

## Appendix: Supplementary Tables, Statistical Code and Calculations

**Table S1 Model Comparisons and Variable Selection**

MODEL (# of variables)	Full (10)*	Full <sub>LT</sub> (10)*	Reduced <sub>LT</sub> , A (8)*	Reduced <sub>LT</sub> , B (6)*	Parsimonious <sub>LT</sub> (5)*	ACR (4)
Included covariates that were not statistically significant	Sex, New headache, Temporal artery abnormality, Diplopia, CRP	Sex, New headache, Temporal artery abnormality, Diplopia, log ESR	New headache, Temporal artery abnormality, log ESR	Temporal artery abnormality		New headache (Age category omitted)
AUROC	.8084	.8196	.8185	.8163	.8164	.6339
p Hosmer-Lemeshow	.3590	.5485	.1722	.0182	.8115	.2227
Brier score	.1381	.1313	.2434	.2426	.1335	.1792
p Spiegelhalter z-test	.4833	.5811	<.001	<.001	.6662	.5213
AIC	480.6595	466.2297	462.7570	460.603	461.0216	576.4964
BIC	527.6611	513.2314	501.2129	490.5135	486.6589	593.5500
Internal validation AUROC		0.803 (95% CI 0.757, 0.849)			0.810 (95% CI 0.766, 0.854).	
External validation AUROC		Range 0.688, 0.824			Range 0.750, 0.845	

LT = log transformed covariates. The CRP and ESR were skewed distributions and log transformed for model fitting.

\*The statistically significant predictors were: Age, Jaw claudication, Ischemic vision loss, Platelets, and CRP in the non-log transformed model. In the log transformed model InCRP replaced the CRP and InESR replaced the ESR

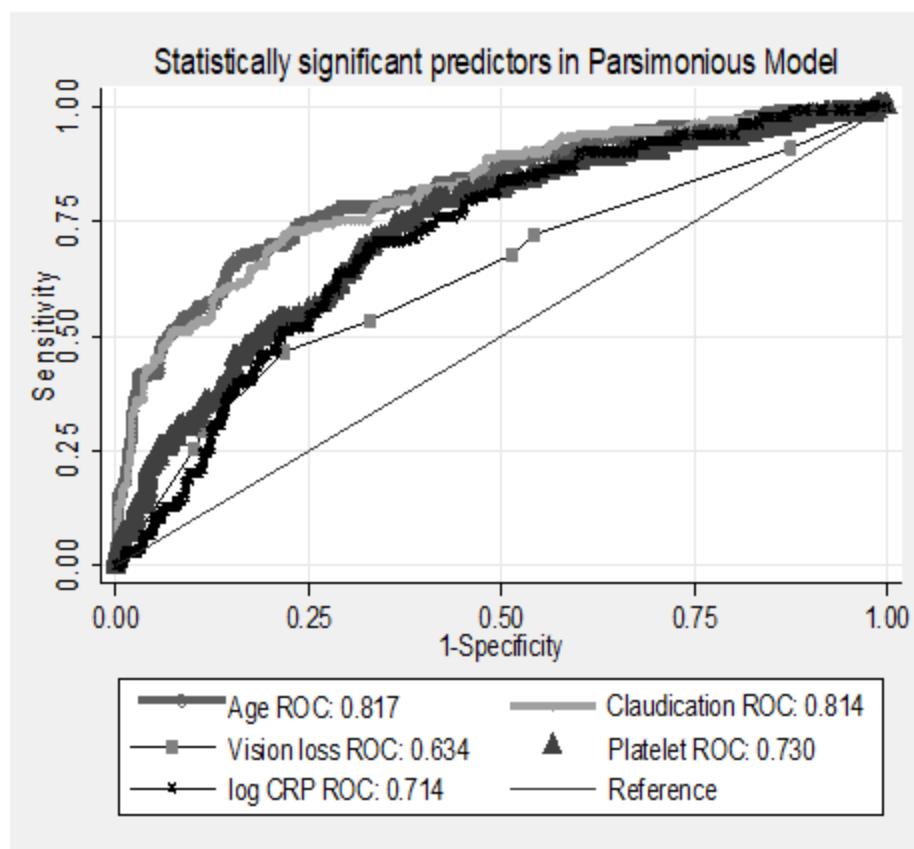
The Brier score has both a calibration and discrimination (sharpness) component. A statistically significant value of Spiegelhalter's z-test (evaluated at  $\alpha=0.05$ ), indicates poor calibration. (1)

ACR= American College of Rheumatology non-histologic criteria. None of the patients with positive temporal artery biopsy were under age 50 years old and the age category was omitted on logistic regression.

**Table S2** Model Performance for each prediction model at its optimal cutpoint (Liu method)

	Full Model	Parsimonious Model	ACR Model
<b>ACR Model Optimal Cutpoint 0.294</b>			
Sensitivity	0.68	0.65	0.47
Specificity	0.81	0.82	0.78
False Negative Rate	0.32	0.35	0.53
Misclassification rate	0.22	0.23	0.30
<b>Parsimonious Model Optimal Cutpoint 0.265</b>			
Sensitivity	0.70	0.73	0.54
Specificity	0.79	0.78	0.67
False Negative Rate	0.21	0.28	0.46
Misclassification rate	.23	0.23	0.35
<b>Full Model Optimal Cutpoint 0.242</b>			
Sensitivity	0.74	0.73	0.67
Specificity	0.77	0.77	0.49
False Negative Rate	0.25	0.27	0.32
Misclassification rate	0.24	0.24	0.46

**Figure S1**



**Table S3 Comparison of Prediction Rule Studies in the Literature**

Author (Year)	N (positive biopsy events)	Statistically Significant Predictors	Odds Ratio
Gabriel (1995)	525 (172)	All claudication Synovitis TA Abn Px Highest ESR	4.55 0.31 2.55 1.01
Rodriguez-Valverde, (1997)	227 (90)	New headache TA Abn Px Jaw claudication Raised liver enzymes < 70 years old at disease onset	13.6 4.2 4.8 2.9 0.11
Rieck (2011)	82 (22)	Jaw claudication Weight loss	4.50 3.76
Gonzalez-Lopez (2013)	335 (81)	Jaw claudication New headache TA Abn Px Pain and stiffness in neck and shoulders Unintentional weight loss Age Biopsy Length ESR	4.6 4.4 2.8 2.3 1.33 1.085 1.079 1.042
De Lott (2015)	239 for logistic regression (?90)	Age Platelets	1.06 1.01
Grossman (2016)	224 (57) [included 25 biopsy negative GCA]	New headache Jaw claudication ESR Platelet count	6.0 4.5 1.5 1.74
Present study (2017)	530 (133)	Jaw Claudication Vision Loss Platelet LogCRP Age	4.0 2.7 1.005 1.35 1.04

GCA = giant cell arteritis

TA Abn Px = temporal artery abnormality on physical exam

## COMMENTS ON THE GCA CALCULATOR

The calculator is meant to objectify decision-making when discussing management options with patients with suspected GCA. The calculator only predicts **biopsy-proven** GCA.

We encourage you to interpret the probability score with respect to the cutpoints. For scores lower than 15%, ask your patient, "Given GCA is a potentially blinding disease, and there is a \_% chance you might have GCA, do you want to proceed with TABx and steroids, or do you want to continue observation?"

None of the 133 biopsy proven GCA patients had a probability score below 2.7%

7.5% of the patients with biopsy proven GCA had a normal platelet, and normal ESR and normal CRP level pre-steroid; the average probability score on this seronegative group was 11%.

Female gender and diplopia were NOT helpful predictors of GCA in this study. You will see the risk score goes down for females and patients with diplopia. More females had GCA than males, but more females also had negative biopsy. In this series 35% with biopsy proven GCA had vision loss. Perhaps the high incidence of vision loss made binocular diplopia unlikely and that may be why the diplopia variable performs this way on the calculator.

**STATISTICAL PROGRAMS:** Stata 14.2, JMP Pro 13

### CODEBOOK FOR VARIABLES

VARIABLE NAME	MEANING		TYPE OF VARIABLE
Outcome variable			
tabxneg0	Temporal artery biopsy pathology result	Positive =1 Negative=0	binary
Predictor variables			
age	Age in years		continuous
female1	Gender	Female=1 Male = 0	binary
newheadache	Recent onset headache	1= new headache 0= no new headache	binary

taabnormality	Tender or pulseless temporal artery or scalp tenderness/nodule	1= present 0= absent	binary
Jawclaudication1	Jaw claudication + (tongue claudication)	1=claudication	binary
discrao	Permanent vision/field loss (disc edema, retinal artery occlusion, PION) In article = VL	1 = vision loss 0= no vision loss	binary
Diplopial	Diplopia symptoms	1= diplopia present	binary
platelet			
lncrp	Natural log CRP + constant	Ln (CRP divided by ULN) +3.69	continuous
lnesr	Natural log Westergren ESR + constant	ln(ESR) + 4.61	continuous
length	Biopsy length in cm	Posthoc analysis (not a predictor)	continuous
f1530pr1	Probability of positive biopsy, full model		continuous
pabicpr1	Probability of positive biopsy, parsimonious model		continuous
acrpobl	Probability of positive biopsy ACR model		continuous
esrcat	ESR>=50 == 1 for ACR		binary
agecat	Age >=50 ==1 for ACR		binary

code3	region		Categorical string
code4	Numeric region code		categorical
nlbloodtabxpos1	Biopsy positive but Platelets<400, ESR<50 and CRP <=1		binary

## METHOD SECTION

### KATTAN NOMOGRAM CONSTRUCTION

Skewed data was normalized by the logarithmic function and constants were added to provide positive results for the Kattan nomogram. For the ESR the constant 4.61 was added to the logESR, and for the CRP the constant 3.69 was added to the logCRP so that there were no negative values for the later Kattan nomogram. For ease of use, the logarithmic transforms were converted back to ESR and CRP values in the nomogram scales.

