

## MULTIPLE REGRESSION WITH INTERACTION TERMS

Luis Castro-de-Araujo<sup>a</sup> 11/07/2022

Virginia Institute for Psychiatric and Behavioral Genetics

<sup>&</sup>lt;sup>a</sup> Post-doc T32. luis.araujo@vcuhealth.org

Multiple regression recap

Interaction terms

Visualizing interactions

Marginal effects

## **MULTIPLE REGRESSION RECAP**

#### **MULTIPLE LINEAR REGRESSION**



 Extension of the simple linear regression model to two or more independent variables

$$y = \beta_0 + \beta_1 x_1 + \beta_2 x_2 + \dots + \beta_p x_p + \epsilon$$

- Expression = Baseline + Age + Tissue + Sex + Error
- Partial Regression Coefficients: effect on the dependent variable when increasing the i<sup>th</sup> independent variable by 1 unit, holding all other predictors constant

#### **CATEGORICAL INDEPENDENT VARIABLES**



- Qualitative variables are easily incorporated in regression framework through dummy variables
- Simple example: sex can be coded as 0/1
- What if my categorical variable contains three levels:

$$x_1 = \begin{cases} 0 \text{ if } AA \\ 1 \text{ if } AG \\ 2 \text{ if } GG \end{cases}$$

## **CATEGORICAL INDEPENDENT VARIABLES**



- Previous coding would result in colinearity
- Solution is to set up a series of dummy variable.
- for k levels you need k-1 dummy variables

$$x_1 = \begin{cases} 1 \text{ if AA} \\ 0 \text{ otherwise} \end{cases}$$
$$x_2 = \begin{cases} 1 \text{ if } AG \\ 0 \text{ otherwise} \end{cases}$$

	x1	x2
AA	1	0
AG	0	1
GG	0	0

#### **ASSUMPTIONS**

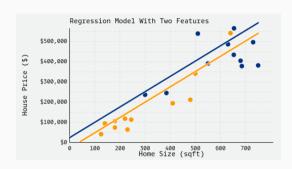


**Validity** Does the data we're modeling matches the problem we're actually trying to solve?

**Representativeness** Is the sample data used in the regression model representative of the population to which it will be applied? **Additivity and Linearity** The deterministic component of a regression model is a linear function of the separate predictors:  $y = B_0 + B_1x_1 + ... + B_px_p$  **Independence of Errors** The errors from our model are independent. **Homoscedasticity** The errors from our model have equal variance. **Normality of Errors** The errors from our model are normally distributed.

#### **MULTIVARIATE REGRESSION**





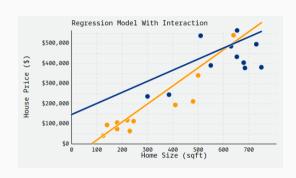
houseprice = -27154 + 757 \* sqft + 51867 \* pool

 In our example, we model home prices as a function of both the size of the house (sqft) and whether or not it has a pool

- intercept: -\$27,154, the predicted average housing price for houses with all x<sub>i</sub> = 0. Or the cost of houses with no pools and a square-footage of zero.
- coefficient of pool: \$51,867, average expected price difference in houses of the same size (in sqft) if they do or do not have a pool. In other words, we expect, on average, houses of the same size to cost \$51,867 more if they have a pool than if they do not.
- coefficient of sqft: \$757, average expected price difference in housing price for houses that have the same value of pool but differ in size by one square-foot.
- We assume the same slope for sqft. Hence, two lines. This isn't always a valid assumption to make.

#### BACK TO OUR HOUSING EXAMPLE, NOW WITH INTERACTIONS





- interaction term: -\$347, represents the difference in the slope for sqft, comparing houses that do and do not have pools. Visually, this represents the difference between the slopes of the two lines.
- intercept: -\$70,296, represents the predicted housing price for houses with no pools and a square-footage of zero.
- coefficient of pool: \$217,111, represents the average expected difference in houses of the same size (0 sqft) that differed in whether or not they had a pool. (It's not super useful since we don't have houses with 0 square-feet).
- coefficient of sqft: \$899, represents the average expected difference in housing price for houses that do not have a pool (pool= 0) but differ in size by one square-foot.

houseprice = -70296 + 899 \* sqft + 217111 \* pool - 347 \* (sqft : pool)

 If we believe that the slope for sqft should differ between houses that do have pools and houses that do not, we can add an interaction term to our model, (sqft:pool).

