## Advanced Regression Methods for Independent Data

# STAT/BIOST 570, 2020

Practical Aspects of Regression Models with Weak Assumptions

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### Back to the FEV Data

- FEV: forced expiratory volume. FEV1: amount of air you can force from your lungs in one second.
- Data from 654 children and youths ages 3–19 in East Boston, 1980. (Childhood Respiratory Disease Study).
- For more information visit: http://www.statsci.org/data/general/fev.html

```
> url <- "http://www.statsci.org/data/general/fev.txt"
> data <- read.table(file = url, header = T, sep="\t", stringsAsFactors = F)
> data$Sex <- factor(data$Sex, levels=c("Male", "Female"), labels=c(0,1))
> data$Smoker <- factor(data$Smoker, levels=c("Non", "Current"), labels=c(0,1))
> data$HeightC <- data$Height - 60
> data$AgeC <- data$Age - 10
> data$HeightCSq <- data$HeightC^2</pre>
```

### An Initial Normal Linear Model

Based on an exploratory analysis, we first looked at a normal linear model with regression function:

```
\begin{split} \mathsf{E}(\mathsf{FEV1} \mid \mathbf{X}) = & \beta_0 + \beta_1 \mathsf{Height} + \beta_2 \mathsf{Height}^2 + \beta_3 \mathsf{Age} + \\ & (\beta_4 + \beta_5 \mathsf{Height} + \beta_6 \mathsf{Height}^2 + \beta_7 \mathsf{Age}) I(\mathsf{Sex=Female}) + \\ & (\beta_8 + \beta_9 \mathsf{Height} + \beta_{10} \mathsf{Height}^2 + \beta_{11} \mathsf{Age})) I(\mathsf{Smoker=Yes}) \end{split}
```

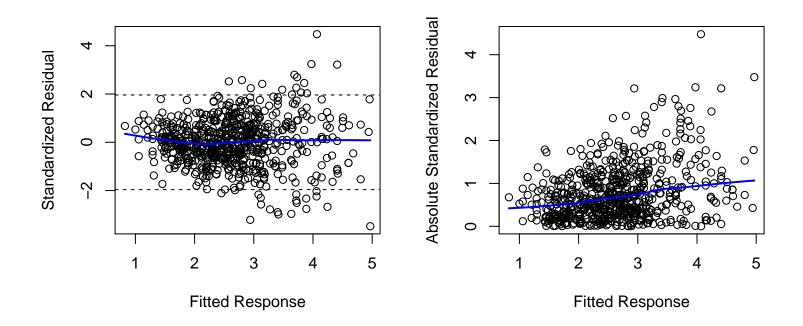
```
> model3 <- lm(
+ FEV ~ (HeightC+HeightCSq+AgeC)*(Sex+Smoker),
+ data=data)</pre>
```

### An Initial Normal Linear Model

Based on the fit of this normal linear model, we looked at the residuals to check whether its assumptions were reasonable

```
> limits3 <- range(data$FEV, fitted(model3))
> par(mfrow = c(1,2))
> library(MASS)
> resid_mod3 <- stdres(model3)
> plot(fitted(model3), resid_mod3,
+    ylab = "Standardized Residual", xlab = "Fitted Response")
> lines(lowess(fitted(model3), resid_mod3), lwd = 2, col = "blue")
> abline(h = 1.96, lty = 2)
> abline(h = -1.96, lty = 2)
> plot(fitted(model3), abs(resid_mod3),
+    ylab = "Absolute Standardized Residual", xlab = "Fitted Response")
> lines(lowess(fitted(model3), abs(resid_mod3)), lwd = 2, col = "blue")
```

### An Initial Normal Linear Model



We concluded that lack of homoscedasticity was an issue, so inferences based on a normal linear model wouldn't be reliable

### A Gamma GLM

We then considered a gamma GLM, for which

$$var(Y_i \mid \mathbf{x}_i) = \alpha V(\mu_i) = \alpha \mu_i^2 = \alpha E(Y_i \mid \mathbf{x}_i)^2,$$

which we thought could be appropriate for our data

We used the identity link, which is fine here since the outcome is away from zero

```
> FEV_Gamma <- glm(
+ FEV ~ (HeightC+HeightCSq+AgeC)*(Sex+Smoker),
+ family = Gamma(link="identity"),
+ data=data)</pre>
```

