

Fecal matter transplant in Huntington's disease (Male) Data Analysis

Saritha Kodikara

January 5, 2022

Contents

E	cecut	tive Su	ımmary	2
1	Dat	a Inpu	ut	2
2	Lin	ear M	ixed Models - Repeated measures analysis	5
	2.1	LMM	for Body Weight, Weight Gain, Rotarod, Fecal Output and Fecal water content	3
	2.2	LMM	for Food and Water intake	5
	2.3	LMM	for CFC Conditioning and Extinction	6
3	Lin	ear M	ixed Models - One Time point analysis	ç
	3.1	Y-ma	ze	6
	3.2	Colon	otypic Variables (Gut Transit Time, Gut permeability, Cecum Weight, Cecum Length, Length, Y-maze difference, Y-maze index, Y-maze Dist trial1, NOR, Brain Weight, e Ratio)	10
	3.3	Metal	bolic Variables (Acetate, Propionate, Isobutyrate,2-Methylbutyrate)	14
		3.3.1	LMM with only main effects as the number of samples were less than four for HD-FMT group	14
		3.3.2	With all two way interactions (except FMT*ATB)	15
	3.4	Cytok	cines Variables	16
		3.4.1	LMM with only main effects due to low samples	16
		3.4.2	With all two way interactions (except FMT*ATB)	17
4	Cla	sping	score	20
	4.1	Ordin	al regression	24
	4.2	Ordin	al regression with random effect	25
	43	Comp	parison between Cumulative Link Model and Cumulative Link Mixed Model	26



5	Post	t-Hoc test	26
	5.1	Body Weight	26
	5.2	Weight Gain	27
	5.3	Fecal Water Content	27
	5.4	Food Intake	28
	5.5	Water Intake	29
	5.6	CFC Extinction	29
	5.7	Ymaze	30
	5.8	Brain Weight	30
	5.9	Propel Brake Ratio- Week 14	31
	5.10	FITC- Week 20	31
	5.11	CecumWeight- Week 14	32
	5.12	Colon Length- Week 20	32
	5.13	Acetate- Week 20	32
	5.14	IL17E- Week 20	33
6	Info	ermation about the used B session	34

Executive Summary

- For repeated measures, Linear Mixed Models (LMM) were fitted with time as a covariate.
- For one time point measures, Linear Mixed Models (LMM) were fitted with all two way interactions except for ATB*FMT (No observations for FMT and no ATB). However, only main effects were considered for variables that had few samples when covariates are considered together. For example IL17A-w14 had only 3 samples in HD-FMT group but when considered separately HD had 15 samples and FMT had 8 samples. Thus, IL17A-w14 was modeled without interactions.
- Clasping score was modeled using a cumulative link mixed model.
- For all models with significant effects on categorical variables (i.e, genotype, FMT, ATB), pairwise comparisons were performed using emmeans package.

1 Data Input

```
# load data from the csv file first. (the csv file is from your xlsx file)
data <- read_excel("Data/CarolGubert_FMT_male_data_July2021_final.xlsx")
data <- as.data.frame(data)
# extract group information
data$genotype <- factor(data$genotype,levels = c("WT", "HD"))
data$ATB<- factor(data$ATB,levels = c("NO", "YES"))
data$FMT<- factor(data$FMT,levels = c("NO", "YES"))
data$BOX <- factor(data$BOX)</pre>
Treatment<-ifelse((data$ATB=="YES"&data$FMT=="YES"),"ATB_FMT",
```



```
ifelse((data$ATB=="YES"&data$FMT=="NO"),"Only ATB","No treatment"))%>%
    factor(levels = c("No treatment","Only ATB","ATB_FMT"))
data.new<-add_column(data, Treatment = Treatment, .after = 5)</pre>
```

2 Linear Mixed Models - Repeated measures analysis

```
data.repeated.1<-data.new[,c(1:6,8:37,61:75,91:116)]
long_data.repeated.1</pre>
gather(v, value, BodyWeight_W6:FecalWaterContent_W20) %>%
separate(v, c("col", "Time"),sep="_W") %>%
arrange(ID) %>%
spread(col, value)
long_data.repeated.1$Time <-as.numeric(long_data.repeated.1$Time)
#filter out Time 6 and 7
long_data.repeated.1</pre>
long_data.repeated.1
cong_data.repeated.1%filter(Time!=6 &Time!=7)
long_data.repeated.1$Time<-long_data.repeated.1$Time-7
```

2.1 LMM for Body Weight, Weight Gain, Rotarod, Fecal Output and Fecal water content

```
#subset(long data.repeated.1, !is.na(BodyWeight)) %>%
    #group_by(genotype, FMT, ATB) %>%
    #tally()
#subset(long data.repeated.1, !is.na(BodyWeight)) %>%
    #group_by(genotype, Treatment, Time) %>%
    #tally()%>% summary()
#Interaction between ATB & FMT not considered as all no ATB had no FMT
BodyWeight.lmm <- lme(BodyWeight ~ genotype+Treatment+Time+</pre>
                         genotype:Treatment+genotype:Time+
                         Treatment:Time,random =(~1|BOX),
                         data = subset(long_data.repeated.1, !is.na(BodyWeight)))
WeightGain.lmm <- lme(WeightGain ~ genotype+Treatment+Time+</pre>
                         genotype:Treatment+genotype:Time+
                         Treatment:Time,random =(~1|BOX),
                         data = subset(long_data.repeated.1, !is.na(WeightGain)))
Rotarod.lmm <- lme(Rotarod ~ genotype+Treatment+Time+</pre>
                         genotype:Treatment+genotype:Time+
                         Treatment:Time,random =(~1|BOX),
                         data = subset(long_data.repeated.1, !is.na(Rotarod)))
FecalOutput.lmm <- lme(FecalOutput ~ genotype+Treatment+Time+
                         genotype:Treatment+genotype:Time+
                         Treatment:Time,random =(~1|BOX),
```



## ##		 =========			
##		1	Dependent va	riable:	
## ##		WeightGain (2)		FecalWaterContent (4)	FecalOutput (5)
## ## ## ##	genotypeHD			-14.1599 p = 0.1225	1.9431 p = 0.2412
## ## ##	J	-7.8967+ p = 0.0662		4.8862 p = 0.6306	-0.7293 p = 0.6924
## ## ##	TreatmentATB_FMT	-3.8837 p = 0.3519		1.3579 p = 0.8897	0.2536 p = 0.8867
## ## ##	Time	2.5328*** p = 0.0000			0.0298 p = 0.7119
## ## ##	<pre>genotypeHD:TreatmentOnly ATB</pre>	11.5724** p = 0.0100			3.6911+ p = 0.0569
## ## ##	genotypeHD:TreatmentATB_FMT	-1.2411 p = 0.8108		21.4059+ $p = 0.0926$	2.2866 p = 0.3023
## ## ##	genotypeHD:Time	-2.8962*** p = 0.0000		-0.2176 p = 0.5202	-0.6899*** p = 0.0000
## ## ##	TreatmentOnly ATB:Time	0.0886 p = 0.6414			0.0506 p = 0.5991
## ## ##	TreatmentATB_FMT:Time	0.2528 p = 0.1792			0.0780 p = 0.4151
## ## ##	Constant	108.6238*** p = 0.0000			12.4161*** p = 0.0000



```
1378
                               715 1346
                                                 710
## Observations
                                                           713
## Log Likelihood
                   -3198.6550 -2497.1390 -7412.5200 -3028.3720
                                                       -2007.4000
                   6421.3090 5018.2780 14849.0400
                                             6080.7430
## Akaike Inf. Crit.
                                                         4038.8000
                   6483.9620 5072.9760 14911.4100 6135.3560
## Bayesian Inf. Crit.
                                                         4093.4640
## Note:
                                    + p<0.1; * p<0.05; ** p<0.01; *** p<0.001
```