# **INTERACTION TERMS**

## WHAT IS AN INTERACTION?



• An interaction is a predictor that is some combination of the other predictors.

#### **CONSTRUCTING AN INTERACTION**



- Interactions are often the product of two or more predictors.
- Can be written as,

$$Y = \beta_0 + \beta_1 X_1 + \beta_2 X_2 + \beta_3 X_1 X_2 + \epsilon$$

#### **CONDITIONAL VS. MARGINAL EFFECTS**



- Conditional effects: the effect of a predictor on the response, holding all other predictors constant.
- Marginal effects: the effect of a predictor on the response, averaged over all values of the other predictors.

#### **CONDITIONAL VS. MARGINAL EFFECTS**



• If the conditional effects of X1 on Y at different levels of X2 are all the same then there is no interaction.





Parame	eter Meaning	Where people (used to) go awry
$\beta_0$	Expected value of the DV when X1 and X2 ==0	People get this
$eta_{ extsf{1}} \ eta_{ extsf{2}}$	Effect of X1 when X2 == 0 Effect of X2 when X1 == 0	Not marginal effects! Not marginal effects!
$eta_3$	The addition to the conditional effect when both X! and X2 are 1	People just look at the significance of the interaction parameter and do not calculate the underlying marginal or conditional effects or standard errors

## IN THE PAST IT WAS COMMON TO SEE STANDARD ERRORS WRONGLY CALCULATED

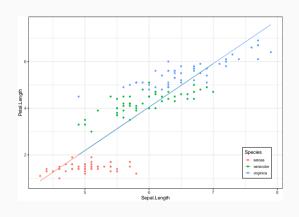


- A common mistake that people make when interpreting interaction models is using the wrong standard errors.
- The standard errors that are printed in every regression table are the positive square roots of the diagonal elements of the variance- covariance matrix of  $\beta$
- This does not matter anymore because of margins()

## INTERACTION AND DUMMMY VARIABLE, IRIS flower DATA SET



$$petal.length_i = \beta_0 + \beta_1 sepal.length_i$$



## LET'S IMPROVE THE MODEL?



## Creating the dummy

$$setosa_i = \begin{cases} 1 \text{ if species of flower } i = setosa, \forall i \in [1, 150] \\ 0 \text{ otherwise} \end{cases}$$

$$versicolor_i = \begin{cases} 1 \text{ if species of flower } i = versicolor, \forall i \in [1, 150] \\ 0 \text{ otherwise} \end{cases}$$

#### Our formula is then

$$petal.length_i = \beta_0 + \beta_1 sepal.length_i + \beta_2 setosa_i + \beta_3 versicolor_i$$

#### BY SUBSTITUTION WE GET THREE LINES WITH SAME SLOPE



#### If it is setosa

$$\begin{split} peta\overline{l.length_i} &= \beta_0 + \beta_1 sepal.length_i + \beta_2 \text{ setosa }_i + \beta_3 \text{ versicolor }_i \\ &= \beta_0 + \beta_1 sepal.length_i + \beta_2 1 + \beta_3 0 \\ &= \beta_0 + \beta_1 sepal.length_i + \beta_2 \\ &= (\beta_0 + \beta_2) + \beta_1 sepal.length_i \end{split}$$

#### If it is versicolor

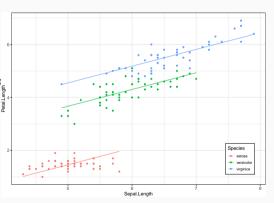
$$\begin{aligned} petal.length_i &= \beta_0 + \beta_1 sepal.length_i + \beta_2 \text{ setosa }_i + \beta_3 \text{ versicolor }_i \\ &= \beta_0 + \beta_1 sepal.length_i + \beta_2 0 + \beta_3 1 \\ &= \beta_0 + \beta_1 sepal.length_i + \beta_3 \\ &= (\beta_0 + \beta_3) + \beta_1 sepal.length_i \end{aligned}$$

## If it is virginica

$$\begin{split} petal.length_i &= \beta_0 + \beta_1 sepal.length_i + \beta_2 \text{ setosa }_i + \beta_3 \text{ versicolor }_i \\ &= \beta_0 + \beta_1 sepal.length_i + \beta_2 0 + \beta_3 0 \\ &= \beta_0 + \beta_1 sepal.length_i \\ &= \beta_0 + \beta_1 sepal.length_i \end{split}$$

## SAME SLOPE, DIFFERENT INTERCEPTS





#### Now we can add an interaction



```
\begin{split} \widehat{petal.length_i} &= \beta_0 + \beta_1 sepal.length_i + \beta_2 \text{ setosa }_i + \beta_3 \text{ versicolor }_i \\ &+ \beta_4 sepal.length_i \text{ setosa }_i + \beta_5 sepal.length_i \text{ versicolor }_i \end{split}
```

- this will result in three unique lines depending on the species of the flower.
- both the intercepts and the slopes will be allowed to be different.