### In this case, $\hat{\alpha}$ was

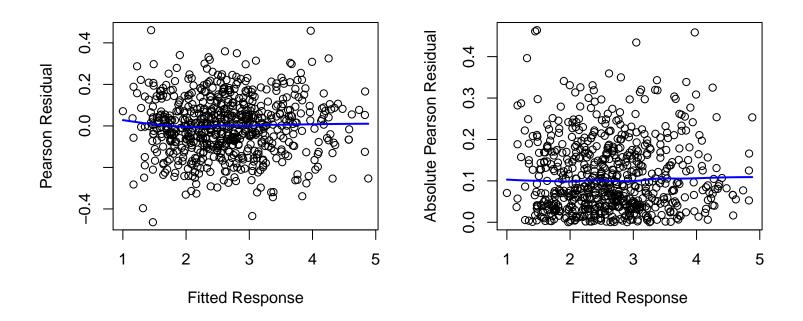
```
> summary(FEV_Gamma)$dispersion
[1] 0.02015759
```

### A Gamma GLM

#### We checked the Pearson residuals

```
> resid_Gamma <- residuals(FEV_Gamma, type = "pearson")
> fitted_Gamma <- fitted(FEV_Gamma)
> limits <- range(data$FEV, fitted_Gamma)
> par(mfrow = c(1,2))
> plot(fitted_Gamma, resid_Gamma,
+ ylab = "Pearson Residual", xlab = "Fitted Response")
> lines(lowess(fitted_Gamma, resid_Gamma), lwd = 2, col = "blue")
> abline(h = 1.96, lty = 2)
> abline(h = -1.96, lty = 2)
> plot(fitted_Gamma, abs(resid_Gamma),
+ ylab = "Absolute Pearson Residual", xlab = "Fitted Response")
> lines(lowess(fitted_Gamma, abs(resid_Gamma)), lwd = 2, col = "blue")
```

### A Gamma GLM



We did not find patterns in the Pearson residuals that indicated violations to the model's assumptions

## Quasi-Likelihood Approaches

In R, some quasi-likelihood approaches are implemented with the glm function

The regression function is taken as in a GLM:

$$\mathsf{E}(Y_i \mid \mathbf{x}_i) = \mu_i = \mu(\mathbf{x}_i, \boldsymbol{\beta}) = g^{-1}(\mathbf{x}_i \boldsymbol{\beta})$$

for a link function  $g(\cdot)$ 

The variance function is taken as

$$var(Y_i \mid \mathbf{x}_i) = \alpha V(\mu_i)$$

for some known function  $V(\cdot)$ 

## Quasi-Likelihood Approaches

To specify a quasi-likelihood model, the family argument of glm can be taken as one of the following default options

```
quasi(link = "identity", variance = "constant")
quasibinomial(link = "logit")
quasipoisson(link = "log")
```

Available link functions: logit, probit, cloglog, identity, inverse, log, 1/mu^2 and sqrt. Also, the function power can be used to create a power link function

Available variance functions for quasi: constant, mu(1-mu), mu, mu^2 and mu^3

Variance function for quasibinomial is proportional to mu(1-mu)

Variance function for quasipoisson is proportional to mu

## Quasi-Likelihood Approaches

Quasi-examples (pun totally intended!):

• glm(y~x1+x2, family = quasi(link="identity", variance="mu^2"))
corresponds to assuming

$$g(\mu_i) = \mu_i, \ V(\mu_i) = \mu_i^2,$$

that is,

$$\mathsf{E}(Y_i \mid \mathbf{x}_i) = \mathbf{x}_i \boldsymbol{\beta}, \ \ \mathsf{var}(Y_i \mid \mathbf{x}_i) = \alpha \mathsf{E}(Y_i \mid \mathbf{x}_i)^2$$