## RESULTS SECTION

TABLE 1

```
table1, by( tabxneg0) vars( age contn \ female1 bin \ newheadache bin
\ taabnormality bin \ jawclaudication1 bin \platelet contn \ origesr
conts \ crp conts \ discrao bin \ Diplopial bin \ length contn)
saving(Table3GCA.xls)

UNIVARIATE LOGISTIC REGRESSION
foreach x of varlist age female1 newheadache taabnormality jawclaudication1 platelet
origesr crp discrao Diplopial {
logistic tabxneg0 `x'
}
*
"SERONEGATIVE" GCA
generate nlbloodtabxpos1 = 0
replace nlbloodtabxpos1 =1 if origesr<50 & crp <=1 & platelet<400 &tabxneg0==1
tabstat age female1 newheadache newheadache taabnormality jawclaudication1 discrao Diplopial
length platelet origesr crp if tabxneg0==1, statistics( mean sd count ) by(nlbloodtabxpos1)
Summary statistics: mean, sd, N
by categories of: nlbloodtabxpos1
nlbloodtabxpos1 |    age    female1   newhea~e  newhea~e  taabno~y  jawcla~1   discrao  Diplop~1
                  length  platelet  origesr      crp
-----+-----+
-----+
          0 |  77.06748  .6585366  .7642276  .7642276  .5121951  .4878049  .3577236
          .0650407  1.915306           395   58.39443  13.77265
```

		7.778464	.47614	.4262167	.4262167	.5018956	.5018956	.4812906	.2476062	
.580322	140.3226	29.93112	14.76352							
123	98	123	123	123	123	123	123	123	123	
278.8	1	73.22	.7	.7	.7	.1	.3	.3	0	2.0125
	22.1	.3998333								
0	.710005	36.57807	13.2619	.2253101						
10	10	10	10	10	10	10	10	10	8	10
Total		76.7782	.6616541	.7593985	.7593985	.481203	.4736842	.3533835	.0601504	
1.922642	386.2632	55.66553	12.76718							
.5876966	138.6938	30.53335	14.62812							
106	133	133	133	133	133	133	133	133	133	

```
foreach x of varlist age female1 newheadache taabnormality jawclaudication1 discrao
Diplopial platelet origesr crp length {
ttest `x' if tabxneg0==1, by(nlbloodtabxpos1)
}
```

### SHORT TEMPORAL ARTERY BIOPSIES

tabstat length if length <1.5, statistics( count mean sd ) by(tabxneg0)

Summary for variables: length

by categories of: tabxneg0

tabxneg0		N	mean	sd
NEG		118	1.051271	.2626673
POS		23	1.030435	.242027
Total		141	1.047872	.2586968

ttesti 118 1.051271 .2626673 23 1.030435 .242027

Two-sample t test with equal variances

		Obs	Mean	Std. Err.	Std. Dev.	[95% Conf. Interval]
x		118	1.051271	.0241805	.2626673	1.003383 1.099159
y		23	1.030435	.0504661	.242027	.9257747 1.135095
combined		141	1.047872	.0217862	.2586968	1.0048 1.090945
diff			.020836	.0591505		-.0961151 .1377871

diff = mean(x) - mean(y) t = 0.3523

Ho: diff = 0 degrees of freedom = 139

Ha: diff < 0 Ha: diff != 0 Ha: diff > 0

Pr(T < t) = 0.6374 Pr(|T| > |t|) = 0.7252 Pr(T > t) = 0.3626

```

sum length, detail, if tabxneg0==0
      length
-----
Percentiles      Smallest
1%          .5          .1
5%          .8          .4
10%         1          .4      Obs             376
25%         1.3         .5      Sum of Wgt.     376
50%         1.8
                           Mean        1.848005
                           Largest     Std. Dev.   .6952717
75%         2.5         3.8
90%         2.5         4.2      Variance       .4834027
95%         2.6         4.3      Skewness       .2796604
99%         3.8         4.6      Kurtosis       3.321012
sum length, detail, if tabxneg0==1
      length
-----
Percentiles      Smallest
1%          .7          .6
5%          .8          .7
10%         1          .8      Obs             106
25%         1.5         .8      Sum of Wgt.     106
50%         2
                           Mean        1.922642
                           Largest     Std. Dev.   .5876966
75%         2.5         2.6
90%         2.5         2.7      Variance       .3453872
95%         2.6         2.8      Skewness       -.5285843
99%         2.8         3      Kurtosis       2.200076

```

#### **CHOICE OF FULL MODEL TO MINIMIZE AIC AND BIC**

Full model: nontransformed esr crp

Model	Obs	ll(null)	ll(model)	df	AIC	BIC
.	530	-298.5857	-229.3297	11	480.6595	527.6611

Full model: lnesr, crp

Model	Obs	ll(null)	ll(model)	df	AIC	BIC
.	530	-298.5857	-226.6329	11	475.2658	522.2674

Full model: esr, lncrp

Model	Obs	ll(null)	ll(model)	df	AIC	BIC
.	530	-298.5857	-222.7568	11	467.5137	514.5153

**Full model: lnesr, lncrp**

Model	Obs	ll(null)	ll(model)	df	AIC	BIC
.	1	530	-298.5857	-222.1149	11 466.2297	513.2314

In the non-transformed full model, the ESR was statistically significant ( $p=0.029$  with  $p_{MICE}=0.30$ ), but the CRP ( $p=0.762$  with  $p_{MICE}=0.651$ ) was not. However the full model with the log transformed CRP ( $p=.003$ ) and log transformed ESR ( $p=.273$ ) was chosen because it had lower AIC and BIC.

**NO SPECIFICATION ERROR (Stata "linktest")**

**FULL MODEL**

. logit tabxneg0 age female1 newheadache taabnormality jawclaudication1 platelet crp origesr discrao Diplopial, nolog	Logistic regression	Number of obs	=	530
		LR chi2(10)	=	138.51
		Prob > chi2	=	0.0000
		Pseudo R2	=	0.2319
Log likelihood = -229.32973				
<hr/>				
tabxneg0   Coef. Std. Err. z P> z  [95% Conf. Interval]				
<hr/>				
age   .0465098 .0128504 3.62 0.000 .0213234 .0716961				
female1   -.2353226 .2571522 -0.92 0.360 -.7393318 .2686865				
newheadache   .2044438 .2854474 0.72 0.474 -.3550228 .7639105				
taabnormality   .4003271 .2554037 1.57 0.117 -.1002551 .9009092				
jawclaudication1   1.335526 .2592266 5.15 0.000 .8274509 1.843601				
platelet   .0060248 .0010619 5.67 0.000 .0039435 .0081061				
crp   .0028098 .0092628 0.30 0.762 -.015345 .0209645				
origesr   .0089525 .0040991 2.18 0.029 .0009183 .0169866				
discrao   .9951831 .2753072 3.61 0.000 .4555591 1.534775				
Diplopial   -.0639864 .5051335 -0.13 0.899 -1.05403 .926057				
_cons   -7.855003 1.121877 -7.00 0.000 -10.05384 -5.656165				
<hr/>				
. linktest, nolog	Logistic regression	Number of obs	=	530
		LR chi2(2)	=	138.71
		Prob > chi2	=	0.0000
		Pseudo R2	=	0.2323
Log likelihood = -229.23054				
<hr/>				
tabxneg0   Coef. Std. Err. z P> z  [95% Conf. Interval]				
<hr/>				
_hat   1.046251 .1490225 7.02 0.000 .7541728 1.33833				
_hatsq   .0294055 .0662836 0.44 <b>0.657</b> -.1005079 .1593189				
_cons   -.0207729 .1574102 -0.13 0.895 -.3292911 .2877454				
<hr/>				