2.2 LMM for Food and Water intake

```
FoodIntake.data<-data.new[,c(1:6,38:49)]
long_FoodIntake.data-FoodIntake.data %>%
  gather(v, value, FoodIntake_W9:FoodIntake_W20) %>%
  separate(v, c("col", "Time"),sep="_W") %>%
  arrange(ID) %>%
  spread(col, value)
long_FoodIntake.data$Time <-as.numeric(long_FoodIntake.data$Time)</pre>
long_FoodIntake.data$Time<-long_FoodIntake.data$Time-8</pre>
WaterIntake.data<-data.new[,c(1:6,50:60)]
long_WaterIntake.data<-WaterIntake.data %>%
  gather(v, value, WaterIntake_W10:WaterIntake_W20) %>%
  separate(v, c("col", "Time"), sep="_W") %>%
 arrange(ID) %>%
  spread(col, value)
long_WaterIntake.data$Time <-as.numeric(long_WaterIntake.data$Time)</pre>
long_WaterIntake.data$Time<-long_WaterIntake.data$Time-9</pre>
```



##	genotypeHD	0.1195* p = 0.0135	0.2391 $p = 0.1308$
## ## ## ##	TreatmentOnly ATB	0.2039*** p = 0.0001	0.5717*** p = 0.0010
	TreatmentATB_FMT	0.2070*** p = 0.0004	0.3296+ $p = 0.0578$
	Time	-0.0290*** p = 0.0000	-0.0469*** p = 0.00005
##	genotypeHD:TreatmentOnly ATB	-0.2220*** p = 0.000005	-0.1428 p = 0.4277
##	genotypeHD:TreatmentATB_FMT	-0.2221*** p = 0.0002	0.0375 p = 0.8518
##	genotypeHD:Time	-0.0034 p = 0.4308	0.0569*** p = 0.000001
##	TreatmentOnly ATB:Time	-0.0020 p = 0.6974	-0.0333* p = 0.0146
##	TreatmentATB_FMT:Time	-0.0047 p = 0.3776	-0.0093 p = 0.4904
## ## ## ##	Constant	0.9929*** p = 0.0000	1.2985*** p = 0.0000
## ## ##	Observations Log Likelihood Akaike Inf. Crit. Bayesian Inf. Crit.	324 154.5813 -285.1626 -240.1699	296 -103.4265 230.8530 274.7249
##	Note:	=======================================	

2.3 LMM for CFC Conditioning and Extinction

```
CFC.data<-data.new[,c(1:6,165:171)]
long_CFC.data</pre>
gather(v, value, CONDT_1:CONDT_6) %>%
separate(v, c("col", "Time"),sep="_") %>%
arrange(ID) %>%
spread(col, value)
long_CFC.data$Time <-as.numeric(long_CFC.data$Time)

CFC.ex.data<-data.new[,c(1:6,172:181)]
long_CFC.ex.data<-CFC.ex.data %>%
gather(v, value, `EXT_1-5`:`EXT_41-45`) %>%
separate(v, c("col", "Time"),sep="_") %>%
```



##

```
arrange(ID) %>%
  spread(col, value)
long_CFC.ex.data$Time <- factor(long_CFC.ex.data$Time,ordered = TRUE,</pre>
                                     levels = c('1-5', '6-10',
                                                 '11-15', '16-20',
                                                 '21-25', '26-30',
                                                 '31-35', '36-40',
                                                 '41-45'))
long_CFC.ex.data$Time = as.numeric(long_CFC.ex.data$Time)
CFC_Conditioning.lmm <- lme(CONDT ~ genotype+Treatment+Time+CONDT_Base+
                         genotype:Treatment+genotype:Time+genotype:CONDT_Base+
                         Treatment:Time+Treatment:CONDT_Base+Time:CONDT_Base,
                         random=(~1|BOX),data =subset(long_CFC.data,!is.na(CONDT)))
CFC_Extinction.lmm <- lme(EXT ~ genotype+Treatment+Time+EXT_Base+</pre>
                         genotype:Treatment+genotype:Time+genotype:EXT_Base+
                         Treatment:Time+Treatment:EXT_Base+Time:EXT_Base,
                         random=(~1|BOX),data =subset(long_CFC.ex.data,!is.na(EXT)))
stargazer(CFC_Conditioning.lmm,CFC_Extinction.lmm, type = "text",
          digits = 4,
          report = ('vc*p'),
          star.char = c("+", "*", "**", "***"),
          star.cutoffs = c(.1, .05, .01, .001),
          digit.separator = "",
          notes = c("+ p<0.1; * p<0.05; ** p<0.01; *** p<0.001"),
          notes.append = FALSE)
```

## =========== ##		Dependent variable:			
##					
##	CONDT	EXT			
##	(1)	(2)			
##					
## genotypeHD	-0.9536	-15.7142**			
##	p = 0.9120	p = 0.0025			
##					
## TreatmentOnly ATB	3.1574	2.2257			
##	p = 0.7355	p = 0.6633			
##					
## TreatmentATB_FMT	3.2605	8.1952			
##	p = 0.7353	p = 0.1311			
##					
## Time	15.7786***	-1.9155***			
##	p = 0.0000	p = 0.0002			
##					
## CONDT_Base	4.4344***				
##	p = 0.0002				
##					
## EXT_Base		1.0142***			
##		p = 0.0000			



