn2/solution;
random intercept month/ subject=ID type=UN g gcorr;
run;
```

Model Information	
Data Set	WORK.HIV_SPLINE
Dependent Variable	CD4
Covariance Structure	Unstructured
Subject Effect	ID
Estimation Method	REML
Residual Variance Method	Profile
Fixed Effects SE Method	Model-Based
Degrees of Freedom Method	Containment

Class Level Information		
Class	Levels	Values
ID	120	1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47 48 49 50 51 52 53 54 55 56 57 58 59 60 61 62 63 64 65 66 67 68 69 70 71 72 73 74 75 76 77 78 79 80 81 82 83 84 85 86 87 88 89 90 91 92 93 94 95 96 97 98 99 100 101 102 103 104 105 106 107 108 109 110 111 112 113 114 115 116 117 118 119 120
Group	3	2 3 1

Dimensions	
Covariance Parameters	4
Columns in X	12
Columns in Z per Subject	2
Subjects	120
Max Obs per Subject	6

Number of Observations	
Number of Observations Read	720
Number of Observations Used	720
Number of Observations Not Used	0

Iteration History			
Iteration	Evaluations	-2 Res Log Like	Criterion
0	1	8452.53002008	
1	1	8148.70194016	0.00000000

Convergence criteria met.

Estimated G Matrix				
Row	Effect	ID	Col1	Col2
1	Intercept	1	3315.86	85.1966
2	Month	1	85.1966	3.7143

Estimated G Correlation Matrix				
Row	Effect	ID	Col1	Col2
1	Intercept	1	1.0000	0.7677
2	Month	1	0.7677	1.0000

Estimated V Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	6945.46	3486.25	3656.64	3827.04	3997.43	4167.82
2	3486.25	7301.10	3856.75	4042.00	4227.25	4412.50
3	3656.64	3856.75	7686.46	4256.97	4457.07	4657.18
4	3827.04	4042.00	4256.97	8101.53	4686.89	4901.86
5	3997.43	4227.25	4457.07	4686.89	8546.32	5146.54
6	4167.82	4412.50	4657.18	4901.86	5146.54	9020.82

Estimated V Correlation Matrix for ID 1						
Row	Col1	Col2	Col3	Col4	Col5	Col6
1	1.0000	0.4896	0.5005	0.5102	0.5188	0.5265
2	0.4896	1.0000	0.5148	0.5256	0.5351	0.5437
3	0.5005	0.5148	1.0000	0.5395	0.5499	0.5593
4	0.5102	0.5256	0.5395	1.0000	0.5633	0.5734
5	0.5188	0.5351	0.5499	0.5633	1.0000	0.5861
6	0.5265	0.5437	0.5593	0.5734	0.5861	1.0000

Covariance Parameter Estimates					
Cov Parm	Subject	Estimate	Standard Error	Z Value	Pr Z
UN(1,1)	ID	3315.86	693.12	4.78	<.0001
UN(2,1)	ID	85.1966	54.9406	1.55	0.1210
UN(2,2)	ID	3.7143	8.0032	0.46	0.3213
Residual		3629.60	235.03	15.44	<.0001

Fit Statistics	
-2 Res Log Likelihood	8148.7
AIC (Smaller is Better)	8156.7
AICC (Smaller is Better)	8156.8
BIC (Smaller is Better)	8167.9

Null Model Likelihood Ratio Test		
DF	Chi-Square	Pr > ChiSq
3	303.83	<.0001

Solution for Fixed Effects						
Effect	Group	Estimate	Standard Error	DF	t Value	Pr > t
Intercept		512.86	12.5586	117	40.84	<.0001
Group	2	8.2469	17.7605	477	0.46	0.6426
Group	3	2.1689	17.7605	477	0.12	0.9029
Group	1	0
Month		-21.5651	3.1056	117	-6.94	<.0001
sp_mn2		1.0849	4.5710	477	0.24	0.8125
Month*Group	2	-2.6579	4.3919	477	-0.61	0.5453
Month*Group	3	-0.9908	4.3919	477	-0.23	0.8216
Month*Group	1	0
sp_mn2*Group	2	22.0859	6.4644	477	3.42	0.0007
sp_mn2*Group	3	32.7530	6.4644	477	5.07	<.0001
sp_mn2*Group	1	0

Type 3 Tests of Fixed Effects				
Effect	Num DF	Den DF	F Value	Pr > F
Group	2	477	0.12	0.8906
Month	1	117	161.44	<.0001
sp_mn2	1	477	53.84	<.0001
Month*Group	2	477	0.19	0.8294
sp_mn2*Group	2	477	13.36	<.0001

Let's compare these two models graphically:

```
proc mixed data=hiv_spline;
class ID month group(ref='1');
model CD4 = group month group*month/solution outpm=pred2;
repeated month/ subject=ID type=CS r rcorr;
run;
```

```
proc sgplot data=pred;
series y=pred x=month / group=group;
run;
```

```
proc sgplot data=pred2;
series y=pred x=month / group=group;
run;
```