**FULL MODEL WITH LOG TRANSFORMED ESR CRP**

```

. logit tabxneg0 age female1 newheadache taabnormality jawclaudication1 platelet lncrp
lnesr discrao Diplopial, nolog
Logistic regression
Number of obs = 530
LR chi2(10) = 152.94
Prob > chi2 = 0.0000
Log likelihood = -222.11487
Pseudo R2 = 0.2561
-----
          tabxneg0 |   Coef.   Std. Err.      z     P>|z|      [95% Conf. Interval]
-----+
        age |   .0449871   .0131698     3.42    0.001     .0191748   .0707993
female1 |  -.1523935   .2610758    -0.58    0.559    -.6640926   .3593057
newheadache |   .2306857   .2895042     0.80    0.426    -.3367321   .7981035
taabnormality |   .3563223   .2609711     1.37    0.172    -.1551716   .8678162
jawclaudication1 |   1.296339   .2636562     4.92    0.000     .7795826   1.813096
platelet |   .0050691   .0010637     4.77    0.000     .0029844   .0071538
lncrp |   .2901259   .0980896     2.96    0.003     .0978738   .4823779
lnesr |   .1870691   .1705210     1.10    0.273    -.1471459   .5212841
discrao |   1.03084   .2806157     3.67    0.000     .4808433   1.580836
Diplopial |  -.229284   .5020145    -0.46    0.648    -1.213214   .7546464
_cons |  -9.96761   1.600932    -6.23    0.000    -13.10538   -6.829841
-----
. linktest, nolog
Logistic regression
Number of obs = 530
LR chi2(2) = 154.36
Prob > chi2 = 0.0000
Log likelihood = -221.40814
Pseudo R2 = 0.2585
-----
          tabxneg0 |   Coef.   Std. Err.      z     P>|z|      [95% Conf. Interval]
-----+
_hat |   1.124489   .1516177     7.42    0.000     .8273235   1.421654
_hat_sq |   .0723567   .0606377     1.19    0.233  -.0464911   .1912045
_cons |  -.0515301   .1562692    -0.33    0.742    -.3578121   .2547519
-----+

```

A SMALL ODDS RATIOS MAY BE SEEMINGLY UNIMPORTANT, BUT PLATELETS WERE A CONTINOUS COVARIATE with wide range. The odds ratio for platelets is **1.005** with  $\beta$  .0050691. Although a 1 unit change in platelets may be a small difference, and the odds ratio for platelets seemingly unimportant a 10 unit increase in platelets has effect  $e^{(\beta \text{platelets} * 10)} = e^{(.0050691 * 10)} = 1.05X$  increase in the odds ratio. A 50 unit increase in platelets results in a  $e^{(.0050691 * 50)} = 1.29X$  increase in the odds.

#### PARSIMONIOUS MODEL

```

logit tabxneg0 age jawclaudication1 platelet lncrp discrao, nolog
Logistic regression
Number of obs = 530
LR chi2(5) = 148.15
Prob > chi2 = 0.0000

```

```

Log likelihood = -224.5108 Pseudo R2 = 0.2481
-----
          tabxneg0 |   Coef.   Std. Err.      z     P>|z| [95% Conf. Interval]
-----+
           age |  .0438666  .0129774      3.38    0.001  .0184314  .0693019
jawclaudication1 |  1.402289  .2527095      5.55    0.000  .9069877  1.897591
      platelet |  .0050816  .0010462  4.86    0.000  .003031  .0071321
      lncrp |  .3581129  .0800815  4.47    0.000  .2011559  .5150698
      discrao |  .9248225  .2686439  3.44    0.001  .3982902  1.451355
      _cons | -8.49572  1.127081   -7.54   0.000 -10.70476 -6.286681
-----
. linktest, nolog
Logistic regression
Number of obs = 530
LR chi2(2) = 148.68
Prob > chi2 = 0.0000
Pseudo R2 = 0.2490
-----
          tabxneg0 |   Coef.   Std. Err.      z     P>|z| [95% Conf. Interval]
-----+
      _hat |  1.073897  .1472921      7.29    0.000  .7852101  1.362585
      _hatsq |  .045486  .0627132  0.73  **0.468** -0.0774296  .1684017
      _cons | -.0343638  .1566749   -0.22    0.826  -.341441  .2727135
-----
ACR MODEL (5 subjects less than 50 years of age, all had negative TABx)
logit tabxneg0 agecat esrcat newheadache taabnormality, nolog
note: agecat != 1 predicts failure perfectly
      agecat dropped and 5 obs not used
Logistic regression
Number of obs = 525
LR chi2(3) = 25.77
Prob > chi2 = 0.0000
Pseudo R2 = 0.0434
-----
          tabxneg0 |   Coef.   Std. Err.      z     P>|z| [95% Conf. Interval]
-----+
      agecat |          0 (omitted)
      esrcat |  .8195679  .2070725  3.96    0.000  .4137132  1.225423
      newheadache |  .0979034  .2451079  0.40    0.690  -.3824992  .5783061
      taabnormality |  .5792912  .2143856  2.70    0.007  .1591032  .9994792
      _cons | -1.738102  .2369044   -7.34   0.000 -2.202426 -1.273778
linktest, nolog
Logistic regression
Number of obs = 525
LR chi2(2) = 27.32
Prob > chi2 = 0.0000
Pseudo R2 = 0.0460
-----
          tabxneg0 |   Coef.   Std. Err.      z     P>|z| [95% Conf. Interval]
-----+
      _hat |  2.036413  .8596832  2.37    0.018  .3514652  3.721361
      _hatsq |  .5234275  .421919   1.24    0.215 -.3035185  1.350374

```

\_cons | .3707679 .3752286 0.99 0.323 -.3646667 1.106203

**NO MULTICOLLINEARITY**

collin age female1 newheadache taabnormality jawclaudication1 platelet crp origesr  
discrao Diplopia1

(obs=530)

Collinearity Diagnostics

Variable	SQRT		R-	
	VIF	VIF	Tolerance	Squared
age	1.02	1.01	0.9803	0.0197
female1	1.05	1.02	0.9549	0.0451
newheadache	1.10	1.05	0.9060	0.0940
taabnormality	1.15	1.07	0.8718	0.1282
jawclaudication1	1.09	1.05	0.9153	0.0847
platelet	1.19	1.09	0.8424	0.1576
crp	1.45	1.21	0.6880	0.3120
origesr	1.39	1.18	0.7220	0.2780
discrao	1.09	1.04	0.9168	0.0832
Diplopia1	1.02	1.01	0.9839	0.0161

Mean VIF 1.15

Eigenval	Cond	
	Index	
1	6.4979	1.0000
2	0.9639	2.5965
3	0.9080	2.6751
4	0.7584	2.9271
5	0.6696	3.1151
6	0.4741	3.7021
7	0.2660	4.9424
8	0.2211	5.4215
9	0.1496	6.5901
10	0.0831	8.8408
11	0.0082	28.0948

Condition Number 28.0948

Eigenvalues & Cond Index computed from scaled raw sscp (w/ intercept)

Det(correlation matrix) 0.4855

. collin age female1 newheadache taabnormality jawclaudication1 platelet lncrp lnesr  
discrao Diplopia1

(obs=530)

Collinearity Diagnostics

Variable	SQRT		R-	
	VIF	VIF	Tolerance	Squared
age	1.03	1.01	0.9740	0.0260
female1	1.03	1.02	0.9686	0.0314

newheadache	1.10	1.05	0.9080	0.0920
taabnormality	1.16	1.08	0.8653	0.1347
jawclaudication1	1.09	1.04	0.9180	0.0820
platelet	1.21	1.10	0.8231	0.1769
lncrp	1.68	1.30	0.5949	0.4051
lnesr	1.49	1.22	0.6724	0.3276
discrao	1.09	1.04	0.9199	0.0801
Diplopia1	1.02	1.01	0.9780	0.0220

-----  
Mean VIF            1.19

Eigenval	Cond	Index
1	7.2943	1.0000
2	0.9529	2.7668
3	0.9001	2.8468
4	0.6771	3.2822
5	0.4764	3.9131
6	0.2867	5.0440
7	0.2299	5.6334
8	0.0937	8.8239
9	0.0700	10.2054
10	0.0131	23.5941
11	0.0059	35.0522

-----  
Condition Number        35.0522

Eigenvalues & Cond Index computed from scaled raw sscp (w/ intercept)

Det(correlation matrix)    0.4313

#### **PARSIMONIOUS MODEL**

collin age jawclaudication1 platelet lncrp discrao  
(obs=530)

#### Collinearity Diagnostics

Variable	SQRT	R-		
	VIF	VIF	Tolerance	Squared
age	1.01	1.01	0.9869	0.0131
jawclaudication1	1.02	1.01	0.9796	0.0204
platelet	1.18	1.09	0.8475	0.1525
lncrp	1.16	1.08	0.8621	0.1379
discrao	1.03	1.01	0.9716	0.0284

-----  
Mean VIF            1.08

Eigenval	Cond	Index
1	4.4175	1.0000
2	0.7825	2.3760
3	0.6201	2.6690
4	0.0930	6.8934
5	0.0784	7.5068

```

6      0.0085      22.7935
-----
Condition Number      22.7935
Eigenvalues & Cond Index computed from scaled raw sscp (w/ intercept)
Det(correlation matrix) 0.8271
ACR MODEL
collin agecat esrcat newheadache taabnormality
(obs=530)
Collinearity Diagnostics
          SQRT                  R-
Variable    VIF VIF    Tolerance   Squared
-----
agecat      1.00  1.00  0.9955  0.0045
esrcat      1.01  1.01  0.9854  0.0146
newheadache 1.06  1.03  0.9432  0.0568
taabnormality 1.06  1.03  0.9453  0.0547
-----
Mean VIF      1.03
          Cond
Eigenval      Index
-----
1      3.7179  1.0000
2      0.5965  2.4965
3      0.5034  2.7178
4      0.1775  4.5764
5      0.0046  28.3254
-----
Condition Number      28.3254
Eigenvalues & Cond Index computed from scaled raw sscp (w/ intercept)
Det(correlation matrix) 0.9365
There was NO specification error in our logistic regression models.
There was no multicollinearity; the mean variance inflation factor
(VIF) was 1.19, all of the predictor VIF were in the range of one
with maximum 1.68 , and lowest observed tolerance 0.595 associated
with the log CRP.

