

STATA code

New Combined Variables

A.1 Levels of Exercise

```
generate atleast_mod =
```

```
.
```

```
replace atleast_mod = 1 if Vigorousrecreationalactivities == 2 & Moderaterecreationalactivities  
== 2 replace atleast_mod = 2 if Moderaterecreationalactivities == 1 &  
Vigorousrecreationalactivities == 2 replace atleast_mod = 3 if Vigorousrecreationalactivities ==  
1 & Moderaterecreationalactivities == 2 replace atleast_mod = 4 if  
Vigorousrecreationalactivities == 1 & Moderaterecreationalactivities == 1
```

```
label define level_of_exercise 1 "not Moderate or Vigorous" 2 "Only Moderate" 3 "Only  
Vigorous" 4
```

```
"Moderate and Vigorous" label values
```

```
atleast_mod level_of_exercise label variable
```

```
atleast_mod "At least moderate exercise"
```

```
2 . tabulate atleast_mod
```

At least moderate exercise	Freq.	Percent	Cum.
not Moderate or Vigorous	3,120	53.28	53.28
Only Moderate	1,302	22.23	75.51
Only Vigorous	428	7.31	82.82
Moderate and Vigorous	1,006	17.18	100.00
Total	5,856	100.00	

A.2 Arthritis and Gout generate

```
combined_gout_and_arthritis = .
```

```
replace combined_gout_and_arthritis = 1 if arthritis == 2 & gout == 2
```

```
replace combined_gout_and_arthritis = 2 if arthritis == 1 & gout == 2
```

```
replace combined_gout_and_arthritis = 3 if gout == 1 & arthritis == 2
```

```
replace combined_gout_and_arthritis = 4 if gout == 1 & arthritis == 1
```

```
label define gout_arthritis 1 "no gout or arthritis " 2 "Only Arthritis" 3 "Only Gout" 4 "Gout and  
Arthritis" label values combined_gout_and_arthritis gout_arthritis label variable  
combined_gout_and_arthritis "Combined gout and arthritis"
```

```
3 . tabulate combined_gout_and_arthritis
```

Combined gout and arthritis	Freq.	Percent	Cum.
no gout or arthritis	3,727	67.21	67.21
Only Arthritis	1,481	26.71	93.92
Only Gout	126	2.27	96.19
Gout and Arthritis	211	3.81	100.00
Total	5,545	100.00	

Descriptive Statistics

Table 1

dtable Ageinyearsatscreening i.Gender BodyMassIndexkgm2 i.atleast_mod
i.combined_gout_and_arthritis i.Atleastonecomorbidity i.trouble_sleeping, by (NSAID)

```
5 . dtable Ageinyearsatscreening i.Gender BodyMassIndexkgm2 i.atleast_mod i.combined_gout_and_arthritis i.Atleastonecomorbidity
> y i.trouble_sleeping, by (NSAID)
```

	No	NSAID Yes	Total
N	5,471 (93.6%)	375 (6.4%)	5,846 (100.0%)
Age in years at screening	49.374 (18.886)	57.176 (15.553)	49.875 (18.786)
Gender			
Male	2,685 (49.1%)	149 (39.7%)	2,834 (48.5%)
Female	2,786 (50.9%)	226 (60.3%)	3,012 (51.5%)
Body Mass Index (kg/m**2)	29.531 (7.387)	31.985 (7.875)	29.691 (7.443)
At least moderate exercise			
not Moderate or Vigorous	2,860 (52.3%)	254 (67.7%)	3,114 (53.3%)
Only Moderate	1,219 (22.3%)	81 (21.6%)	1,300 (22.2%)
Only Vigorous	412 (7.5%)	14 (3.7%)	426 (7.3%)
Moderate and Vigorous	980 (17.9%)	26 (6.9%)	1,006 (17.2%)
Combined gout and arthritis			
no gout or arthritis	3,618 (70.0%)	103 (28.3%)	3,721 (67.2%)
Only Arthritis	1,248 (24.1%)	231 (63.5%)	1,479 (26.7%)
Only Gout	119 (2.3%)	7 (1.9%)	126 (2.3%)
Gout and Arthritis	186 (3.6%)	23 (6.3%)	209 (3.8%)
At least one comorbidity			
No	2,388 (43.6%)	47 (12.5%)	2,435 (41.7%)
Yes	3,083 (56.4%)	328 (87.5%)	3,411 (58.3%)
Ever told doctor had trouble sleeping?			
Yes	1,388 (25.4%)	188 (50.3%)	1,576 (27.0%)
No	4,079 (74.6%)	186 (49.7%)	4,265 (73.0%)