• glm(y~x1+x2, family = quasipoisson(link = "log"))
corresponds to assuming

$$g(\mu_i) = \log(\mu_i), \ V(\mu_i) = \mu_i,$$

that is,

$$\mathsf{E}(Y_i \mid \mathbf{x}_i) = \exp(\mathbf{x}_i \boldsymbol{\beta}), \ \ \mathsf{var}(Y_i \mid \mathbf{x}_i) = \alpha \mathsf{E}(Y_i \mid \mathbf{x}_i)$$

Continuing with the FEV data, let us first consider a quasi-likelihood approach with

$$var(Y_i \mid \mathbf{x}_i) = \alpha V(\mu_i) = \alpha \mu_i = \alpha E(Y_i \mid \mathbf{x}_i)$$

```
> FEV_Quasi1 <- glm(
+ FEV ~ (HeightC+HeightCSq+AgeC)*(Sex+Smoker),
+ family = quasi(link="identity", variance="mu"),
+ data=data)</pre>
```

### In this case, $\hat{\alpha}$ is

> summary(FEV\_Quasi1)\$dispersion

[1] 0.05291511

The quasi-likelihood approach relies on correct specification of the variance function,  $var(Y_i \mid \mathbf{X}_i) = \alpha V(\mu_i)$ 

Note that if the mean-variance relationship holds, then the Pearson residuals

$$e_i^{\star} = \frac{Y_i - \mu_i(\widehat{\boldsymbol{\beta}})}{\sqrt{\widehat{\alpha}V(\mu_i(\widehat{\boldsymbol{\beta}}))}}$$

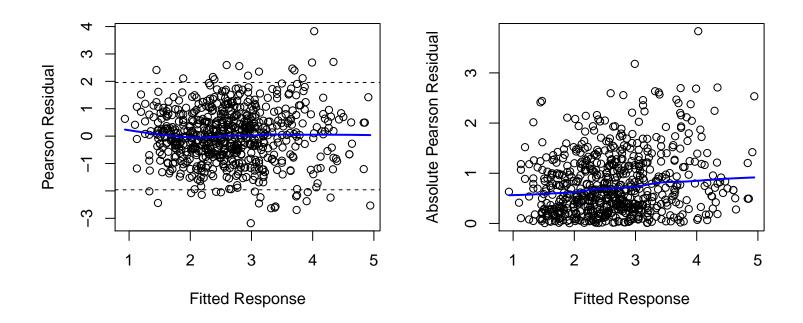
should fluctuate around 0 over the whole range of the covariates

As a function of  $\mu_i(\widehat{\beta})$ , the Pearson residuals should also fluctuate around 0 with homogeneous variance

Note that in R, the residuals function doesn't use  $\hat{\alpha}$  in computing Pearson residuals

#### Computation and plotting of the Pearson residuals

```
> resid_FEV_Quasi1 <- residuals(FEV_Quasi1, type = "pearson")/sqrt(summary(FEV_Quasi1)$dispersion)
> fitted_FEV_Quasi1 <- fitted(FEV_Quasi1)
> limits <- range(data$FEV, fitted_FEV_Quasi1)
> par(mfrow = c(1,2))
> plot(fitted_FEV_Quasi1, resid_FEV_Quasi1,
+ ylab = "Pearson Residual", xlab = "Fitted Response")
> lines(lowess(fitted_FEV_Quasi1, resid_FEV_Quasi1), lwd = 2, col = "blue")
> abline(h = 1.96, lty = 2)
> abline(h = -1.96, lty = 2)
> plot(fitted_FEV_Quasi1, abs(resid_FEV_Quasi1),
+ ylab = "Absolute Pearson Residual", xlab = "Fitted Response")
> lines(lowess(fitted_FEV_Quasi1, abs(resid_FEV_Quasi1)), lwd = 2, col = "blue")
```



The Pearson residuals reveal that the assumption  $\text{var}(Y_i \mid \mathbf{x}_i) = \alpha \mu_i$  does not quite capture the dependence of the variability of the response as a function of the mean, as we can still see an increase in the variance of the residuals as a function of the fitted means

Let us now consider a quasi-likelihood approach with

$$var(Y_i \mid \mathbf{x}_i) = \alpha V(\mu_i) = \alpha \mu_i^2 = \alpha E(Y_i \mid \mathbf{x}_i)^2$$

```
> FEV_Quasi2 <- glm(
+ FEV ~ (HeightC+HeightCSq+AgeC)*(Sex+Smoker),
+ family = quasi(link="identity", variance="mu^2"),
+ data=data)</pre>
```