```

### **MULTIPLE IMPUTATION of non-log transformed data**

```

mi set wide
mi register imputed platelet crp origesr newheadache taabnormality discrao Diplopial
jawclaudication1
.
mi register regular tabxneg0 age female1
mi impute chained (regress) platelet (regress) crp (regress) origesr (logit)
newheadache (logit)taabnormality (logit) discrao
> (logit) Diplopial (logit) jawclaudication1 = tabxneg0 age female1, add(250)
rseed(4409)
note: variables discrao Diplopial contain no soft missing (.) values; imputing nothing
Conditional models:

```

```

newheadache: logit newheadache i.discrao i.Diplopial i.jawclaudication1
i.taabnormality platelet crp origesr tabxneg0
    age female1
jawclaudicat~1: logit jawclaudication1 i.discrao i.Diplopial i.newheadache
i.taabnormality platelet crp origesr tabxneg0
    age female1
taabnormality: logit taabnormality i.discrao i.Diplopial i.newheadache
i.jawclaudication1 platelet crp origesr tabxneg0
    age female1
platelet: regress platelet i.discrao i.Diplopial i.newheadache
i.jawclaudication1 i.taabnormality crp origesr
    tabxneg0 age female1
crp: regress crp i.discrao i.Diplopial i.newheadache i.jawclaudication1
i.taabnormality platelet origesr
    tabxneg0 age female1
origesr: regress origesr i.discrao i.Diplopial i.newheadache i.jawclaudication1
i.taabnormality platelet crp
    tabxneg0 age female1
Performing chained iterations ...
Multivariate imputation                               Imputations = 250
Chained equations                                     added = 250
Imputed: m=1 through m=250                           updated = 0
Initialization: monotone                            Iterations = 2500
                                                       burn-in = 10
platelet: linear regression
crp: linear regression
origesr: linear regression
newheadache: logistic regression
taabnormality: logistic regression
discrao: logistic regression
Diplopial: logistic regression
jawclaudicat~1: logistic regression
-----
|                                Observations per m
|-----
Variable |   Complete   Incomplete   Imputed |   Total
|-----+-----+-----+-----+
platelet |      561       127       127 |      688
crp      |      530       158       158 |      688
origesr |      530       158       158 |      688
newheadache |     686        2        2 |      688
taabnormality |    679         9         9 |      688
discrao |     688        0        0 |      688
Diplopial |     688        0        0 |      688
jawclaudicat~1 |    683         5         5 |      688
|-----+
(complete + incomplete = total; imputed is the minimum across m
of the number of filled-in observations.)
.mi misstable patterns platelet crp origesr newheadache taabnormality
jawclaudication1 discrao Diplopial

```

Missing-value patterns  
(1 means complete)

Percent	Pattern					
	1	2	3	4	5	6
77%	1	1	1	1	1	1
17	1	1	1	0	0	0
3	1	1	1	1	0	0
<1	1	1	0	1	0	0
<1	1	0	1	0	0	0
<1	1	1	0	0	0	0
<1	0	0	0	0	0	0
<1	0	1	1	1	0	0
<1	1	0	1	1	0	0
100%						

Variables are (1) newheadache (2) jawclaudication1 (3) taabnormality (4) platelet  
(5) crp (6) origesr

mi estimate: logit tabxneg0 age female1 newheadache taabnormality jawclaudication1  
platelet crp origesr discrao Diplopial

Multiple-imputation estimates

Imputations	=	<b>250</b>
Number of obs	=	688
Average RVI	=	0.0912
Largest FMI	=	0.2232

DF adjustment: Large sample

DF:	min	=	5,013.54
	avg	=	204,747.63
	max	=	632,370.74

Model F test: Equal FMI

F( 10, 313092.7)	=	11.46
------------------	---	-------

Within VCE type: OIM

Prob > F	=	0.0000
----------	---	--------

tabxneg0 | Coef. Std. Err. t P>|t| [95% Conf. Interval]

age   .0522735 .0113708 4.60 0.000 .029987 .0745599
female1   .0268705 .2319645 0.12 0.908 -.4277741 .4815152
newheadache   .0575915 .2500353 0.23 0.818 -.4324713 .5476543
taabnormality   .4809197 .2290257 2.10 0.036 .0320287 .9298107
jawclaudication1   1.39079 .2254004 6.17 0.000 .949009 1.83257
platelet   .0058186 .0010184 5.71 0.000 .0038221 .007815
crp   .0041638 .0092029 0.45 0.651 -.0138769 .0222044
origesr   .0088631 .0040743 2.18 0.030 .0008757 .0168505
discrao   .9189811 .2453546 3.75 0.000 .4380926 1.39987
Diplopial   .0395116 .4506416 0.09 0.930 -.8437314 .9227546
_cons   -8.28233 1.007949 -8.22 0.000 -10.25789 -6.306771

logit tabxneg0 age female1 newheadache taabnormality jawclaudication1 platelet crp  
origesr discrao Diplopial

Iteration 0: log likelihood = -298.58568

Iteration 1: log likelihood = -233.04312

Iteration 2: log likelihood = -229.34022

```

Iteration 3:  log likelihood = -229.32973
Iteration 4:  log likelihood = -229.32973
Logistic regression                                         Number of obs      =      530
                                                               LR chi2(10)       =     138.51
                                                               Prob > chi2      =     0.0000
                                                               Pseudo R2        =     0.2319
Log likelihood = -229.32973
-----
```

tabxneg0	Coef.	Std. Err.	z	P> z	[95% Conf. Interval]
age   .0465098	.0128504		3.62	0.000	.0213234 .0716961
female1   -.2353227	.2571522	-0.92	0.360	-.7393318	.2686865
newheadache   .2044438	.2854474		0.72	0.474	-.3550228 .7639105
taabnormality   .4003271	.2554037		1.57	0.117	-.100255 .9009092
jawclaudication1   1.335526	.2592266		5.15	0.000	.8274508 1.843601
platelet   .0060248	.0010619	5.67	0.000	.0039435	.0081061
crp   .0028098	.0092628		0.30	0.762	-.015345 .0209645
origesr   .0089525	.0040991	2.18	0.029	.0009183	.0169866
discrao   .9951831	.2753072	3.61	0.000	.455591	1.534775
Diplopial   -.0639863	.5051335	-0.13	0.899	-1.05403	.9260571
_cons   -7.855003	1.121877		-7.00	0.000	-10.05384 -5.656165

(MULTIPLE IMPUTATION CHAINED EQUATIONS OF LOG TRANSFORMED DATA IS usually NOT RECOMMENDED BY , BUT INCLUDED IN APPENDIX FOR COMPLETENESS SAKE)

*MICE of log transformed data is not usually recommended, but performed on the log transformed full model to examine for trends. There was no change in the significance of the predictor variables in the log transformed full model; multiple imputation estimate for the temporal artery abnormality predictor was not statistically significant ( $p_{original} = .172$ ,  $p_{MICE}=.065$ ) .*

```

mi set wide
mi register imputed platelet lncrp lnesr newheadache taabnormality discrao Diplopial
jawclaudication1
mi register regular tabxneg0 age female1
mi impute chained (regress) platelet (regress) lncrp (regress) lnesr (logit)
newheadache (logit)taabnormality (logit) discrao (logit) Diplopial (logit)
jawclaudication1 = tabxneg0 age female1, add(250) rseed(4409)
note: variables discrao Diplopial contain no soft missing (.) values; imputing nothing
Conditional models:
    newheadache: logit newheadache i.discrao i.Diplopial i.jawclaudication1
i.taabnormality lnesr platelet lncrp tabxneg0
        age female1
    jawclaudicat~1: logit jawclaudication1 i.discrao i.Diplopial i.newheadache
i.taabnormality lnesr platelet lncrp tabxneg0
        age female1
    taabnormality: logit taabnormality i.discrao i.Diplopial i.newheadache
i.jawclaudication1 lnesr platelet lncrp tabxneg0

```

age female1  
 lnesr: regress lnesr i.discrao i.Diplopial i.newheadache  
 i.jawclaudication1 i.taabnormality platelet lncrp  
 tabxneg0 age female1  
 platelet: regress platelet i.discrao i.Diplopial i.newheadache  
 i.jawclaudication1 i.taabnormality lnesr lncrp  
 tabxneg0 age female1  
 lncrp: regress lncrp i.discrao i.Diplopial i.newheadache  
 i.jawclaudication1 i.taabnormality lnesr platelet  
 tabxneg0 age female1

Performing chained iterations ...