##			
	<pre>genotypeHD:TreatmentOnly ATB</pre>	-1.3767	-0.7753
##		p = 0.8802	p = 0.8935
## ##	genotypeHD:TreatmentATB_FMT	2.5462	-9.6438
##	-	p = 0.7870	p = 0.1087
##	monotonoUD. Timo	1.0692	2.6709***
## ##	genotypeHD:Time	p = 0.4813	p = 0.000000
##		•	1
	genotypeHD:CONDT_Base	0.2661	
## ##		p = 0.7985	
##	genotypeHD:EXT_Base		0.1023
##			p = 0.1881
## ##	TreatmentOnly ATB:Time	-0.4149	-0.2114
##		p = 0.8200	p = 0.7130
##	The state of the s	0.4007	0.0000
## ##	TreatmentATB_FMT:Time	-0.1887 p = 0.9198	-0.6628 p = 0.2583
##		P 0.0100	P 0.2000
	TreatmentOnly ATB:CONDT_Base	0.2672	
## ##		p = 0.8551	
##	TreatmentATB_FMT:CONDT_Base	-2.6367*	
##		p = 0.0133	
## ##	Time:CONDT_Base	-0.2468+	
##	-	p = 0.0835	
##	TrontmontOnler ATD.EVT Dogo		0.3060***
## ##	TreatmentOnly ATB:EXT_Base		-0.3969*** p = 0.000001
##			1
	TreatmentATB_FMT:EXT_Base		0.1283
## ##			p = 0.2153
##	Time:EXT_Base		-0.0703***
##			p = 0.0000
## ##	Constant	-19.9046**	19.2334***
##		p = 0.0064	p = 0.000003
## ##			
	Observations	522	783
	Log Likelihood	-2491.0010	-3354.7000
	Akaike Inf. Crit.	5016.0030	6743.4000
	Bayesian Inf. Crit.	5087.8880 ===========	6822.3440
	Note:		** p<0.01; *** p<0.00



3 Linear Mixed Models - One Time point analysis

3.1 Y-maze

```
##
  _____
##
                                      Dependent variable:
##
##
                                            Ymaze
                                          -17.1598*
##
  genotypeHD
##
                                         p = 0.0431
##
## TreatmentOnly ATB
                                           -4.0338
##
                                          p = 0.6500
##
## TreatmentATB_FMT
                                           8.2060
##
                                         p = 0.3454
##
## ArmFamiliar
                                           -2.8522
##
                                         p = 0.7350
##
  genotypeHD:TreatmentOnly ATB
                                           14.2992
##
                                         p = 0.1543
  genotypeHD:TreatmentATB_FMT
                                           2.7253
##
##
                                          p = 0.7797
```



```
##
  genotypeHD:ArmFamiliar
                                                    0.8503
##
                                                  p = 0.9136
##
## TreatmentOnly ATB:ArmFamiliar
                                                  -10.6377
##
                                                 p = 0.2800
##
## TreatmentATB FMT:ArmFamiliar
                                                   -15.3970
##
                                                  p = 0.1141
##
## Constant
                                                  73.8479***
##
                                                  p = 0.0000
##
## Observations
                                                      170
## Log Likelihood
                                                   -759.3475
## Akaike Inf. Crit.
                                                   1544.6950
## Bayesian Inf. Crit.
                                                   1584.6720
## Note:
                                  + p<0.1; * p<0.05; ** p<0.01; *** p<0.001
```

3.2 Phenotypic Variables (Gut Transit Time, Gut permeability, Cecum Weight, Cecum Length, ColonLength, Y-maze difference, Y-maze index, Y-maze Dist trial1, NOR, Brain Weight, Brake Ratio)

```
dep_vars <-c("Ymaze_index", "Ymaze_Dist_trial1", "NOR_RI_5min",</pre>
             "BrainWeight", "Digigait PropelBrakeRatio W14",
             "GutTransitTime_W14", "GutTransitTime_W20", "FITC_w14",
             "FITC w20", "CecumWeight w14", "CecumWeight w20",
             "CecumLength_w14", "CecumLength_w20",
             "ColonLength_w14", "ColonLength_w20")
p_val<-lapply(dep_vars, function(r) {</pre>
  m <- lme(formula(paste(r, "genotype+Treatment+genotype:Treatment", sep = "~")),</pre>
           random=(~1|BOX),data = subset(data.new,!is.na(eval(parse(text = r)))))
  assign(paste0(r,".lmm_"),m, envir = .GlobalEnv)
  summary(m)$tTable[,5]})
stargazer(Ymaze_index.lmm_,Ymaze_Dist_trial1.lmm_,
          NOR_RI_5min.lmm_,BrainWeight.lmm_,Digigait_PropelBrakeRatio_W14.lmm_,
          type = "text",
          digits = 4,
          report = ('vc*p'),
          star.char = c("+", "*", "**", "***"),
          star.cutoffs = c(.1, .05, .01, .001),
          digit.separator = "",
          notes = c("+ p<0.1; * p<0.05; ** p<0.01; *** p<0.001"),
          notes.append = FALSE,
          dep.var.labels.include = FALSE,
          column.labels=c("YmazeIndex","YmazeDist","NOR",
```



```
"BrainWeight", "PropelBrakeRatio"),
align=TRUE)
```

```
______
##
                                        Dependent variable:
##
##
                        YmazeIndex YmazeDist
                                             NOR
                                                   BrainWeight PropelBrakeRatio
##
                         (1)
                                  (2)
                                             (3)
                                                    (4)
                        -0.2238 -3097.5080 -0.9825 -0.0700**
                                                             1.4822**
  genotypeHD
                       p = 0.1310 p = 0.4623 p = 0.8634 p = 0.0033 p = 0.0044
##
## TreatmentOnly ATB
                        0.0526
                                -501.3699 0.7903 -0.0075
                                                              0.5922
##
                       p = 0.7190 p = 0.9005 p = 0.8882 p = 0.7280
                                                              p = 0.2368
                        0.0870
                                1392.4650 4.5609 0.0336
## TreatmentATB FMT
                                                              0.2232
                        p = 0.5483 p = 0.7601 p = 0.4261 p = 0.1181
##
                                                             p = 0.6388
##
## genotypeHD:TreatmentOnly ATB 0.1328 929.3390 -7.6252 0.0190
                                                              -1.8307*
                        p = 0.5218 p = 0.8743 p = 0.3359 p = 0.5240
##
                                                              p = 0.0110
##
                                                   0.0056
##
  genotypeHD:TreatmentATB_FMT 0.2237
                                2227.8640 -8.0220
                                                              -0.4834
##
                        p = 0.2801 p = 0.7183 p = 0.3154 p = 0.8512
                                                              p = 0.4914
##
                        1.1261*** 27415.8500*** 59.8142*** 0.4275***
                                                             3.4404***
## Constant
                                                             p = 0.0000
##
                        p = 0.0000 p = 0.0000 p = 0.0000 p = 0.0000
##
                         94
## Observations
                                84 92 52
                                                                 93
## Log Likelihood
                       -36.8822 -830.9299 -364.6142 73.4326
                                                             -157.8347
                                1677.8600 745.2284 -130.8652
## Akaike Inf. Crit.
                       89.7644
                                                              331.6694
                       109.5831
## Bayesian Inf. Crit.
                                 1696.7140 764.8632 -116.2361
                                                              351.3967
## Note:
                                         + p<0.1; * p<0.05; ** p<0.01; *** p<0.001
```