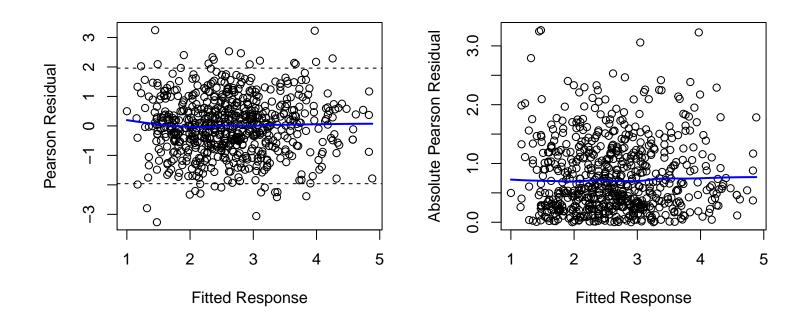
### In this case, $\hat{\alpha}$ is

> summary(FEV\_Quasi2)\$dispersion

[1] 0.02015759

#### Computation and plotting of the Pearson residuals

```
> resid_FEV_Quasi2 <- residuals(FEV_Quasi2, type = "pearson")/sqrt(summary(FEV_Quasi2)$dispersion)
> fitted_FEV_Quasi2 <- fitted(FEV_Quasi2)
> limits <- range(data$FEV, fitted_FEV_Quasi2)
> par(mfrow = c(1,2))
> plot(fitted_FEV_Quasi2, resid_FEV_Quasi2,
+ ylab = "Pearson Residual", xlab = "Fitted Response")
> lines(lowess(fitted_FEV_Quasi2, resid_FEV_Quasi2), lwd = 2, col = "blue")
> abline(h = 1.96, lty = 2)
> abline(h = -1.96, lty = 2)
> plot(fitted_FEV_Quasi2, abs(resid_FEV_Quasi2),
+ ylab = "Absolute Pearson Residual", xlab = "Fitted Response")
> lines(lowess(fitted_FEV_Quasi2, abs(resid_FEV_Quasi2)), lwd = 2, col = "blue")
```



The Pearson residuals indicate that the assumption  ${\rm var}(Y_i\mid {\bf x}_i)=\alpha\mu_i^2$  seems reasonable

Question: how does this compare with the gamma GLM?

The gamma GLM and the quasi-likelihood approach with  $var(Y_i \mid \mathbf{x}_i) = \alpha \mu_i^2$  are indistinguishable from their estimating functions!

```
> identical(summary(FEV_Quasi2)$dispersion, summary(FEV_Gamma)$dispersion)
[1] TRUE
> identical(fitted(FEV_Quasi2), fitted(FEV_Gamma))
[1] TRUE
> identical(vcov(FEV_Quasi2), vcov(FEV_Gamma))
[1] TRUE
```

One more quasi-likelihood approach: let us now consider

$$var(Y_i \mid \mathbf{x}_i) = \alpha V(\mu_i) = \alpha \mu_i^3 = \alpha E(Y_i \mid \mathbf{x}_i)^3$$

```
> FEV_Quasi3 <- glm(
+ FEV ~ (HeightC+HeightCSq+AgeC)*(Sex+Smoker),
+ family = quasi(link="identity", variance="mu^3"),
+ data=data)</pre>
```

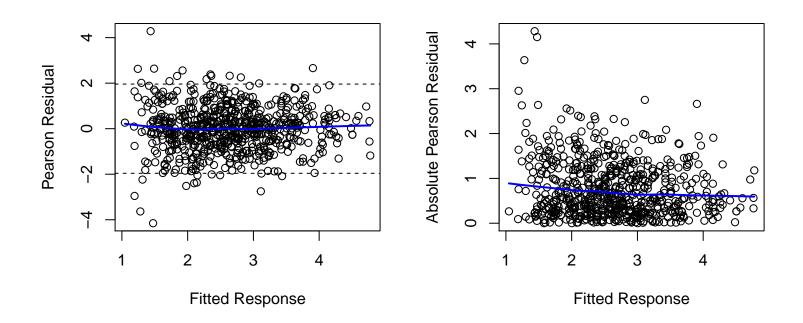
### In this case, $\hat{\alpha}$ is

```
> summary(FEV_Quasi3)$dispersion
```