Multivariate imputation	Imputations =	250
Chained equations	added =	250
Imputed: m=1 through m=100	updated =	0
Initialization: monotone	Iterations =	2500
	burn-in =	10

platelet: linear regression  
 lncrp: linear regression  
 lnesr: linear regression  
 newheadache: logistic regression  
 taabnormality: logistic regression  
 discrao: logistic regression  
 Diplopial: logistic regression  
 jawclaudicat~1: logistic regression

---

Variable	Observations per m			Total
	Complete	Incomplete	Imputed	
platelet	561	127	127	688
lncrp	559	129	129	688
lnesr	595	93	93	688
newheadache	686	2	2	688
taabnormality	679	9	9	688
discrao	688	0	0	688
Diplopial	688	0	0	688
jawclaudicat~1	683	5	5	688

---

(complete + incomplete = total; imputed is the minimum across m of the number of filled-in observations.)

mi estimate: logit tabxneg0 age female1 newheadache taabnormality jawclaudication1  
 platelet lncrp lnesr discrao Diplopial

Multiple-imputation estimates

Imputations	=	250
Number of obs	=	688
Average RVI	=	0.0895
Largest FMI	=	0.2416

DF adjustment: Large sample

DF:	min	=	4,279.96
	avg	=	176,897.51
	max	=	765,589.65

Model F test: Equal FMI  $F(10, 328159.3) = 12.29$   
 Within VCE type: OIM  $\text{Prob} > F = 0.0000$

tabxneg0	Coef.	Std. Err.	t	P> t	[95% Conf. Interval]
age	.0514132	.0117052	4.39	0.000	.0284713 .074355
female1	.1148088	.2367164	0.49	0.628	-.3491502 .5787677
newheadache	.0739943	.2562119	0.29	0.773	-.4281759 .5761644
taabnormality	.4321508	.234619	1.84	0.065	-.0277023 .892004
jawclaudication1	1.344632	.2293685	5.86	0.000	.8950734 1.79419
platelet	.0047802	.0010181	4.70	0.000	.0027844 .0067761
lncrp	.3254603	.095798	3.40	0.001	.1376466 .513274
lnesr	.1723798	.1546375	1.11	0.265	-.1307628 .4755223
discrao	.9623211	.2515404	3.83	0.000	.4693082 1.455334
Diplopial	-.0911896	.4530914	-0.20	0.840	-.9792338 .7968546
_cons	-10.46421	1.454274	-7.20	0.000	-13.31468 -7.613738

logit tabxneg0 age female1 newheadache taabnormality jawclaudication1 platelet lncrp  
 lnesr discrao Diplopial

Iteration 0: log likelihood = -298.58568  
 Iteration 1: log likelihood = -227.04763  
 Iteration 2: log likelihood = -222.16089  
 Iteration 3: log likelihood = -222.11488  
 Iteration 4: log likelihood = -222.11486

Logistic regression  
 Number of obs = 530  
 LR chi2(10) = 152.94  
 Prob > chi2 = 0.0000  
 Pseudo R2 = 0.2561

tabxneg0	Coef.	Std. Err.	z	P> z	[95% Conf. Interval]
age	.0449871	.0131698	3.42	0.001	.0191748 .0707993
female1	-.1523934	.2610758	-0.58	0.559	-.6640926 .3593057
newheadache	.2306857	.2895042	0.80	0.426	-.3367321 .7981035
taabnormality	.3563223	.2609711	1.37	0.172	-.1551716 .8678162
jawclaudication1	1.296339	.2636562	4.92	0.000	.7795826 1.813096
platelet	.0050691	.0010637	4.77	0.000	.0029844 .0071538
lncrp	.2901259	.0980896	2.96	0.003	.0978738 .482378
lnesr	.1870691	.170521	1.10	0.273	-.1471459 .5212841
discrao	1.03084	.2806157	3.67	0.000	.4808433 1.580836
Diplopial	-.229284	.5020145	-0.46	0.648	-1.213214 .7546464
_cons	-9.96761	1.600932	-6.23	0.000	-13.10538 -6.82984

Multiple imputation analysis of the log transformed model: n=688,  
 250 imputations

Predictor	$\beta$	Odds ratio (O.R.)	p value	95% confidence interval (O.R.)	$\beta$ Multiple imputation estimates	p Multiple imputation estimates

Age	0.045	1.046	0.001	1.019, 1.074	0.052	<.001
Gender (female)	-0.152	0.859	0.559	0.514, 1.432	0.114	0.628
New headache	0.231	1.259	0.426	0.714, 2.221	0.078	0.761
Jaw claudication	1.296	3.656	<0.001	2.181, 6.129	1.342	<0.001
TA abnormality	0.356	1.428	0.172	0.856, 2.382	0.430	0.067
Vision Loss	1.031	2.803	<0.001	1.617, 4.859	0.967	<0.001
Diplopia	-0.229	0.795	0.648	0.297, 2.127	-0.090	0.842
lnESR	0.187	1.206	0.273	0.863, 1.684	0.167	0.280
lnCRP	0.290	1.337	0.003	1.103, 1.620	0.331	0.001
Platelets	0.005	1.005	<0.001	1.003, 1.007	0.005	<0.001
Constant	-9.967				-10.450	

TA abnormality: temporal artery abnormality on clinical exam

lnESR: natural logarithm of ESR

lnCRP: natural logarithm CRP

#### BOOTSTRAP SAMPLING (3000 random samples) OF PARSIMONIOUS MODEL

```
logit tabxneg0 age jawclaudication1 platelet lncrp discrao, vce (bootstrap, reps(3000)
bca seed(1))
```

```
estat bootstrap
```

#### INTERNAL VALIDATION PARSIMONIOUS MODEL

```
cvauroc tabxneg0 age jawclaudication1 platelet lncrp discrao, kfold(10) seed(3)
```

```
1-fold.....
```

```
2-fold.....
```

```
3-fold.....
```

```
4-fold.....
```

```
5-fold.....
```

```
6-fold.....
```

```
7-fold.....
```

```
8-fold.....
```

```
9-fold.....
```

```
10-fold.....
```

```
(530 real changes made)
```

	ROC	-Asymptotic Normal--	
Obs	Area	Std. Err.	[95% Conf. Interval]

530	0.8102	0.0224	0.76634	0.85397
-----	--------	--------	---------	---------

#### INTERNAL VALIDATION FULL MODEL

```

cvAUROC tabxneg0 female1 age newheadache taabnormality
jawclaudication1 platelet lnesr lncrp discrao, kfold(10) seed(1)
cvAUROC tabxneg0 female1 age newheadache taabnormality jawclaudication1 platelet lnesr
lncrp discrao Diplopia1, kfold(10) seed
> (2)
1-fold.....
2-fold.....
3-fold.....
4-fold.....
5-fold.....
6-fold.....
7-fold.....
8-fold.....
9-fold.....
10-fold.....
(530 real changes made)

          ROC           -Asymptotic Normal--
      Obs     Area     Std. Err.   [95% Conf. Interval]
-----
      530    0.8030    0.0233    0.75726    0.84875

EXTERNAL VALIDATION further examples
Full model; Combined centers below
Full model: Toronto below
Full model: Swiss below
Parsimonious model: Combined centers, below

```

## **External Validation: Comparison with the development data of the distribution of important variables (demographics, predictors and outcome with ANOVA.**

```

by code3, sort : tabstat Age female1 New_Headache TA_tender_noPulse Jaw_claudication
CRP Platelets Vision_Loss ESR Diplopia length, statistics( mean sd count )
Combined
  stats |      Age    female1  New_He~e  TA~Pulse  Jaw_cl~n        CRP  Platel~s
Vision~s          ESR  Diplopia  length
-----+
-----+
  mean |  73.04762  .5873016  .7619048  .4761905  .3492063  5.18452  381.5079
.5079365  8.335137  .047619  1.973684
  sd |  10.24133  .4962739  .4293388  .5034444  .4805484  1.518077  139.7956
.5039526  .7852612  .2146694  .1147079
  N |       63      63      63      63      63      63      63      63      63
19
-----+
-----+
Kingston
  stats |      Age    female1  New_He~e  TA~Pulse  Jaw_cl~n        CRP  Platel~s
Vision~s          ESR  Diplopia  length