align=TRUE)

```
##
                                            Dependent variable:
##
                           GutTransitTime-W14 GutTransitTime-W20 FITC-W14 FITC-W20
##
                                         (2)
##
                                          -12.2482 0.0751 0.5967*
p = 0.8391 p = 0.8132 p = 0.0250
                              36.0361
## genotypeHD
                              p = 0.2631
##
##
## TreatmentOnly ATB
                              11.7028
                                              -60.2482
                                                           -0.3718 -0.0200
##
                              p = 0.7143
                                              p = 0.3242
                                                           p = 0.2326 p = 0.9293
##
                                                           -0.5159 -0.0348
## TreatmentATB FMT
                                9.2306
                                               31.7518
                                              p = 0.5997
                                                           p = 0.1303 p = 0.8775
##
                              p = 0.7487
##
  genotypeHD:TreatmentOnly ATB
                              -52.6194
                                               92.7482
                                                            0.2728 -0.3062
                                             92.7482
p = 0.2775
##
                              p = 0.2533
                                                           p = 0.5377 p = 0.3657
##
  genotypeHD:TreatmentATB_FMT
                              -46.6216
                                               2.1232
                                                            -0.0355
##
                                                                      0.2578
                                             p = 0.9799
##
                              p = 0.2771
                                                           p = 0.9399 p = 0.4377
##
                                                           0.9959*** 0.4059*
## Constant
                             154.3806***
                                             271.7482***
                                           p = 0.000003 p = 0.0002 p = 0.0262
##
                              p = 0.0000
##
  ______
                                                             41
## Observations
                                 87
                                                 47
                                                                        30
                              -461.5515
## Log Likelihood
                                             -256.1310
                                                            -28.6413 -8.7940
## Akaike Inf. Crit.
                              939.1031
                                                            73.2826
                                              528.2621
                                                                      33.5880
## Bayesian Inf. Crit.
                              958.2587
                                              541.9707
                                                           85.7254
## Note:
                                          + p<0.1; * p<0.05; ** p<0.01; *** p<0.001
stargazer(CecumWeight_w14.lmm_,CecumWeight_w20.lmm_,CecumLength_w14.lmm_,
        CecumLength_w20.1mm_,
        type = "text",
        digits = 4,
        report = ('vc*p'),
        star.char = c("+", "*", "**", "***"),
        star.cutoffs = c(.1, .05, .01, .001),
        digit.separator = "",
        notes = c("+ p<0.1; * p<0.05; ** p<0.01; *** p<0.001"),
        notes.append = FALSE,
        dep.var.labels.include = FALSE,
        column.labels=c("CecumWeight-W14", "CecumWeight-W20",
                      "CecumLength-W14", "CecumLength-W20"),
```

```
##
## -----
## Dependent variable:
##

CecumWeight-W14 CecumWeight-W20 CecumLength-W14 CecumLength-W20
##

(1)
(2)
(3)
(4)
```



```
-0.0697
## genotypeHD
                           -0.0593
                                      0.0061
                                                                -0.1464
##
                          p = 0.1218
                                      p = 0.9154
                                                 p = 0.7425
                                                              p = 0.5702
##
                           -0.0300
                                      0.0893
                                                   0.0250
## TreatmentOnly ATB
                                                                -0.3250
##
                         p = 0.4047
                                      p = 0.1339
                                                 p = 0.9042
                                                              p = 0.1993
##
## TreatmentATB FMT
                          -0.0850*
                                      0.0102
                                                   -0.0208
                                                                0.3583
                                      p = 0.8559 p = 0.9260
##
                         p = 0.0393
                                                              p = 0.1469
##
## genotypeHD:TreatmentOnly ATB
                          0.0493
                                      -0.0649
                                                   -0.2178
                                                               0.1464
##
                          p = 0.3431
                                      p = 0.4092
                                                  p = 0.4667
                                                              p = 0.6752
##
## genotypeHD:TreatmentATB FMT
                          0.0373
                                      0.0921
                                                   -0.1770
                                                               -0.2869
##
                          p = 0.5138
                                    p = 0.2456
                                                 p = 0.5878
                                                              p = 0.4127
##
                          0.4250***
                                     0.4857***
                                                  2.7375***
                                                              2.9750***
## Constant
                                    p = 0.0000
##
                         p = 0.0000
                                                 p = 0.0000
                                                             p = 0.0000
##
## -----
## Observations
                                        51
## Log Likelihood
                          38.8129
                                      30.4576
                                                   -16.4150
                                                               -38.2558
## Akaike Inf. Crit.
                          -61.6257
                                      -44.9151
                                                   48.8300
                                                               92.5117
                         -48.9576
## Bayesian Inf. Crit.
                                      -30.4618
                                                  61.4982
                                                               106.9650
## Note:
                                         + p<0.1; * p<0.05; ** p<0.01; *** p<0.001
```

	Dependent v	variable:
_	Calanian + 1114	
	O	ColonLength-W20 (2)
	(1)	
genotypeHD	-0.3969	-1.2244**
8	p = 0.3566	p = 0.0085
	•	•
TreatmentOnly ATB	-0.0750	-0.7750+
	p = 0.8561	p = 0.0740
TreatmentATB_FMT	-0.3458	-0.3600
	p = 0.4427	p = 0.3785
	·	$p = 0.3566$ TreatmentOnly ATB -0.0750 $p = 0.8561$ TreatmentATB_FMT -0.3458



```
##
## genotypeHD:TreatmentOnly ATB
                         0.1469
                                         0.7761
##
                         p = 0.8044
                                       p = 0.1917
##
## genotypeHD:TreatmentATB_FMT
                         0.2493
                                         0.3441
##
                         p = 0.7031
                                       p = 0.5558
##
## Constant
                          8.9625***
                                       8.6250***
##
                         p = 0.0000
                                        p = 0.0000
##
## -----
## Observations
                           42
                                           51
## Log Likelihood
                        -45.5742
                                        -60.2668
## Akaike Inf. Crit.
                        107.1483
                                        136.5335
                      119.8165
                                     150.9868
## Bayesian Inf. Crit.
## Note:
                      + p<0.1; * p<0.05; ** p<0.01; *** p<0.001
```