Modeling Unadjusted

1.logistic NSAID

atleast_mod

2 . logistic NSAID i.atleast_mod

Logistic regression

Number of obs = 5,846

LR chi2(3) = 54.36

Prob > chi2 = 0.0000

Pseudo R2 = 0.0195

Log likelihood = -1365.497

NSAID	Odds ratio	Std. err.	z	P> z	[95% conf. interval]	
atleast_mod						
Only Moderate	.748193	.0988427	-2.20	0.028	.5775141	.9693144
Only Vigorous	.382616	.1069563	-3.44	0.001	.221217	.661771
Moderate and Vigorous	.2987305	.0624973	-5.78	0.000	.198244	.4501518
_cons	.0888112	.0058147	-36.98	0.000	.0781155	.1009713

Note: _cons estimates baseline odds.

1. logistic NSAID Ageinyearsatscreening

2 . logistic NSAID Ageinyearsatscreening

Logistic regression

Number of obs = 5,846

LR chi2(1) = 61.96

Prob > chi2 = 0.0000

Pseudo R2 = 0.0222

Log likelihood = -1361.6981

NSAID	Odds ratio	Std. err.	z	P> z	[95% conf. interval]	
Ageinyearsatscreening	1.023362	.0030887	7.65	0.000	1.017326	1.029433
_cons	.0199946	.0035859	-21.81	0.000	.0140688	.0284164

Note: _cons estimates baseline odds.

2. logistic NSAID Gender

3 . logistic NSAID Gender

Logistic regression

Log likelihood = **-1386.493**

Number of obs = **5,846**

LR chi2(1) = **12.37**

Prob > chi2 = **0.0004**

Pseudo R2 = **0.0044**

NSAID	Odds ratio	Std. err.	z	P> z	[95% conf. interval]	
Gender	1.461791	.1592451	3.49	0.000	1.180747	1.80973
_cons	.0554935	.0046706	-34.35	0.000	.0470544	.0654461

Note: **_cons** estimates baseline odds.

4.logistic NSAID BodyMassIndexkgm2

Logistic regression

Log likelihood = **-1289.6823**

Number of obs = **5,425**

LR chi2(1) = **32.16**

Prob > chi2 = **0.0000**

Pseudo R2 = **0.0123**

NSAID	Odds ratio	Std. err.	z	P> z	[95% conf. interval]	
BodyMassIndexkgm2	1.038538	.0066194	5.93	0.000	1.025645	1.051593
_cons	.021819	.0045727	-18.25	0.000	.0144692	.0329021

Note: **_cons** estimates baseline odds.

5. logistic NSAID combined gout and arthritis

5 . logistic NSAID combined_gout_and_arthritis

Logistic regression

Number of obs = 5,535

LR chi2(1) = 111.09

Prob > chi2 = 0.0000

Pseudo R2 = 0.0414

Log likelihood = -1286.9132

NSAID	Odds ratio	Std. err.	z	P> z	[95% conf. interval]	
combined_gout_and_arthritis	1.889315	.1058224	11.36	0.000	1.692886	2.108536
_cons	.0254875	.0029125	-32.11	0.000	.0203732	.0318858

Note: _cons estimates baseline odds.