[1] 0.00845347

#### Computation and plotting of the Pearson residuals

```
> resid_FEV_Quasi3 <- residuals(FEV_Quasi3, type = "pearson")/sqrt(summary(FEV_Quasi3)$dispersion)
> fitted_FEV_Quasi3 <- fitted(FEV_Quasi3)
> limits <- range(data$FEV, fitted_FEV_Quasi3)
> par(mfrow = c(1,2))
> plot(fitted_FEV_Quasi3, resid_FEV_Quasi3,
+ ylab = "Pearson Residual", xlab = "Fitted Response")
> lines(lowess(fitted_FEV_Quasi3, resid_FEV_Quasi3), lwd = 2, col = "blue")
> abline(h = 1.96, lty = 2)
> abline(h = -1.96, lty = 2)
> plot(fitted_FEV_Quasi3, abs(resid_FEV_Quasi3),
+ ylab = "Absolute Pearson Residual", xlab = "Fitted Response")
> lines(lowess(fitted_FEV_Quasi3, abs(resid_FEV_Quasi3)), lwd = 2, col = "blue")
```



These Pearson residuals now indicate that assuming  $var(Y_i \mid \mathbf{x}_i) = \alpha \mu_i^3$  is not appropriate, as it makes the variance of the responses grow too fast, which can be seen from the variability of the Pearson residuals decreasing as a function of the fitted means

The usual table with test statistics for the individual parameters can be obtained from summary or from coeftest in the lmtest package

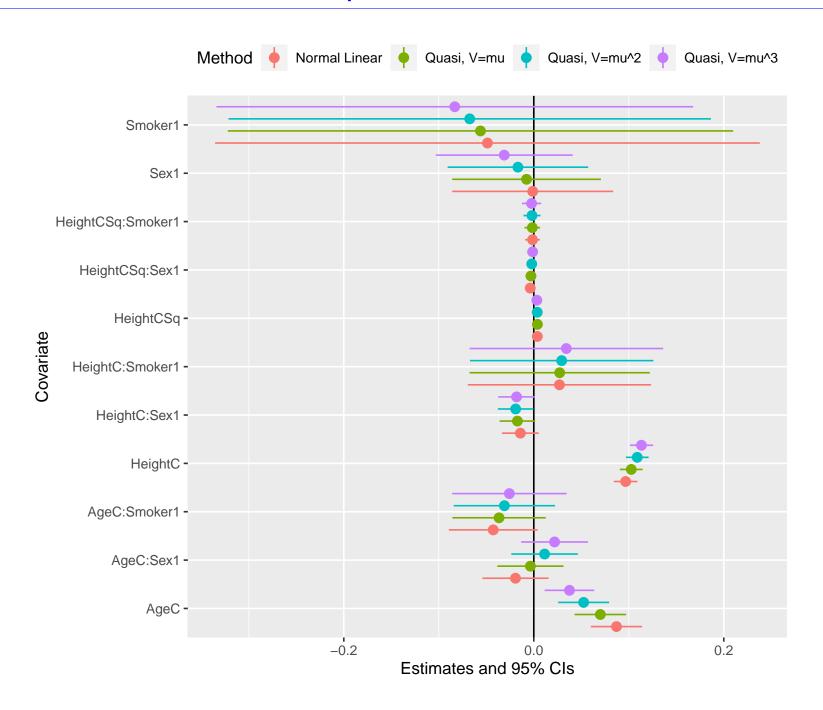
```
> round(coeftest(FEV_Quasi2),3)
```

z test of coefficients:

	Estimate Std.	Error z value	Pr(> z )	
(Intercept)	2.436	0.030 82.239	<2e-16 ***	
HeightC	0.109	0.006 17.851	<2e-16 ***	
HeightCSq	0.004	0.000 7.499	<2e-16 ***	
AgeC	0.052	0.014 3.833	<2e-16 ***	
Sex1	-0.017	0.038 -0.443	0.657	
Smoker1	-0.067	0.130 -0.521	0.603	
HeightC:Sex1	-0.019	0.010 -1.999	0.046 *	
HeightC:Smoker1	0.029	0.049 0.596	0.551	
HeightCSq:Sex1	-0.002	0.001 -2.882	0.004 **	
HeightCSq:Smoker1	-0.002	0.005 -0.421	0.674	
AgeC:Sex1	0.011	0.018 0.636	0.525	
AgeC:Smoker1	-0.031	0.027 -1.138	0.255	
Signif. codes: 0	'***' 0.001 '	**' 0 01 '*' 0	.05 '.' 0.1 ' '	1

Let's compare the point estimates and confidence intervals obtained from each quasi-likelihood approach

```
> normal_CIs <- data.frame(coefci(model3, vcov = vcov(model3)), method="Normal Linear")
> quasi1_CIs <- data.frame(coefci(FEV_Quasi1, vcov = vcov(FEV_Quasi1)), method="Quasi, V=mu")
> quasi2_CIs <- data.frame(coefci(FEV_Quasi2, vcov = vcov(FEV_Quasi2)), method="Quasi, V=mu^2")
> quasi3_CIs <- data.frame(coefci(FEV_Quasi3, vcov = vcov(FEV_Quasi3)), method="Quasi, V=mu^3")
> all_CIs <- rbind(normal_CIs[-1,],quasi1_CIs[-1,],quasi2_CIs[-1,],quasi3_CIs[-1,])
> all_CIs <- data.frame(all_CIs, param=rownames(normal_CIs)[-1],
         pointest = c(model3$coeff[-1],FEV_Quasi1$coeff[-1],
                                  FEV_Quasi2$coeff[-1],FEV_Quasi3$coeff[-1])
> names(all_CIs)[1:2] <- c("lower", "upper")
> myplot <- ggplot(data = all_CIs, aes(x = param, y = pointest,
                            ymin = lower, ymax = upper, group = method, color = method))
+
> myplot + geom_hline(vintercept=0, color="black") +
          geom_pointrange(position = position_dodge(width = 1)) +
          theme(legend.position = "top") + coord_flip() +
          labs(x = "Covariate", y = "Estimates and 95% CIs", color = "Method")
+
```



We had the following question: is there any association between FEV and smoking after controlling for other variables?

Based on the model for the mean

E(FEV1 | X) = 
$$\beta_0 + \beta_1$$
Height +  $\beta_2$ Height<sup>2</sup> +  $\beta_3$ Age+  
( $\beta_4 + \beta_5$ Height +  $\beta_6$ Height<sup>2</sup> +  $\beta_7$ Age) $I(Sex=Female)$ +  
( $\beta_8 + \beta_9$ Height +  $\beta_{10}$ Height<sup>2</sup> +  $\beta_{11}$ Age)) $I(Smoker=Yes)$ 

we can test the hull hypothesis

$$H_0: \beta_8 = \beta_9 = \beta_{10} = \beta_{11} = 0$$

to try to answer the question

Let us do this comparing the fit of the quasi-likelihood with quadratic variance FEV\_Quasi2 with a reduced model that does not include the Smoker variable, say FEV\_Quasi2\_red:

```
> FEV_Quasi2_red <- glm(
+ FEV ~ (HeightC+HeightCSq+AgeC)*Sex,
+ family = quasi(link="identity", variance="mu^2"),
+ data=data)</pre>
```

```
> waldtest(FEV_Quasi2_red, FEV_Quasi2, test = "Chisq") # Wald test
Wald test
Model 1: FEV ~ (HeightC + HeightCSq + AgeC) * Sex
Model 2: FEV ~ (HeightC + HeightCSq + AgeC) * (Sex + Smoker)
 Res.Df Df Chisq Pr(>Chisq)
    646
1
    642 4 3.0416
                      0.5509
> anova(FEV_Quasi2_red, FEV_Quasi2, test = "Rao") # Score test
Analysis of Deviance Table
Model 1: FEV ~ (HeightC + HeightCSq + AgeC) * Sex
Model 2: FEV ~ (HeightC + HeightCSq + AgeC) * (Sex + Smoker)
 Resid. Df Resid. Dev Df Deviance
                                       Rao Pr(>Chi)
        646
                13.462
2
        642
                13.404 4 0.058237 0.056386
                                             0.5923
```