```

```

-----+
-----+
      mean |  74.43605   .6686047      .75   .3953488   .2674419   5.307306  317.6105
.2325581  8.343498   .0930233   1.756977
      sd |  9.499245   .4720892   .434277   .490353   .4439171   1.657388  117.5335
.4236963  .868971   .2913131   .5978599
      N |      172       172       172       172       172       172       172       172
      172       172       172
-----+
-----+
Ottawa
      stats |      Age   female1  New_He~e  TA~Pulse  Jaw_cl~n        CRP  Platelets
Vision~s          ESR   Diplopia length
-----+
-----+
      mean |  74.36219   .7058824   .7815126   .2016807   .210084   3.541939  315.0168
.2436975  7.546964   .0252101           2.5
      sd |  9.659166   .4575717   .4149671   .4029517   .4090905   1.534551  119.9266
.4311275  1.430981   .1574255           0
      N |      119       119       119       119       119       119       119       119
      119       119       119
-----+
Rochester_MN
      stats |      Age   female1  New_He~e  TA~Pulse  Jaw_cl~n        CRP  Platelets
Vision~s          ESR   Diplopia length
-----+
-----+
      mean |  74.21154   .6923077   .6923077   .1730769   .1538462   4.553688  279.5192
.1153846  8.10026   .0576923   1.317308
      sd |  10.15437   .4660414   .4660414   .3820047   .3643213   1.310443  94.99731
.3226025  1.139259   .2354355   .4869759
      N |      52        52        52        52        52        52        52        52
      52
-----+
-----+
Toronto
      stats |      Age   female1  New_He~e  TA~Pulse  Jaw_cl~n        CRP  Platelets
Vision~s          ESR   Diplopia length
-----+
-----+
      mean |  72.66935   .6935484   .6774194   .5080645   .2822581   4.312429  280.2742
.0967742  7.8404   .0241935   1.607917
      sd |  10.47701   .4628898   .4693602   .5019631   .4519242   1.621985  107.6076
.2968494  1.034101   .154273   .7713623
      N |      124       124       124       124       124       124       124       124
      124       124       120
-----+
-----+
by tabxneg0, sort : tabstat Age female1 New_Headache TA_tender_noPulse
Jaw_claudication CRP Platelets Vision_Loss ESR Diplopia length, statistics( mean sd
count ) by(code3)

```

**Negative Biopsy Group**

Summary statistics: mean, sd, N

by categories of: code3

code3	Age	female1	New_He~e	TA~Pulse	Jaw_cl~n	CRP	Platel~s
Vision~s							
	ESR	Diplopia	length				
-----+-----							
Combined	67.83333	.5416667	.6666667		.375	.1666667	4.714693
323.7083	.4166667	8.215517	.0833333	1.916667			
	11.84563	.5089774	.4815434	.4945354	.3806935	1.613385	114.5727
.5036102	.8071274	.2823299	.2041241				
	24	24	24	24	24	24	24
24	6						
-----+-----							
Kingston	73.24286	.6928571	.7357143	.3785714	.1857143	5.104099	303.3857
.2142857	8.299645	.0928571	1.734286				
	9.733035	.4629656	.4425354	.4867726	.3902723	1.644053	108.3706
.4117993	.8720117	.2912743	.6051067				
	140	140	140	140	140	140	140
140	140	140	140				
-----+-----							
Ottawa	73.23478	.673913	.7934783	.1630435	.173913	3.134447	282.3261
.2282609	7.308337	.0217391	2.5				
	9.928618	.4713482	.4070274	.3714295	.3811116	1.322775	92.46765
.4220114	1.51303	.1466296	0				
	92	92	92	92	92	92	92
92							
-----+-----							
Rochester_MN	73.86842	.6842105	.7368421	.1578947	.0526316	4.329098	272.5
.1052632	7.989262	.0263158	1.307895				
	10.50912	.4710691	.4462583	.369537	.2262943	1.257937	96.22321
.3110117	1.210942	.1622214	.469837				
	38	38	38	38	38	38	38
38	38						
-----+-----							
Toronto	72.67961	.6990291	.6699029	.4563107	.2427184	4.056551	266.9417
.0679612	7.726733	.0194175	1.6085				
	10.95554	.460923	.4725473	.5005232	.4308227	1.628432	96.77626
.25291	1.071258	.1386618	.8055068				
	103	103	103	103	103	103	103
103	103	103	100				
-----+-----							
Total	72.82771	.6801008	.7279597	.3274559	.1838791	4.278151	287.3224
.1813602	7.886487	.0503778	1.848005				

		10.34455	.4670258	.4455721	.4698773	.3878744	1.703687	102.1441
.3858025	1.189804	.2189994	.6952717					
		397	397	397	397	397	397	397
397		397	397	376				

---

### Positive Biopsy Group

Summary statistics: mean, sd, N

by categories of: code3

	code3	Age	female1	New_He~e	TA~Pulse	Jaw_cl~n	CRP	Platel~s
Vision~s		ESR	Diplopia	length				
<hr/>								
	Combined	76.25641	.6153846	.8205128	.5384615	.4615385	5.473644	
417.0769	.5641026	8.408749	.025641		2			
		7.645838	.4928641	.3887764	.5050354	.5050354	1.399769	143.2898
.5023561	.7727799	.1601282		0				
		39	39	39	39	39	39	39
13								
<hr/>								
	Kingston	79.65625	.5625	.8125		.46875	.625	6.196337
379.8438	.3125	8.535355		.09375	1.85625			
		6.198904	.5040161	.3965578	.5070073	.4918694	1.424984	136.4897
.4709291	.8420514	.2961446		.563364				
		32	32	32	32	32	32	32
32								
<hr/>								
	Ottawa	78.2037	.8148148	.7407407	.3333333	.3333333	4.930431	426.4074
.2962963	8.360062	.037037		2.5				
		7.654585	.3958474	.4465761	.4803845	.4803845	1.404883	136.6799
.4653216	.626882	.1924501		0				
		27	27	27	27	27	27	27
27								
<hr/>								
	Rochester_MN	75.14286	.7142857	.5714286	.2142857	.4285714	5.16329	298.5714
.1428571	8.40154	.1428571	1.342857					
		9.428738	.4688072	.5135526	.4258153	.5135526	1.298283	92.28527
.3631365	.8857189	.3631365		.5487248				
		14	14	14	14	14	14	14
14								
<hr/>								
	Toronto	72.61905	.6666667	.7142857	.7619048	.4761905	5.567447	345.6667
.2380952	8.39791	.047619		1.605				
		7.933954	.4830459		.46291	.4364358	.5117663	.815159
.4364358	.5752944	.2182179		.5880163				134.3244

		21	21	21	21	21	21	21	21	21	21
20											
<hr/>											
	Total	76.7782	.6616541	.7593985	.481203	.4736842	5.519391	386.2632			
	.3533835	8.426856	.0601504	1.922642							
		7.824779	.4749357	.4290648	.5015356	.5011947	1.375809	138.6938			
	.4798278	.7398071	.2386642	.5876966							
		133	133	133	133	133	133	133	133	133	133
133		133	133	106							
<hr/>											

```

foreach x of varlist age female1 newheadache taabnormality
jawclaudication1 platelet origesr crp discrao Diplopial length {
oneway `x' code3, bonferroni
}
*
foreach x of varlist age female1 newheadache taabnormality jawclaudication1 platelet
origesr crp discrao Diplopial length {
2.
.oneway `x' code3, bonferroni
3.
.}

Analysis of Variance

```

Source	SS	df	MS	F	Prob > F
<hr/>					
Between groups	309.98749	4	77.4968724	0.79	0.5340
Within groups	51702.6105	525	98.4811628		
Total	52012.598	529	98.3224914		

Bartlett's test for equal variances: chi2(4) = 1.7275 Prob>chi2 = 0.786  
Comparison of AGE by code3  
(Bonferroni)

Row Mean -	Col Mean	Combined	Kingston	Ottawa	Roches..
<hr/>					
Kingston	1.38843				
	1.000				
Ottawa	1.31457	-.073861			
	1.000	1.000			
Roches..	1.16392	-.224508	-.150647		
	1.000	1.000	1.000		
Toronto	-.378264	-1.76669	-1.69283	-1.54218	
	1.000	1.000	1.000	1.000	

Source	SS	df	MS	F	Prob > F
<hr/>					

-----

Between groups	.66318155	4	.165795387	0.75	0.5560
Within groups	115.517951	525	.220034191		

-----

Total	116.181132	529	.219624068		
-------	------------	-----	------------	--	--

Bartlett's test for equal variances: chi2(4) = 0.6108 Prob>chi2 = 0.962

Comparison of **SEX(female)** by code3  
(Bonferroni)