6.logistic NSAID Atleastonecomorbidity

6 . logistic NSAID Atleastonecomorbidity

Logistic regression

Number of obs = 5,846

LR chi2(1) = 161.61

Prob > chi2 = 0.0000

Pseudo R2 = 0.0580

Log likelihood = -1311.8702

NSAID	Odds ratio	Std. err.	z	P> z	[95% conf. interval]	
Atleastonecomorbidity	5.405507	.8558555	10.66	0.000	3.963378	7.372375
_cons	.0196818	.002899	-26.67	0.000	.0147465	.0262688

Note: _cons estimates baseline odds.

7.logistic NSAID trouble_sleeping

4 . logistic NSAID trouble_sleeping

Logistic regression

Number of obs = 5,841

LR chi2(1) = 98.21

Prob > chi2 = 0.0000

Pseudo R2 = 0.0353

Log likelihood = -1340.5583

NSAID	Odds ratio	Std. err.	z	P> z	[95% conf. interval]	
trouble_sleeping	.3366595	.0363547	-10.08	0.000	.272441	.4160152
_cons	.4023255	.0694288	-5.28	0.000	.2868711	.5642457

Note: _cons estimates baseline odds.

Adjusted

logistic NSAID i.atleast_mod i.Gender BodyMassIndexkgm2 Ageinyearsatscreening
i.combined_gout_and_arthritis i.Atleastonecomorbidity i.trouble_sleeping

```
2 . logistic NSAID i.atleast_mod i.Gender BodyMassIndexkgm2 Ageinyearsatscreening i.combined_gout_and_arthritis i.Atleastoneco
> morbidity i.trouble_sleeping
```

Logistic regression

Number of obs = 5,145

LR chi2(11) = 300.55

Prob > chi2 = 0.0000

Log likelihood = -1107.2505

Pseudo R2 = 0.1195

NSAID	Odds ratio	Std. err.	z	P> z	[95% conf. interval]	
atleast_mod						
Only Moderate	.760945	.1090436	-1.91	0.057	.5746135	1.007699
Only Vigorous	.7367289	.2160868	-1.04	0.298	.4146154	1.309092
Moderate and Vigorous	.5062747	.1134909	-3.04	0.002	.3262674	.7855952
Gender						
Female	1.126903	.1353883	0.99	0.320	.8904744	1.426105
BodyMassIndexkgm2	1.014508	.0073901	1.98	0.048	1.000127	1.029097
Ageinyearsatscreening	.9977981	.0042015	-0.52	0.601	.9895973	1.006067
combined_gout_and_arthritis						
Only Arthritis	3.89333	.6297433	8.40	0.000	2.835567	5.345674
Only Gout	1.298742	.5794323	0.59	0.558	.5417019	3.113763
Gout and Arthritis	2.687076	.7369995	3.60	0.000	1.569698	4.599852
Atleastonecomorbidity						
Yes	1.750032	.3722645	2.63	0.009	1.153399	2.655293
trouble_sleeping						
No	.5581071	.0668639	-4.87	0.000	.4413057	.7058226
_cons	.0281218	.010441	-9.62	0.000	.0135835	.0582205

Note: _cons estimates baseline odds.