## Does it make sense to retain the interaction?

```
inter <- lm(Petal.Length - Sepal.Length + Species + Sepal.Length:Species, data = iris)</pre>
```

#### DOES IT MAKE SENSE TO RETAIN THE INTERACTION?



```
broom::tidy(inter) |> kable()
```

tidy.summary.glht jtools

Registered S3 methods overwritten by 'broom':
method from
tidy.glht jtools

term	estimate	std.error	statistic	p.value
(Intercept)	0.803	0.531	1.512	0.133
Sepal.Length	0.132	0.106	1.244	0.216
Speciesversicolor	-0.618	0.684	-0.904	0.368
Speciesvirginica	-0.193	0.658	-0.293	0.770
Sepal.Length:Speciesversicolor	0.555	0.128	4.330	0.000
Sepal.Length:Speciesvirginica	0.618	0.121	5.111	0.000

anova(nospecies, w\_species, inter)

Analysis of Variance Table

148 111.5

```
Model 1: Petal.Length - Sepal.Length + Species

Model 2: Petal.Length - Sepal.Length + Species

Model 3: Petal.Length - Sepal.Length + Species + Sepal.Length:Species

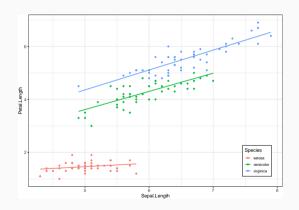
Res.Df RSS Df Sum of Sq F Pr(>F)
```

#### Now we can add an interaction



$$\begin{split} \widehat{petal.length_i} &= \beta_0 + \beta_1 sepal.length_i + \beta_2 \ setosa_i + \beta_3 \ versicolor_i \\ &+ \beta_4 sepal.length_i \ setosa_i + \beta_5 sepal.length_i \ versicolor_i \end{split}$$

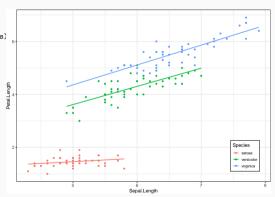
- this will result in three unique lines depending on the species of the flower.
- both the intercepts and the slopes will be allowed to be different.
- ggplot geom\_smooth does this by default if color is used



## Now we can add an interaction



```
ggplot(iris, aes(x = Sepal.Length, y = Petal.Length, color = Species)
geom_point() +
geom_smooth(method = "lm", se = FALSE) +
theme_luis()+
theme( legend.position = c(0.9, 0.15))
```



# **VISUALIZING INTERACTIONS**

## MTCARS DATA SET - REGRESSION OF SPEED ON WT\*CYL



```
fit1 <- glm(qsec ~ wt*as.factor(cyl), data = mtcars)
broom::tidy(fit1) |> kable()
```

 Note: not significant, but we will return to this later summary(margins(fit1))

term	estimate	std.error	statistic	p.value
(Intercept)	14.83	1.500	9.884	0.000
wt	1.89	0.639	2.951	0.007
as.factor(cyl)6	-9.78	4.395	-2.226	0.035
as.factor(cyl)8	-1.44	2.273	-0.632	0.533
wt:as.factor(cyl)6	2.26	1.464	1.546	0.134
wt:as.factor(cyl)8	-1.04	0.764	-1.360	0.185

## **REGRESSION OF MPG ON WT\*CYL**



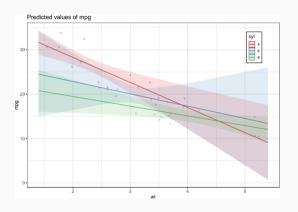
```
fit <- glm(mpg ~ wt*as.factor(cyl), data = mtcars)
broom::tidy(fit) |> kable()
```

term	estimate	std.error	statistic	p.value
(Intercept)	39.57	3.19	12.39	0.000
wt	-5.65	1.36	-4.15	0.000
as.factor(cyl)6	-11.16	9.36	-1.19	0.244
as.factor(cyl)8	-15.70	4.84	-3.25	0.003
wt:as.factor(cyl)6	2.87	3.12	0.92	0.366
wt:as.factor(cyl)8	3.46	1.63	2.12	0.043