Row Mean-|

Col Mean	Combined	Kingston	Ottawa	Roches..
Kingston   .081303				
1.000				
Ottawa   .118581		.037278		
1.000		1.000		
Roches..   .105006		.023703	-.013575	
1.000		1.000	1.000	
Toronto   .106247		.024944	-.012334	.001241
1.000		1.000	1.000	1.000

-----

Analysis of Variance

Source	SS	df	MS	F	Prob > F
Between groups	.847271494	4	.211817874	1.09	0.3615
Within groups	102.171596	525	.194612565		

-----

Total	103.018868	529	.194742661		
-------	------------	-----	------------	--	--

Bartlett's test for equal variances: chi2(4) = 2.3152 Prob>chi2 = 0.678

Comparison of **NEWHEADACHE** by code3  
(Bonferroni)

Row Mean-|

Col Mean	Combined	Kingston	Ottawa	Roches..
Kingston   -.011905				
1.000				
Ottawa   .019608		.031513		
1.000		1.000		
Roches..   -.069597		-.057692	-.089205	
1.000		1.000	1.000	
Toronto   -.084485		-.072581	-.104093	-.014888
1.000		1.000	0.665	1.000

-----

Analysis of Variance

Source	SS	df	MS	F	Prob > F
Between groups	8.56420742	4	2.14105185	9.82	0.0000

Within groups 114.424472 525 .217951375  
-----  
Total 122.988679 529 .232492777  
Bartlett's test for equal variances: chi2(4) = 11.1540 Prob>chi2 = 0.025

Comparison of **taabnormal~y** by code3  
(Bonferroni)

Row Mean-	Combined	Kingston	Ottawa	Roches..
Col Mean				
Kingston	-.080842			
	1.000			
Ottawa	-.27451	-.193668		
	0.002	0.005		
Roches..	-.303114	-.222272	-.028604	
	0.006	0.028	1.000	
Toronto	.031874	.112716	.306384	.334988
	1.000	0.409	0.000	0.000

Analysis of Variance

Source	SS	df	MS	F	Prob > F
Between groups	1.44865439	4	.362163596	1.91	0.1077
Within groups	99.6532324	525	.189815681		
Total	101.101887	529	.191118879		

Bartlett's test for equal variances: chi2(4) = 5.5112 Prob>chi2 = 0.239

Comparison of **jawclaudic~1** by code3  
(Bonferroni)

Row Mean-	Combined	Kingston	Ottawa	Roches..
Col Mean				
Kingston	-.081764			
	1.000			
Ottawa	-.139122	-.057358		
	0.409	1.000		
Roches..	-.19536	-.113596	-.056238	
	0.170	1.000	1.000	
Toronto	-.066948	.014816	.072174	.128412
	1.000	1.000	1.000	0.750

Analysis of Variance

Source	SS	df	MS	F	Prob > F
Between groups	490529.653	4	122632.413	9.00	0.0000
Within groups	7155506.27	525	13629.5358		

Total 7646035.92 529 14453.7541  
 Bartlett's test for equal variances: chi2(4) = 10.0430 Prob>chi2 = 0.040  
 Comparison of platelet by code3  
 (Bonferroni)

Row Mean -	Col Mean   Combined	Kingston	Ottawa	Roches..
Kingston   -63.8975				
0.002				
Ottawa   -66.4911 -2.59366				
0.003 1.000				
Roches..   -101.989 -38.0912 -35.4976				
0.000 0.397 0.680				
Toronto   -101.234 -37.3363 -34.7426 .754963				
0.000 0.069 0.208 1.000				

#### Analysis of Variance

Source	SS	df	MS	F	Prob > F
Between groups	56924.9564	4	14231.2391	15.10	0.0000
Within groups	494873.465	525	942.616123		
Total	551798.421	529	1043.0972		

Bartlett's test for equal variances: chi2(4) = 43.4064 Prob>chi2 = 0.000  
 Comparison of origestr by code3  
 (Bonferroni)

Row Mean -	Col Mean   Combined	Kingston	Ottawa	Roches..
Kingston   3.82676				
1.000				
Ottawa   -21.4612 -25.2879				
0.000 0.000				
Roches..   -5.79117 -9.61793 15.67				
1.000 0.483 0.022				
Toronto   -15.5368 -19.3636 5.92435 -9.74566				
0.011 0.000 1.000 0.552				

#### Analysis of Variance

Source	SS	df	MS	F	Prob > F
Between groups	11197.2915	4	2799.32288	17.29	0.0000
Within groups	85022.0439	525	161.94675		
Total	96219.3355	529	181.889103		

Bartlett's test for equal variances: chi2(4) = 174.8649 Prob>chi2 = 0.000

Comparison of **crp** by code3  
(Bonferroni)

Row Mean-	Combined	Kingston	Ottawa	Roches..
Col Mean				
Kingston	4.82371			
	0.103			
Ottawa	-6.68107	-11.5048		
	0.008	0.000		
Roches..	-4.13833	-8.96204	2.54274	
	0.832	0.000	1.000	
Toronto	-3.80594	-8.62966	2.87512	.332386
	0.538	0.000	0.789	1.000

Analysis of Variance

Source	SS	df	MS	F	Prob > F
Between groups	7.75825082	4	1.9395627	12.05	0.0000
Within groups	84.5228813	525	.160995964		
Total	92.2811321	529	.174444484		

Bartlett's test for equal variances: chi2(4) = 32.3925 Prob>chi2 = 0.000

Comparison of **DiscRAO** by code3  
(Bonferroni)

Row Mean-	Combined	Kingston	Ottawa	Roches..
Col Mean				
Kingston	-.275378			
	0.000			
Ottawa	-.264239	.011139		
	0.000	1.000		
Roches..	-.392552	-.117174	-.128313	
	0.000	0.656	0.549	
Toronto	-.411162	-.135784	-.146923	-.01861
	0.000	0.042	0.045	1.000

Analysis of Variance

Source	SS	df	MS	F	Prob > F
Between groups	.473271773	4	.118317943	2.38	0.0503
Within groups	26.0474829	525	.049614253		
Total	26.5207547	529	.050133752		

Bartlett's test for equal variances: chi2(4) = 79.0289 Prob>chi2 = 0.000

Comparison of **Diplopial** by code3  
(Bonferroni)

Row Mean-	Combined	Kingston	Ottawa	Roches..
Col Mean				
Kingston	.045404			
	1.000			
Ottawa	-.022409	-.067813		
	1.000	0.110		
Roches..	.010073	-.035331	.032482	
	1.000	1.000	1.000	
Toronto	-.023425	-.06883	-.001017	-.033499
	1.000	0.090	1.000	1.000

#### Analysis of Variance

Source	SS	df	MS	F	Prob > F
Between groups	73.7444151	4	18.4361038	60.96	0.0000
Within groups	144.25787	477	.302427401		
Total	218.002285	481	.453227204		

Bartlett's test for equal variances: chi2(3) = 61.1587 Prob>chi2 = 0.000

note: Bartlett's test performed on cells with positive variance:

1 multiple-observation cells not used

Comparison of **length** by **code3**  
(Bonferroni)

Row Mean-	Combined	Kingston	Ottawa	Roches..
Col Mean				
Kingston	-.216707			
	1.000			
Ottawa	.526316	.743023		
	0.001	0.000		
Roches..	-.656377	-.439669	-1.18269	
	0.000	0.000	0.000	
Toronto	-.365768	-.14906	-.892083	.290609
	0.073	0.231	0.000	0.016

**foreach x of varlist age female1 newheadache taabnormality**

**jawclaudication1 platelet origesr crp discrao Diplopia1 {**

**anova `x' code4**

**}**

**\***

Number of obs = 530 R-squared = 0.0060  
Root MSE = 9.92377 Adj R-squared = -0.0016

Source	Partial SS	df	MS	F	Prob>F
--------	------------	----	----	---	--------



Total		101.10189		529	.19111888		
Number of obs	=		530	R-squared	=	0.0642	
Root MSE	=		116.746	Adj R-squared	=	0.0570	
Source		Partial SS	df	MS	F	Prob>F	
-----+-----							
Plat		490529.65	4	122632.41	9.00	0.0000	
code4		490529.65	4	122632.41	9.00	0.0000	
Residual		7155506.3	525	13629.536			
-----+-----							
Total		7646035.9		529	14453.754		
Number of obs	=		530	R-squared	=	0.1032	
Root MSE	=		30.7021	Adj R-squared	=	0.0963	
Source		Partial SS	df	MS	F	Prob>F	
-----+-----							
ESR		56924.956	4	14231.239	15.10	0.0000	
code4		56924.956	4	14231.239	15.10	0.0000	
Residual		494873.46	525	942.61612			
-----+-----							
Total		551798.42		529	1043.0972		
Number of obs	=		530	R-squared	=	0.1164	
Root MSE	=		12.7258	Adj R-squared	=	0.1096	
Source		Partial SS	df	MS	F	Prob>F	
-----+-----							
CRP		11197.292	4	2799.3229	17.29	0.0000	
code4		11197.292	4	2799.3229	17.29	0.0000	
Residual		85022.044	525	161.94675			
-----+-----							
Total		96219.335		529	181.8891		
Number of obs	=		530	R-squared	=	0.0841	
Root MSE	=		.401243	Adj R-squared	=	0.0771	
Source		Partial SS	df	MS	F	Prob>F	
-----+-----							
Vision		7.7582508	4	1.9395627	12.05	0.0000	
code4		7.7582508	4	1.9395627	12.05	0.0000	
Residual		84.522881	525	.16099596			
-----+-----							
Total		92.281132		529	.17444448		
Number of obs	=		530	R-squared	=	0.0178	
Root MSE	=		.222743	Adj R-squared	=	0.0104	
Source		Partial SS	df	MS	F	Prob>F	
-----+-----							
Diplopia		.47327177	4	.11831794	2.38	0.0503	

code4	.47327177		4	.11831794		2.38	0.0503
Residual	26.047483		525	.04961425			
			-----+-----				
Total	26.520755		529	.05013375			