- 3.3 Metabolic Variables (Acetate, Propionate, Isobutyrate, 2-Methylbutyrate)
- 3.3.1 LMM with only main effects as the number of samples were less than four for HD-FMT group

```
#subset(data.new, !is.na(Acetate w14)) %>%
    #group_by(genotype, Treatment) %>%
    #tally()
dep_vars <-c("Acetate_w14", "Propionate_w14",</pre>
             "Isobutyrate_w14", "Methylbutyrate_w14")
p_val<-lapply(dep_vars, function(r) {</pre>
  m <- lme(formula(paste(r, "genotype+Treatment", sep = "~")),</pre>
           random=(~1|BOX),data = subset(data.new,!is.na(eval(parse(text = r)))))
  assign(paste0(r,".lmm"),m, envir = .GlobalEnv)
  summary(m)$tTable[,5]})
stargazer(Acetate_w14.lmm,Propionate_w14.lmm,
          Isobutyrate_w14.lmm, Methylbutyrate_w14.lmm,
          type = "text",
          digits = 4,
          report = ('vc*p'),
          star.char = c("+", "*", "**", "***").
          star.cutoffs = c(.1, .05, .01, .001),
          digit.separator = "",
          notes = c("+ p<0.1; * p<0.05; ** p<0.01; *** p<0.001"),</pre>
          notes.append = FALSE,
          dep.var.labels.include = FALSE,
          column.labels=c("Acetate-14", "Propionate-14",
                          "Isobutyrate-14", "Methylbutyrate-14"),
          align=TRUE)
```



```
##
##
                                       Dependent variable:
##
                     Acetate-14 Propionate-14 Isobutyrate-14 Methylbutyrate-14
##
##
                      (1) (2) (3) (4)
                    29.5120* 0.9088 0.5904 0.1806
## genotypeHD
                    p = 0.0425 p = 0.5602 p = 0.2188 p = 0.2247
##
##
                    4.7346 1.7019 0.3049
p = 0.7555 p = 0.3241 p = 0.5616
## TreatmentOnly ATB 4.7346
                                                              -0.1399
##
                                                            p = 0.3874
##
## TreatmentATB_FMT
                    11.0272 -1.5247
                                              -0.3897
                                                              -0.2389
                     11.0272 -1.5247 -0.3897 -0.2389 p = 0.5474 p = 0.4770 p = 0.5441 p = 0.2377
##
##
               64.8784*** 2.9463*
                                            2.5912***
                                                             0.7841***
## Constant
                     p = 0.0001 p = 0.0469 p = 0.00002
                                                            p = 0.00002
##
## Observations 36 36 35

## Log Likelihood -161.9543 -97.9223 -55.1535

## Akaike Inf. Crit. 335.9086 207.8446 122.3070

## Bayesian Inf. Crit. 344.7030 216.6390 130.9109
                                                              -20.7752
                                                             53.5503
                                                              62.3447
## Note:
                                    + p<0.1; * p<0.05; ** p<0.01; *** p<0.001
```

3.3.2 With all two way interactions (except FMT*ATB)

```
dep_vars <-c("Acetate_w20", "Propionate_w20", "Isobutyrate_w20",</pre>
             "Methylbutyrate w20")
p_val<-lapply(dep_vars, function(r) {</pre>
  m <- lme(formula(paste(r, "genotype+Treatment+genotype:Treatment", sep = "~")),</pre>
           random=(~1|BOX),data = subset(data.new,!is.na(eval(parse(text = r)))))
  assign(pasteO(r,".lmm"),m, envir = .GlobalEnv)
  summary(m)$tTable[,5]})
stargazer(Acetate_w20.lmm,Propionate_w20.lmm,
          Isobutyrate_w20.lmm, Methylbutyrate_w20.lmm,
          type = "text",
          digits = 4,
          report = ('vc*p'),
          star.char = c("+", "*", "**", "***").
          star.cutoffs = c(.1, .05, .01, .001),
          digit.separator = "",
          notes = c("+ p<0.1; * p<0.05; ** p<0.01; *** p<0.001"),
          notes.append = FALSE,
          dep.var.labels.include = FALSE,
          column.labels=c("Acetate-20", "Propionate-20",
                           "Isobutyrate-20", "Methylbutyrate-20"),
```



align=TRUE)

## ##		Depende	ent variable:	
##		-		Methylbutyrate-20
##	(1)	(2)	(3)	(4)
## genotypeHD	-9.5017	-0.0525	0.0254	0.2137
##			p = 0.9763	p = 0.7970
##				
## TreatmentOnly ATB		0.2268	0.7819	1.4797+
##	p = 0.5463	p = 0.9496	p = 0.3360	p = 0.0644
## TreatmentATB_FMT	-20.1300	-7.4825+	1.1532	-0.1066
##		p = 0.0945		p = 0.8859
##	•	•	•	•
$\verb ## genotypeHD:TreatmentOnly ATB \\$			1.4744	-0.9962
##	p = 0.0435	p = 0.2574	p = 0.2282	p = 0.3887
<pre>## ## genotypeHD:TreatmentATB_FMT</pre>	39 6838	10.5947+	0.6947	0.3270
##		p = 0.0990		
##	1	1	1	1
## Constant	70.8392***	10.4182**	2.6521***	0.6934
##	p = 0.00004	p = 0.0011	p = 0.0003	p = 0.1986
## ##				
## Observations	32	 26	32	 32
## Log Likelihood	-129.3232		-44.8358	-47.8591
## Akaike Inf. Crit.	274.6464	151.5456	105.6716	111.7182
## Bayesian Inf. Crit.	284.7112	159.5114	115.7364	121.7829

3.4 Cytokines Variables

3.4.1 LMM with only main effects due to low samples



```
##
Dependent variable:
                 ______
##
                 IL17A-14 IL17E-14 IL21-14 IL7R-14
##
                  (1) (2) (3) (4)
                0.0589 0.4301 -0.0558 1.0230*
## genotypeHD
##
               p = 0.6692 p = 0.1134 p = 0.4973 p = 0.0162
##
## TreatmentOnly ATB -0.1186 0.1212 -0.0518 -0.5898
##
                p = 0.4588 p = 0.6958 p = 0.5905 p = 0.2102
##
## TreatmentATB_FMT
                 -0.0928 0.3462 0.1637 -0.2757
                p = 0.5949 p = 0.2649 p = 0.1212 p = 0.5620
##
##
## Constant
                 0.2550+ 0.5904* 0.3083*** 0.7143+
##
               p = 0.0615 p = 0.0206 p = 0.0009 p = 0.0565
##
## Observations 31 26 31
                                            25
## Log Likelihood -12.6930 -21.9144 3.9388 -28.7985
## Akaike Inf. Crit. 37.3860 55.8288 4.1224 69.5970
## Bayesian Inf. Crit. 45.1610 62.3750 11.8975
                                          75.8642
## Note:
                   + p<0.1; * p<0.05; ** p<0.01; *** p<0.001
```