## **REGRESSION OF MPG ON WT\*CYL**



```
pred <- ggpredict(fit, terms = c("wt", "cyl"))
plot(pred, add.data = TRUE)+
    theme_luis()+
    theme( legend.position = c(0.1, 0.15))</pre>
```



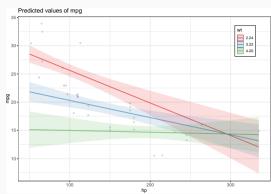
## AN INTERACTION MAY NOT BE SIG ACROSS THE ENTIRE RANGE OF THE PREDICTOR



## Let's see mpg ~ hp + wt

```
fit <- glm(mpg - hp*wt, data = mtcars)
pred <- ggpredict(fit, terms = c("hp", "wt"))
# plot(pred, add.data = TRUE) +
# theme_luis() +
# theme( legend.position = c(0.9, 0.85))
broom::tidy(fit) |> kable()
```

term	estimate	std.error	statistic	p.value
(Intercept)	49.808	3.605	13.82	0.000
hp	-0.120	0.025	-4.86	0.000
wt	-8.217	1.270	-6.47	0.000
hp:wt	0.028	0.007	3.75	0.001

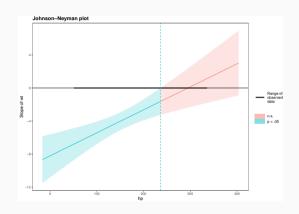


# Now, from what point the slope becomes non signficant?



#### JOHNSON-NEYMAN INTERVAL

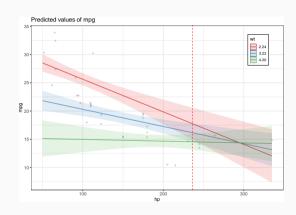
```
jn <- johnson_neyman(fit, wt, hp , plot = TRUE)
jn</pre>
```



## Now, from what point the slope becomes non signficant?



# JOHNSON-NEYMAN INTERVAL Overlayed on data

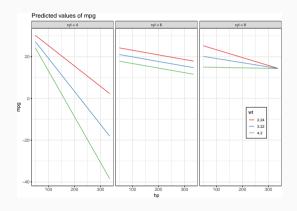


## **THREE-WAY INTERACTIONS**



```
fit <- glm(mpg - hp*wt*as.factor(cyl), data = mtcars)

dat <- ggpredict(fit, terms = c("hp", "wt", "cyl"))
plot(dat, ci = FALSE)+
    theme( legend.position = c(0.9, 0.85))</pre>
```



# **THE SUMMARY**



broom::tidy(fit) |> kable()

term	estimate	std.error	statistic	p.value
(Intercept)	34.972	10.883	3.213	0.004
hp	0.042	0.129	0.324	0.750
wt	0.054	5.010	0.011	0.991
as.factor(cyl)6	-2.340	77.278	-0.030	0.976
as.factor(cyl)8	5.829	42.608	0.137	0.893
hp:wt	-0.062	0.058	-1.068	0.298
hp:as.factor(cyl)6	-0.064	0.660	-0.098	0.923
hp:as.factor(cyl)8	-0.120	0.227	-0.527	0.604
wt:as.factor(cyl)6	-3.321	27.372	-0.121	0.905
wt:as.factor(cyl)8	-6.168	12.492	-0.494	0.627
hp:wt:as.factor(cyl)6	0.062	0.236	0.264	0.794
hp:wt:as.factor(cyl)8	0.080	0.078	1.026	0.317

# **MARGINAL EFFECTS**

#### MARGINAL EFFECTS



- Marginal effects: the effect of a predictor on the response, averaged over all values of the other predictors.
- also, the instantaneous effect of "x" on "y". Calculated with a derivative of x in respect to y, expressed mathematically as dy/dx.
  - the "instantaneous rate of change of y with respect to x."