### STATA NOMOGRAM

```

logit tabxneg0 CRP_divide_ULN Platelets Vision_Loss Jaw_claudication Age
nomolog, title(Nomogram: Probability of Giant Cell Arteritis) varlblsize(2.8)
vli1(Vision_Loss,0,1,1,0) vli4(Jaw_claudication,0,1,1,0) vli5(Age,40,100,10,0)
vli6(Platelets,50,950,75,0) vli9(CRP_divide_by_ULN,0,8.11,1,1) prlabsize(3.5)
CALIBRATION PLOT are visual representation of H-L statistic.
drop prob
drop probgroup
logit tabxneg0 age female1 newheadache taabnormality jawclaudication1 platelet lncrp
lnesr discrao Diplopia1
predict prob
egen probgroup=cut(prob), at(0(0.1)1)
tab probgroup tabxneg0
preserve
collapse (mean) tabxneg0, by(probgp)
replace probgroup=probgroup + 0.05
scatter tabxneg0 probgroup || line probgroup probgroup, legend(order(1 "Observed" 2
"Predicted"))
1 newheadache taabnormality jawclaudication1 platelet lncrp lnesr discrao Diplopia1
lroc
predict xb1, xb
logit tabxneg0 age jawclaudication1 platelet lncrp discrao
lroc
predict xb2, xb
logistic tabxneg0 agecat esrcat newheadache taabnormality
lroc
predict xb3, xb
roccomp tabxneg0 xb1 xb2 xb3, graph summary
roccomp tabxneg0 xb1 xb2 xb3, graph summary
      ROC          -Asymptotic Normal--
      Obs   Area    Std. Err.      [95% Conf. Interval]
-----+
xb1      525  0.8174  0.0223  0.77377  0.86111
xb2      525  0.8142  0.0221  0.77080  0.85755
xb3      525  0.6339  0.0287  0.57753  0.69018
-----+
n=525 (in the ACR model 5 patients under the age of 50 years were
excluded)
ACR = American College of Rheumatology Classification Criteria
(non-histologic)
drop xb xb2 xb3 xb4 xb5

```

```

logistic tabxneg0 age
predict xb1, xb
logit tabxneg0 jawclaudication1
lroc
predict xb2, xb
logistic tabxneg0 discrao
lroc
predict xb3, xb
logistic tabxneg0 platelet
lroc
predict xb4, xb
logistic tabxneg0 lncrp
lroc
predict xb5, xb
roccomp tabxneg0 xb1 xb2 xb3 xb4 xb5, graph summary

```

**Bootstrap Sensitivity Analysis: Full model**

```

logistic tabxneg0 age female1 newheadache jawclaudication1 platelet lnesr lncrp discrao
Diplopial

```

Logistic regression	Number of obs	=	526
	LR chi2(9)	=	150.06
	Prob > chi2	=	0.0000
Log likelihood = -222.3942	Pseudo R2	=	0.2523

	Odds Ratio	Std. Err.	z	P> z	[95% Conf. Interval]
age	1.04388	.0136856	3.28	0.001	1.017398 1.071051
female1	.8514999	.2223184	-0.62	0.538	.5104393 1.420447
newheadache	1.365738	.3870027	1.10	0.271	.7837297 2.379953
jawclaudication1	3.979413	1.027158	5.35	0.000	2.399433 6.599778
platelet	1.005126	.0010682	4.81	0.000	1.003034 1.007221
lnesr	1.173762	.2118964	0.89	0.375	.8239766 1.672035
lncrp	1.362775	.1355529	3.11	0.002	1.12139 1.65612
discrao	2.622504	.7265376	3.48	0.001	1.5237 4.513701
Diplopial	.7751479	.3885454	-0.51	0.611	.2902132 2.07039
_cons	.0000968	.0001336	-6.70	0.000	6.48e-06 .0014462

```

logistic tabxneg0 age female1 newheadache jawclaudication1 platelet lnesr lncrp discrao
Diplopial, vce (bootstrap, reps(5000) bca seed(1))

```

Number of obs	=	526	
Replications	=	5,000	
Wald chi2(9)	=	90.23	
Prob > chi2	=	0.0000	
Log likelihood = -222.3942	Pseudo R2	=	0.2523

	Observed	Bootstrap	Normal-based			
	Odds Ratio	Std. Err.	z	P> z	[95% Conf. Interval]	
age	1.04388	.0135409	3.31	0.001	1.017675	1.07076
female1	.8514999	.2350384	-0.58	0.560	.495711	1.462651
newheadache	1.365738	.4293089	0.99	0.321	.7375625	2.528924
jawclaudication1	3.979413	1.074401	5.12	0.000	2.344246	6.755147
platelet	1.005126	.0010727	4.79	0.000	1.003025	1.00723
lnesr	1.173762	.1960574	0.96	0.337	.8460601	1.628393
lncrp	1.362775	.1277916	3.30	0.001	1.133977	1.637736

```

discrao | 2.622504 .7398522 3.42 0.001 1.508614 4.55884
Diplopial | .7751479 .4196151 -0.47 0.638 .2682866 2.239599
_cons | .0000968 .0001394 -6.42 0.000 5.77e-06 .0016263
-----
Bootstrap Sensitivity Analysis: Parsimonious model
logistic tabxneg0 age jawclaudication1 platelet lncrp discrao
Logistic regression
Number of obs = 526
LR chi2(5) = 147.61
Prob > chi2 = 0.0000
Log likelihood = -223.62022 Pseudo R2 = 0.2481
-----
tabxneg0 | Odds Ratio Std. Err. z P>|z| [95% Conf. Interval]
-----+
age | 1.044307 .0135879 3.33 0.001 1.018012 1.071281
jawclaudication1 | 4.080527 1.03364 5.55 0.000 2.483694 6.704009
platelet | 1.00508 .0010511 4.85 0.000 1.003022 1.007142
lncrp | 1.430682 .1142496 4.48 0.000 1.223401 1.673082
discrao | 2.487281 .6682512 3.39 0.001 1.469044 4.211288
_cons | .0002163 .0002437 -7.49 0.000 .0000238 .0019693
logistic tabxneg0 age jawclaudication1 platelet lncrp discrao, vce (bootstrap, reps(5000)
Logistic regression
Number of obs = 526
Replications = 5,000
Wald chi2(5) = 93.78
Prob > chi2 = 0.0000
Log likelihood = -223.62022 Pseudo R2 = 0.2481
-----
| Observed Bootstrap Normal-based
tabxneg0 | Odds Ratio Std. Err. z P>|z| [95% Conf. Interval]
-----+
age | 1.044307 .0130375 3.47 0.001 1.019064 1.070175
jawclaudication1 | 4.080527 1.051293 5.46 0.000 2.462724 6.761094
platelet | 1.00508 .001026 4.96 0.000 1.003071 1.007093
lncrp | 1.430682 .1129918 4.53 0.000 1.225511 1.670202
discrao | 2.487281 .6674884 3.40 0.001 1.469927 4.208757
_cons | .0002163 .0002465 -7.40 0.000 .0000232 .0020197
-----+

```

## NOMOGRAM CONSIDERATIONS

In this paper, there is only a nomogram for the parsimonious model.

A nomogram of the full model is available for readers interested in the graphic visualization of the importance of all 10 variables.

lnCRP	CRP	lnESR	ESR
0.00112	0.025	0.00483	0.01
1.35187	0.096508	3.34377	0.281892
2.70262	0.372552	6.68271	7.94633
4.05337	1.438168	10.0217	224.001
5.40412	5.551787		
6.75487	21.43167		
8.10562	82.73315		

$$\text{lnCRP} = \log \text{CRP} + 3.69 \quad \text{and} \quad \text{lnESR} = \log \text{ESR} + 4.61$$

The addition of constants was performed to avoid non-negative values, as the nomogram cannot otherwise be generated. The values are transformed back to ESR and CRP.