Sensitivity Analysis

Area Under the Curve

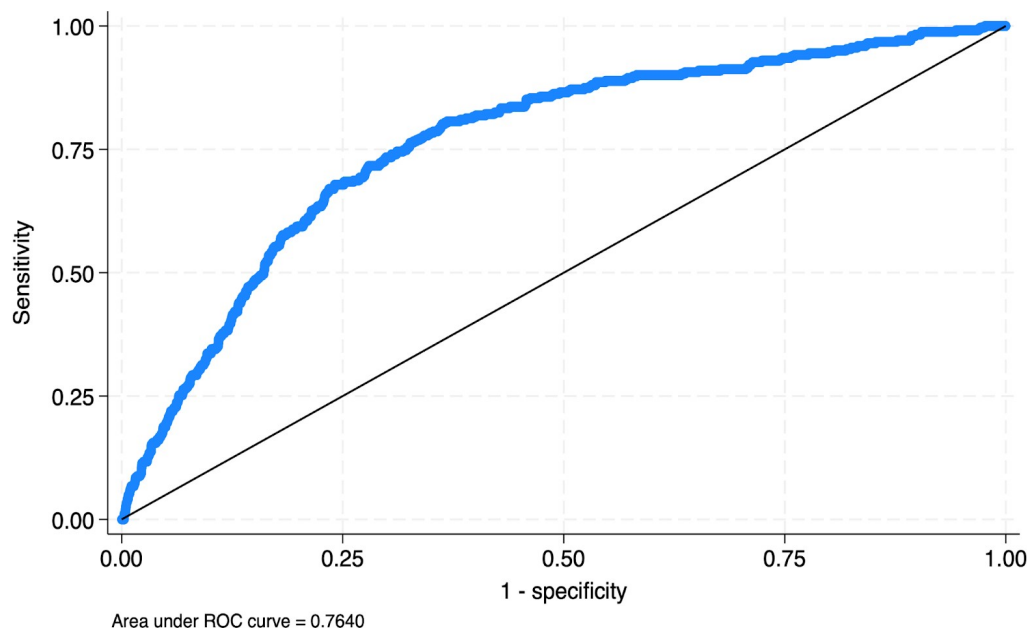
Lroc

```
13 . lroc
```

Logistic model for NSAID

Number of observations = **5145**

Area under ROC curve = **0.7640**



Goodness of fit

estat gof

14 . estat gof

Goodness-of-fit test after logistic model
Variable: **NSAID**

Number of observations = **5,145**
Number of covariate patterns = **5,071**
Pearson chi2(**5059**) = **5008.11**
Prob > chi2 = **0.6918**

Multicollinearity

vif, uncentered

9 . vif, uncentered

Variable	VIF	1/VIF
atleast_mod		
2	1.41	0.709611
3	1.13	0.882457
4	1.29	0.772941
1.Gender	2.08	0.481075
BodyMassIn~2	8.56	0.116782
Ageinyears~g	9.22	0.108493
combined_g~s		
2	2.11	0.473696
3	1.10	0.908538
4	1.17	0.852503
1.Atleasto~y	4.05	0.246649
2.trouble_~g	3.34	0.299080
Mean VIF	3.23	

Model Adjusted to exclude Multicollinear variables

logistic NSAID i.atleast_mod i.Gender i.combined_gout_and_arthritis i.Atleastonecomorbidity
i.trouble_sleeping

```
10 . logistic NSAID i.atleast_mod i.Gender i.combined_gout_and_arthritis i.Atleastonecomorbidity i.trouble_sleeping
```

Logistic regression

Number of obs = 5,530

LR chi2(9) = 320.10

Prob > chi2 = 0.0000

Log likelihood = -1179.41

Pseudo R2 = 0.1195

NSAID	Odds ratio	Std. err.	z	P> z	[95% conf. interval]	
atleast_mod						
Only Moderate	.7910378	.1096459	-1.69	0.091	.6028539	1.037964
Only Vigorous	.7178137	.2077826	-1.15	0.252	.407022	1.265918
Moderate and Vigorous	.5029818	.1084994	-3.19	0.001	.3295623	.7676566
Gender						
Female	1.147779	.1329078	1.19	0.234	.91473	1.440203
combined_gout_and_arthritis						
Only Arthritis	3.867344	.5881246	8.89	0.000	2.87056	5.210254
Only Gout	1.457999	.6044268	0.91	0.363	.6469697	3.28572
Gout and Arthritis	2.629854	.6845381	3.71	0.000	1.578949	4.380213
Atleastonecomorbidity						
Yes	1.785778	.3648324	2.84	0.005	1.196542	2.665182
trouble_sleeping						
No	.5321758	.0613782	-5.47	0.000	.4245044	.6671568
_cons	.0380184	.0074763	-16.63	0.000	.0258586	.0558961