## WHAT ARE THE MARGINAL EFFECTS FOR mpg ~ hp\*wt



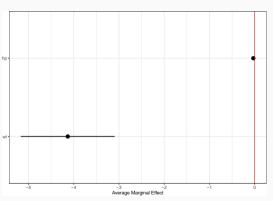
```
fit_m <- margins(fit2)
summary(fit_m)</pre>
```

```
factor AME SE z p lower upper hp -0.0305 0.0075 -4.0661 0.0000 -0.0452 -0.0158 wt -4.1316 0.5296 -7.8021 0.0000 -5.1696 -3.0937
```

• Interpretation: for each unit increase of mpg an average reduction of  ${\tt hp}$  3% is expected.

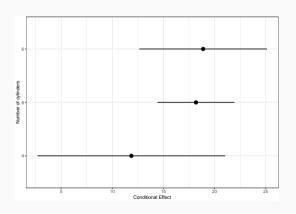
#### **PLOTTING THE MARGINAL EFFECTS**





## **CAN WE PLOT THE CONDITIONAL EFFECTS TOO?**





#### **ODDITIES OF INTERACTIONS IN LINEAR REGRESSIONS**



- At the beginning I said, keep the interaction that is significant, but<sup>1</sup>
  - In the transformation from log-odds to probabilities (AME) the interaction may not be significant anymore, or worse
  - The interaction may be significant in the AME, but not in conditional effects

<sup>&</sup>lt;sup>1</sup>Bruin, "Deciphering Interactions in Logistic Regression," *Introduction to SAS. UCLA: Statistical Consulting Group.*; Vanhove, 2019, "Interactions in Logistic Regression Models," (2019).

## Example when the conditional is not significant while the AME is



# speed on wt\*cyl

fit1 <- glm(qsec - wt\*as.factor(cyl), data = mtcars)
broom::tidy(fit1) |> kable()

term	estimate	std.error	statistic	p.value
(Intercept)	14.83	1.500	9.884	0.000
wt	1.89	0.639	2.951	0.007
as.factor(cyl)6	-9.78	4.395	-2.226	0.035
as.factor(cyl)8	-1.44	2.273	-0.632	0.533
wt:as.factor(cyl)6	2.26	1.464	1.546	0.134
wt:as.factor(cyl)8	-1.04	0.764	-1.360	0.185

summary(margins(fit1)) |> kable()

factor	AME	SE	z	р	lower	upper
cyl6	-2.50	0.825	-3.03	0.002	-4.12	-0.884
cyl8	-4.78	0.823	-5.81	0.000	-6.39	-3.169
wt	1.93	0.406	4.74	0.000	1.13	2.721

## IN OPENMX, MAXIMUM LIKELIHOOD



```
library(umx)

model <- "
    mpg ~ hp*hp + wt*wt
    moderation := hp*wt
"

m1 <- umxRAM(model, data = mtcars)
m1 <- umxCI(m1)
umxSummary(m1, std = T)</pre>
```

Table 10: Parameter loadings for model 'm1'

	name	Std.Estimate	Std.SE	CI	type
5	hp_with_wt	0.66	0.1	0.66 [0.46, 0.85]	Manifest Cov
1	hp	-0.36	0.1	-0.36 [-0.56, -0.17]	Manifest path
2	wt	-0.63	0.09	-0.63 [-0.81, -0.45]	Manifest path
7	one_to_mpg	6.28	0.7	6.28 [4.91, 7.64]	Mean
8	one_to_hp	2.17	0.32	2.17 [1.54, 2.81]	Mean
9	one_to_wt	3.34	0.45	3.34 [2.45, 4.23]	Mean
3	mpg_with_mpg	0.17	0.06	0.17 [0.06, 0.28]	Residual
4	hp_with_hp	1.00	0	1 [1, 1]	Residual
6	wt_with_wt	1.00	0	1 [1, 1]	Residual

Model Fit: Chi2(0) = 0, p = 1.000; CFI = 1; TLI = 1; RMSEA = 0 Algebra'moderation' = 0.123Cl95[0.074, 0.173]. p-value < 0.001

#### CONCLUSION



- We started reviewing multiple regression
- Then discussed the syntax and interpretation of parameters when an interaction term is included
- Finally, we discussed how to extract the marginal effects of the interaction term
- Luckly the package margins() makes this extremely simple.

## **ACKNOWLEDGEMENTS**



#### **Team**

- Charles Gardner (2015)
- Brad Verhulst (2013)
- Joshua Pritkin.
- Rob Kirkpatrick.
- Michael C Neale.
- NIH grant no R01 DA049867 and 5T32MH-020030

## Contact



# • THANK YOU