Therefore the larger model is *not* a better fit, and so we lack evidence that FEV is associated with smoking status after controlling for other variables in this model

Note that the likelihood-ratio test does not make sense under quasi-likelihood, as there is no likelihood

We can base our inferences on sandwich estimators, so as not to rely on correct specification of the mean-variance relationship

The R package sandwich contains functions to extract the pieces required to compute sandwich estimators from the fit of the function glm, lm and others, see Zeileis (2004, 2006, Journal of Statistical Software)

The package sandwich still relies on the regression function being correctly specified

It computes the most general version of  $\widehat{f B}_n$  for uncorrelated data as

$$\widehat{\mathbf{B}}_n = \frac{1}{n} \widehat{\mathbf{D}}^T \widehat{\mathbf{V}}^{-1} \operatorname{diag}\{ [Y_i - \mu_i(\widehat{\beta}_n)]^2 \} \widehat{\mathbf{V}}^{-1} \widehat{\mathbf{D}},$$

which is known as the Eicker-Huber-White (HC0) estimator, and the most general version of  $\widehat{\mathbf{A}}_n$  under correct specification of the mean model

$$\hat{\mathbf{A}}_n = rac{1}{n}\hat{\mathbf{D}}^T\hat{\mathbf{V}}^{-1}\hat{\mathbf{D}}$$

See slides9.pdf, p. 39

```
> library(sandwich) # Default is the Eicker-Huber-White estimator
> coeftest(FEV_Quasi2, vcov = sandwich) # Same as vcov = vcovHC(FEV_Quasi2, type = "HCO")
z test of coefficients:
                 Estimate Std. Error z value Pr(>|z|)
(Intercept)
               2.43615775 0.03226790 75.4979 < 2.2e-16 ***
HeightC
              0.10888453  0.00556352  19.5712 < 2.2e-16 ***
HeightCSq
            AgeC
             -0.01673334 0.03929730 -0.4258
Sex1
                                           0.67024
        -0.06745471 0.14136928 -0.4772
                                           0.63325
Smoker1
HeightC:Sex1 -0.01910880 0.00987039 -1.9360
                                           0.05287 .
HeightC:Smoker1 0.02935413 0.04568911 0.6425
                                           0.52056
HeightCSq:Sex1
             -0.00217012 0.00088874 -2.4418
                                           0.01461 *
HeightCSq:Smoker1 -0.00192167 0.00382853 -0.5019
                                           0.61571
         0.01136177 0.01707719 0.6653
AgeC:Sex1
                                           0.50585
AgeC:Smoker1 -0.03095755 0.02233657 -1.3860
                                           0.16576
Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
```

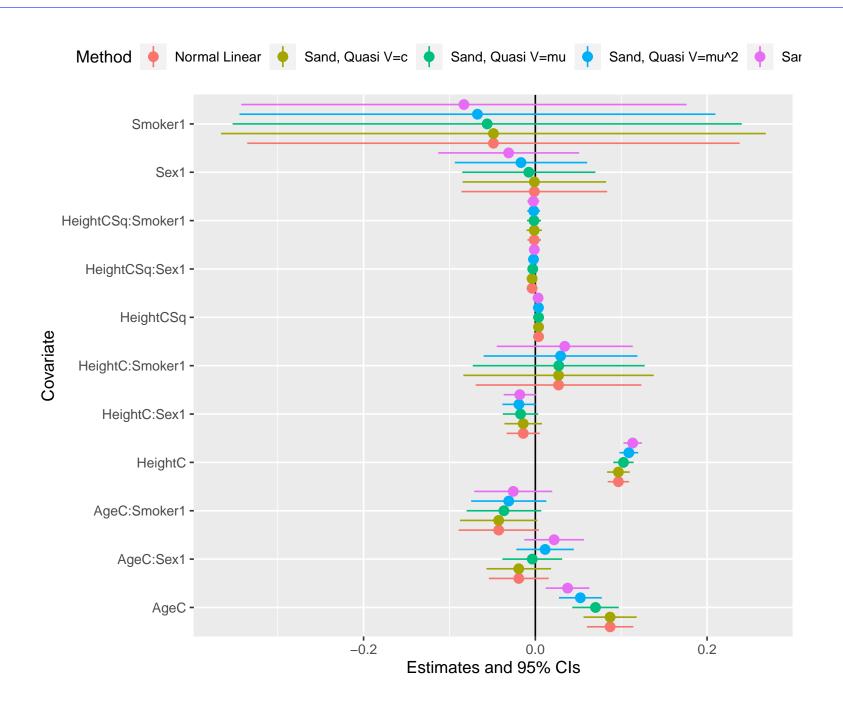
The argument type can also be specified as HC1, which replaces  $e_i^2 := [Y_i - \mu_i(\hat{\beta}_n)]^2$  with  $\frac{n}{n-k-1}e_i^2$  in  $\hat{\mathbf{B}}_n$ ; other options are available, see Zeileis (2004)