3.4.2 With all two way interactions (except FMT*ATB)



```
##
##
                                             Dependent variable:
##
                           IFNg-14
                                             IL22-14
                                                       IL6-14
                                                               TNFa-14
##
                                    IL1b-14
                                                                        IFNg-20
##
                           (1)
                                    (2)
                                             (3)
                                                       (4)
                                                               (5)
                                                                        (6)
                          -0.0100
                                  -0.0718
                                                               0.3622
                                           -0.0097
                                                    -0.2248
                                                                        -0.0247+
  genotypeHD
                          p = 0.4778 p = 0.7599 p = 0.9651 p = 0.3504 p = 0.9964 p = 0.0591
##
##
                                   0.2181 -0.0400
                                                      0.0233
                                                              -0.0461
## TreatmentOnly ATB
                          -0.0015
                                                                        -0.0107
                         p = 0.9147 p = 0.3593 p = 0.8577 p = 0.9219 p = 0.9996 p = 0.4101
##
##
                                    0.1976
                                            0.1203
                                                     -0.0779 0.3905
## TreatmentATB FMT
                          -0.0030
                          p = 0.8382 p = 0.4268 p = 0.6081 p = 0.7546 p = 0.9964 p = 0.1501
##
##
##
  genotypeHD:TreatmentOnly ATB 0.0175
                                   -0.1764
                                             -0.1169
                                                      0.1759
                                                               0.2442
##
                          p = 0.3724 p = 0.5900 p = 0.7061 p = 0.5949 p = 0.9983 p = 0.1869
                                                      0.6705+
  genotypeHD:TreatmentATB_FMT
                           0.0245
                                    0.0438
                                             -0.0483
                                                               150.3674 -0.0180
##
                          p = 0.2475 p = 0.8999 p = 0.8875 p = 0.0803 p = 0.2201 p = 0.3410
##
##
## Constant
                           0.0362** 0.3428+ 0.5447** 0.8044*** 0.8442 0.0405***
                          p = 0.0021 p = 0.0529 p = 0.0032 p = 0.0003 p = 0.9876 p = 0.0003
##
##
## ------
                             35
                                      35
                                              34
                                                        34
                                                                 33
                                                                          37
## Observations
                          61.9874 -19.8067 -17.7358 -19.5024 -174.8050 67.8828
## Log Likelihood
                                  55.6134
                                           51.4717 55.0047
## Akaike Inf. Crit.
                         -107.9749
                                                               365.6099 -119.7656
## Bayesian Inf. Crit.
                          -97.0365
                                   66.5518 62.1293 65.6624
                                                               375.9766 -108.2937
## Note:
                                             + p<0.1; * p<0.05; ** p<0.01; *** p<0.001
```



```
##
                                      Dependent variable:
##
##
                          IL17A-20 IL17E-20 IL1b-20 IL21-20 IL22-20
##
                           (1) (2) (3)
                                                      (4)
                          -0.0107 0.1313 0.1719 -0.2490 -0.0978
## genotypeHD
                         p = 0.8701 p = 0.5654 p = 0.7236 p = 0.1524 p = 0.7838
##
## TreatmentOnly ATB
                          -0.0146
                                  -0.0321 0.3688 -0.1318
##
                          p = 0.8388 p = 0.8612 p = 0.4511 p = 0.3863 p = 0.5895
##
                                                      0.0579
                          0.0685 0.4044* -0.1591
## TreatmentATB_FMT
                          p = 0.3454 p = 0.0432 p = 0.7535 p = 0.7138 p = 0.7868
##
##
## genotypeHD:TreatmentOnly ATB 0.0787 -0.1703 0.3263 0.1306 0.5237
                          p = 0.4319 p = 0.5683 p = 0.6426 p = 0.5842 p = 0.3159
##
##
## genotypeHD:TreatmentATB_FMT 0.0824 -0.5396+ 0.1651 0.1002
                          p = 0.4164 p = 0.0900 p = 0.8172 p = 0.6710 p = 0.4593
##
                           0.1448** 0.6645*** 0.6860+ 0.7783*** 1.1129***
## Constant
                          p = 0.0062 p = 0.0001 p = 0.0581 p = 0.000003 p = 0.0004
##
## ------
                        35 33 39 30 39
17.3611 -11.4072 -47.8369 -6.1180 -38.4919
-18.7222 38.8143 111.6737 28.2359 92.9839
## Observations
## Log Likelihood
## Akaike Inf. Crit.
## Bayesian Inf. Crit.
                          -7.7838 49.1810 123.6458 37.6604
+ p<0.1; * p<0.05; ** p<0.01; *** p<0.001
stargazer(IL6_w20.lmm, TNFa_w20.lmm,
       type = "text",
        digits = 4,
        report = ('vc*p'),
        star.char = c("+", "*", "**", "***"),
        star.cutoffs = c(.1, .05, .01, .001),
        digit.separator = "",
```

notes = c("+ p<0.1; * p<0.05; ** p<0.01; *** p<0.001"),



```
notes.append = FALSE,
dep.var.labels.include = FALSE,
column.labels=c("IL6-20","TNFa-20"),
align=TRUE)
```

##						
##		Dependent variable:				
## ##		IL6-20	TNFa-20			
##		(1)	(2)			
	genotypeHD	-0.2050	-0.9973			
##	<u> </u>	p = 0.7827	p = 0.5521			
##	m	0.0000	0.7000			
##	TreatmentOnly ATB	0.0898 $p = 0.9038$	-0.7883 p = 0.6377			
##		p 0.3000	p 0.0077			
##	TreatmentATB_FMT	1.0858	-0.8374			
##		p = 0.1762	p = 0.6308			
##	<pre>genotypeHD:TreatmentOnly ATB</pre>	0.3921	0.6315			
##	genetypenz: II edomentenzy miz	p = 0.7163	p = 0.7933			
##						
	genotypeHD:TreatmentATB_FMT	-0.3485	4.3945+			
##		p = 0.7500	p = 0.0956			
	Constant	0.8100	1.7749			
##		p = 0.1356	p = 0.1458			
##						
##	Observations	39	38			
	Log Likelihood	-59.1479	-86.8274			
##	Akaike Inf. Crit.	134.2957	189.6548			
	Bayesian Inf. Crit.	146.2678	201.3807			
	Note:	+ p<0.1; * p<0.05; *	* p<0.01; *** p<0.001			

4 Clasping score

```
data_clasping <- data.new[,c(1:6,78:90)] # extract the columns of data that will become Y
long_clasping<-data_clasping %>%
  gather(v, value, Clasping_W8:Clasping_W20) %>%
  separate(v, c("col", "Time"),sep="_W") %>%
  arrange(ID) %>%
  spread(col, value)
long_clasping$Time <-as.numeric(long_clasping$Time)-7
long_clasping$Clasping<-factor(long_clasping$Clasping, ordered = TRUE , levels = c(0:4))
t<-table(long_clasping$Time,long_clasping$Clasping)
t1<-table(long_clasping$Time,long_clasping$Clasping)%>%prop.table(margin = 1)
```

#subset(long_clasping, !is.na(Clasping)) %>%
#group_by(genotype, Treatment, Time) %>%
#tally() %>% summary()