Sample Characteristic	NSAID used in last 30 days		
	No n(%)	Yes n(%)	Total n(%)
Number of participants - n (%)	4,803 (93.4%)	342 (6.6%)	5,145 (100.0%)
Age in years at screening - Mean (SD)	50.754 (17.752)*	57.637 (14.558)*	51.211 (17.640)*
Sex			
Male - n (%)	2,343 (48.8%)	138 (40.4%)	2,481 (48.2%)
Female - n (%)	2,460 (51.2%)	204 (59.6%)	2,664 (51.8%)
Body mass index (kg/m²) - Mean (SD)	29.692 (7.342)*	31.961 (7.747)*	29.842 (7.390)*
Level of recreational exercise			
Not moderate or vigorous - n (%)	2,515 (52.4%)	230 (67.3%)	2,745 (53.4%)
Only moderate - n (%)	1,121 (23.3%)	73 (21.3%)	1,194 (23.2%)
Only vigorous - n (%)	333 (6.9%)	14 (4.1%)	347 (6.7%)
Moderate and vigorous - n (%)	834 (17.4%)	25 (7.3%)	859 (16.7%)
Combined gout and arthritis			
No gout or Arthritis - n (%)	3,369 (70.1%)	99 (28.9%)	3,468 (67.4%)
Only arthritis - n (%)	1,152 (24.0%)	215 (62.9%)	1,367 (26.6%)
Only gout - n (%)	112 (2.3%)	6 (1.8%)	118 (2.3%)
Gout and Arthritis - n (%)	170 (3.5%)	22 (6.4%)	192 (3.7%)
At least one comorbidity			
No - n (%)	2,001 (41.7%)	42 (12.3%)	2,043 (39.7%)
Yes - n (%)	2,802 (58.3%)	300 (87.7%)	3,102 (60.3%)
Ever told doctor had trouble sleeping?			
No - n (%)	3,540 (73.7%)	170 (49.7%)	3,710 (72.1%)
Yes - n (%)	1,263 (26.3%)	172 (50.3%)	1,435 (27.9%)

*Mean (standard deviation)

Table 2. Unadjusted and adjusted models			
<i>Variables</i>	Odds ratio (95 %CI)	Standard error	P value
2.1 Unadjusted model			
atleast_mod*			
Only moderate	0.74 (0.57, 0.96)	0.098	0.028
Only vigorous	0.38 (0.22, 0.66)	0.106	0.001
Moderate and vigorous	0.29 (0.19, 0.45)	0.062	0.000
Age at years at screening	1.02 (1.01, 1.02)	0.003	0.000
Gender			
Female	1.46 (1.18, 1.80)	0.159	0.000
BodyMassIndexkg m2	1.04 (1.02, 1.05)	0.006	0.000
Combined arthritis and gout	1.88 (1.69, 2.11)	0.105	0.000
Trouble sleeping			
No	0.34 (0.27, 0.41)	0.036	0.000
At least one comorbidity			
Yes	5.40 (3.96, 7.37)	0.855	0.000
2.2 Adjusted model			
atlesat_mod			
Only moderate	0.76 (0.57, 1.00)	0.109	0.057
Only vigorous	0.73 (0.41, 1.30)	0.216	0.298

Moderate and vigorous	0.50 (0.32, 0.78)	0.113	0.002
Gender			
Female	1.12 (0.89, 1.42)	0.135	0.320
BodyMassIndexkg m2	1.01 (1.00, 1.02)	0.007	0.048
Age at years at screening	0.99 (0.98, 1.00)	0.004	0.601
Combined gout and arthritis			
Only arthritis	3.89 (2.83, 5.34)	0.629	0.000
Only gout	1.29 (0.54, 3.11)	0.579	0.558
Gout and arthritis	2.68 (1.56, 4.59)	0.736	0.000
Trouble sleeping			
No	0.55 (0.44, 0.70)	0.066	0.000
At least one comorbidity			
Yes	1.75 (1.15, 2.65)	0.372	0.009

*atleast_mod (at least moderate exercise): represents the level of recreational exercise