#### Compare these two:

Relies on correct mean-variance specification > waldtest(FEV\_Quasi2\_red, FEV\_Quasi2, test = "Chisq") Wald test Model 1: FEV ~ (HeightC + HeightCSq + AgeC) \* Sex Model 2: FEV ~ (HeightC + HeightCSq + AgeC) \* (Sex + Smoker) Res.Df Df Chisq Pr(>Chisq) 646 642 4 3.0416 0.5509 Doesn't assume correct variance specification > waldtest(FEV\_Quasi2\_red, FEV\_Quasi2, test = "Chisq", vcov = sandwich) Wald test Model 1: FEV ~ (HeightC + HeightCSq + AgeC) \* Sex Model 2: FEV ~ (HeightC + HeightCSq + AgeC) \* (Sex + Smoker) Res.Df Df Chisq Pr(>Chisq) 646 642 4 3.9253 0.4162

Let's compare the point estimates and confidence intervals obtained from each quasi-likelihood approach

```
> normal_CIs <- data.frame(coefci(model3, vcov = vcov(model3)), method="Normal Linear")
> sand_quasi0_CIs <- data.frame(coefci(model3, vcov = sandwich), method="Sand, Quasi V=c")
> sand_quasi1_CIs <- data.frame(coefci(FEV_Quasi1, vcov = sandwich), method="Sand, Quasi V=mu")
> sand_quasi2_CIs <- data.frame(coefci(FEV_Quasi2, vcov = sandwich), method="Sand, Quasi V=mu^2")
> sand_quasi3_CIs <- data.frame(coefci(FEV_Quasi3, vcov = sandwich), method="Sand, Quasi V=mu^3")
> all_CIs <- rbind(normal_CIs[-1,],sand_quasi0_CIs[-1,],sand_quasi1_CIs[-1,],
          sand_quasi2_CIs[-1,],sand_quasi3_CIs[-1,])
> all_CIs <- data.frame(all_CIs, param=rownames(normal_CIs)[-1],
         pointest = c(model3$coeff[-1], model3$coeff[-1], FEV_Quasi1$coeff[-1],
                                  FEV_Quasi2$coeff[-1],FEV_Quasi3$coeff[-1])
> names(all_CIs)[1:2] <- c("lower", "upper")
> myplot <- ggplot(data = all_CIs, aes(x = param, y = pointest,
                            ymin = lower, ymax = upper, group = method, color = method))
> myplot + geom_hline(yintercept=0, color="black") +
          geom_pointrange(position = position_dodge(width = 1)) +
          theme(legend.position = "top") + coord_flip() +
          labs(x = "Covariate", y = "Estimates and 95% CIs", color = "Method")
```



### **Final Comments**

How can we use all of this in practice?

- Think carefully about the regression function specification; make sure it makes scientific sense
- ullet Try to choose a reasonable  $V(\cdot)$  function for your variance, to base your estimating function
- Use sandwich estimators, just in case you are wrong!
- If you want to rely on stronger assumptions (say in small samples, for efficiency reasons), make sure you can justify them given your data