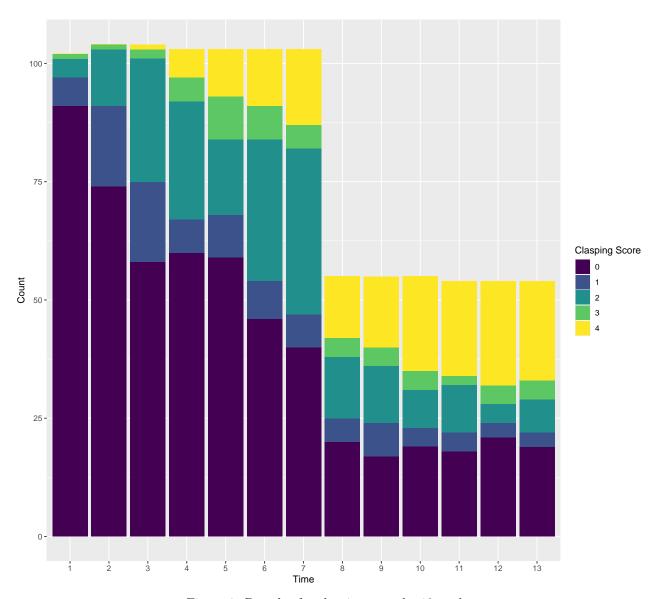


Figure 1: Bar plot for clasping over the 12 weeks

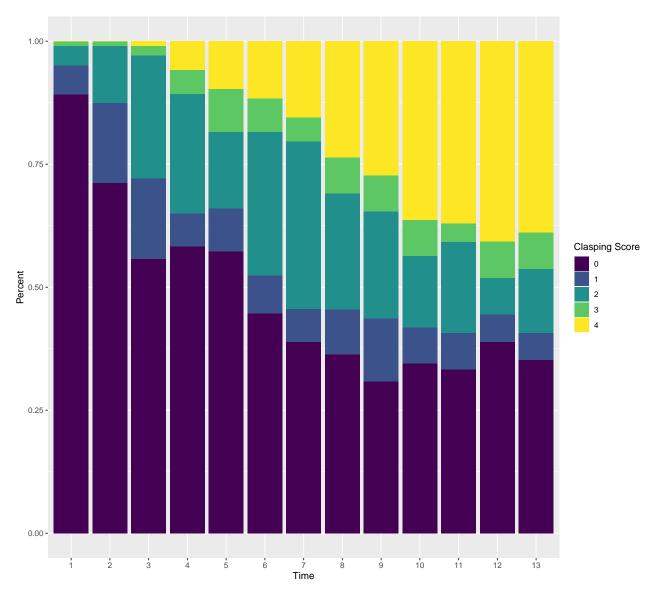


Figure 2: Bar plot for clasping over the 12 weeks

```
# HD mice clasping score
long_clasping_hd<-data_clasping %>%
  gather(v, value, Clasping_W8:Clasping_W20) %>%
  separate(v, c("col", "Time"),sep="_W") %>%
  arrange(ID) %>%
  spread(col, value)%>%filter(genotype=="HD")
long_clasping_hd$Time <-as.numeric(long_clasping_hd$Time)-7
long_clasping_hd$Clasping<-factor(long_clasping_hd$Clasping, ordered = TRUE , levels = c(0:4))
t1<-table(long_clasping_hd$Time,long_clasping_hd$Clasping)%>%prop.table(margin = 1)
```

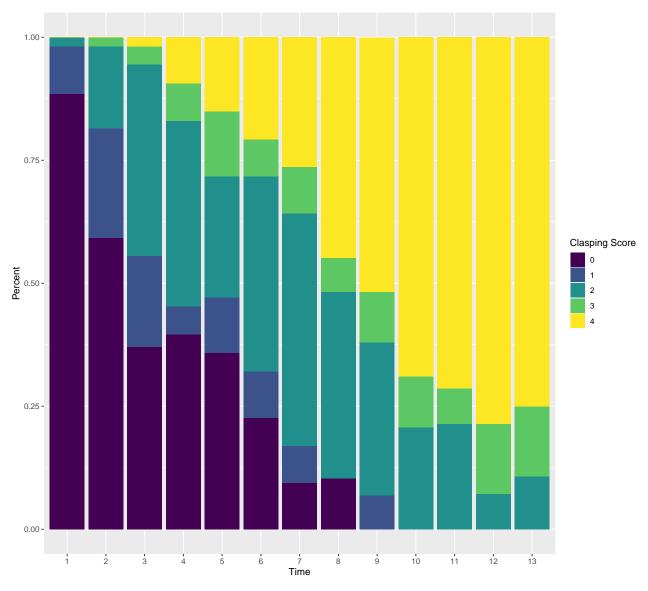


Figure 3: Bar plot for clasping over the 12 weeks for HD mice

```
# WT mice clasping score
long_clasping_wt<-data_clasping %>%
  gather(v, value, Clasping_W8:Clasping_W20) %>%
  separate(v, c("col", "Time"),sep="_W") %>%
  arrange(ID) %>%
  spread(col, value)%>%filter(genotype=="WT")
long_clasping_wt$Time <-as.numeric(long_clasping_wt$Time)-7
long_clasping_wt$Clasping<-factor(long_clasping_wt$Clasping, ordered = TRUE , levels = c(0:4))
t1<-table(long_clasping_wt$Time,long_clasping_wt$Clasping)%>%prop.table(margin = 1)
```

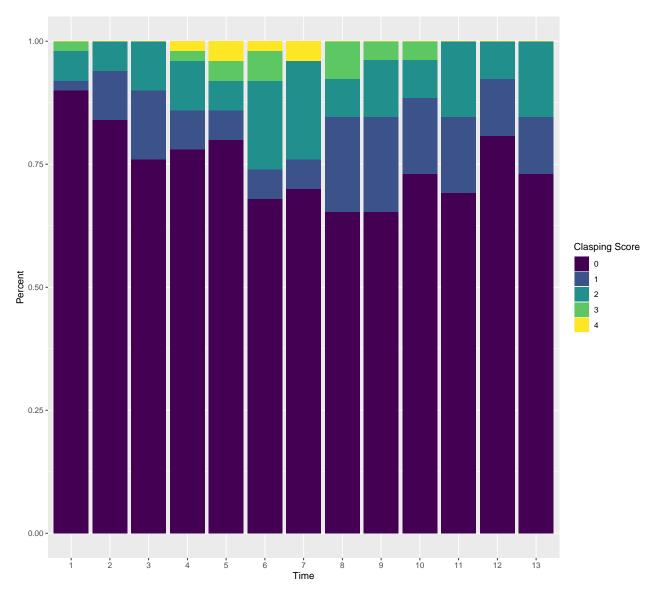


Figure 4: Bar plot for clasping over the 12 weeks for WT mice

4.1 Ordinal regression



```
##
                    Estimate Std. Error z value Pr(>|z|)
## genotypeHD
                                0.15712 17.519
                     2.75256
                                                <2e-16 ***
## TreatmentOnly ATB 0.34900
                                0.15791
                                          2.210
                                                  0.0271 *
## TreatmentATB_FMT
                     0.06226
                                0.15916
                                          0.391
                                                  0.6957
                     0.30508
                                0.02033 15.009
## Time
                                                  <2e-16 ***
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
## Threshold coefficients:
      Estimate Std. Error z value
## 0|1
        3.4581
                   0.2182 15.85
        4.0397
                   0.2278
                            17.73
## 1|2
## 213
       5.5397
                   0.2630
                            21.07
        6.0657
                   0.2761
                            21.96
## (927 observations deleted due to missingness)
```

4.2 Ordinal regression with random effect

```
Clasping.clmm = clmm(Clasping~genotype + Treatment + Time+(1|BOX),
          data=long_clasping, Hess=TRUE)
summary(Clasping.clmm)
## Cumulative Link Mixed Model fitted with the Laplace approximation
##
## formula: Clasping ~ genotype + Treatment + Time + (1 | BOX)
           long_clasping
## data:
## link threshold nobs logLik AIC
                                       niter
                                                  max.grad cond.H
  logit flexible 1049 -996.58 2011.17 523(3468) 6.98e-05 3.6e+03
##
## Random effects:
## Groups Name
                      Variance Std.Dev.
                               1.613
          (Intercept) 2.6
## Number of groups: BOX 51
## Coefficients:
##
                    Estimate Std. Error z value Pr(>|z|)
                                0.51567 7.507 6.05e-14 ***
## genotypeHD
                     3.87110
## TreatmentOnly ATB 0.32919
                                0.40377
                                          0.815
                                                   0.415
## TreatmentATB_FMT
                                                   0.538
                     0.33935
                                0.55081
                                          0.616
## Time
                     0.44920
                                0.02752 16.325 < 2e-16 ***
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
## Threshold coefficients:
      Estimate Std. Error z value
       5.1074
                   0.5320 9.601
## 0|1
## 1|2
       5.8632
                   0.5418 10.822
       7.7182
## 2|3
                   0.5720 13.494
        8.3527
                   0.5824 14.341
## 3|4
## (927 observations deleted due to missingness)
```



4.3 Comparison between Cumulative Link Model and Cumulative Link Mixed Model

```
anova( Clasping.clmm,Clasping.clm)
## Likelihood ratio tests of cumulative link models:
##
##
                formula:
                                                                   link:
## Clasping.clm Clasping ~ genotype + Treatment + Time
                                                                   logit
## Clasping.clmm Clasping ~ genotype + Treatment + Time + (1 | BOX) logit
                threshold:
## Clasping.clm flexible
## Clasping.clmm flexible
##
##
                no.par
                          AIC
                                logLik LR.stat df Pr(>Chisq)
## Clasping.clm
                     8 2201.9 -1092.97
                     9 2011.2 -996.58 192.78 1 < 2.2e-16 ***
## Clasping.clmm
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
```

Cumulative link mixed model is better than the cumulative link model with a lower AIC value.

5 Post-Hoc test

5.1 Body Weight

Table 1: Post-hoc results for genotype *Treatment in body weight

contrast	estimate	SE	df	t.ratio	p.value
WT No treatment - HD No treatment	5.5503	0.9342	71	5.9415	0.0000
WT No treatment - WT Only ATB	2.8106	1.0413	1297	2.6992	0.0760
WT No treatment - HD Only ATB	3.9178	0.9354	71	4.1886	0.0011
WT No treatment - WT ATB_FMT	0.5025	1.0205	71	0.4924	0.9963
WT No treatment - HD $\operatorname{ATB_FMT}$	4.9871	1.0245	71	4.8679	0.0001
HD No treatment - WT Only ATB	-2.7396	0.9361	71	-2.9266	0.0503
HD No treatment - HD Only ATB	-1.6325	0.5282	1297	-3.0908	0.0249



Table 1: Post-hoc results for gen	otype *Treatmen	nt in bod	y weig	tht (contin	nued)
contrast	estimate	SE	df	t ratio	n valu

contrast	estimate	SE	df	t.ratio	p.value
HD No treatment - WT ATB_FMT	-5.0478	0.9141	71	-5.5224	0.0000
HD No treatment - HD ATB_FMT	-0.5632	0.9119	71	-0.6176	0.9894
WT Only ATB - HD Only ATB	1.1072	0.9336	71	1.1859	0.8421
WT Only ATB - WT ATB_FMT	-2.3082	1.0205	71	-2.2617	0.2236
WT Only ATB - HD ATB_FMT	2.1765	1.0245	71	2.1244	0.2869
HD Only ATB - WT ATB_FMT	-3.4153	0.9129	71	-3.7411	0.0048
HD Only ATB - HD ATB_FMT	1.0693	0.9113	71	1.1734	0.8480
WT ATB $_$ FMT - HD ATB $_$ FMT	4.4846	0.9988	71	4.4900	0.0004

5.2 Weight Gain

Table 2: Post-hoc results for genotype * Treatment in Weight Gain

contrast	estimate	SE	df	t.ratio	p.value
WT No treatment - HD No treatment	17.6839	3.5853	23	4.9324	0.0007
WT No treatment - WT Only ATB	7.2766	4.0791	682	1.7839	0.4769
WT No treatment - HD Only ATB	13.3880	3.5875	23	3.7319	0.0123
WT No treatment - WT ATB_FMT	2.1142	3.8698	23	0.5463	0.9935
WT No treatment - HD $\operatorname{ATB_FMT}$	21.0392	3.8809	23	5.4212	$\boldsymbol{0.0002}$
HD No treatment - WT Only ATB	-10.4073	3.5853	23	-2.9028	0.0759
HD No treatment - HD Only ATB	-4.2958	1.8509	682	-2.3209	0.1870
HD No treatment - WT ATB_FMT	-15.5697	3.3452	23	-4.6544	0.0014
HD No treatment - HD ATB_FMT	3.3553	3.3580	23	0.9992	0.9135
WT Only ATB - HD Only ATB	6.1114	3.5875	23	1.7035	0.5432
WT Only ATB - WT ATB_FMT	-5.1624	3.8698	23	-1.3340	0.7637
WT Only ATB - HD ATB_FMT	13.7626	3.8809	23	3.5463	0.0188
HD Only ATB - WT ATB_FMT	-11.2738	3.3476	23	-3.3678	0.0281
HD Only ATB - HD ATB_FMT	7.6512	3.3604	23	2.2769	0.2434
WT ATB_FMT - HD ATB_FMT	18.9250	3.6602	23	5.1704	0.0004

5.3 Fecal Water Content

```
x<-emmeans(FecalWaterContent.lmm, pairwise ~ genotype*Treatment,
    at = list(Time = c(1:13)))</pre>
```



Table 3: Post-hoc results for genotype * Treatment in Fecal Water Content

contrast	estimate	SE	df	t.ratio	p.value
WT No treatment - HD No treatment	15.6830	8.5175	23	1.8413	0.4608
WT No treatment - WT Only ATB	0.6866	9.7349	677	0.0705	1.0000
WT No treatment - HD Only ATB	-9.6736	8.5237	23	-1.1349	0.8618
WT No treatment - WT ATB_FMT	4.6863	9.2371	23	0.5073	0.9954
WT No treatment - $HD ATB_FMT$	-1.0366	9.2566	23	-0.1120	1.0000
HD No treatment - WT Only ATB	-14.9964	8.5175	23	-1.7607	0.5086
HD No treatment - HD Only ATB	-25.3566	4.1105	677	-6.1687	0.0000
HD No treatment - WT ATB_FMT	-10.9967	7.9437	23	-1.3843	0.7356
HD No treatment - HD ATB_FMT	-16.7196	7.9664	23	-2.0988	0.3224
WT Only ATB - HD Only ATB	-10.3603	8.5237	23	-1.2155	0.8249
WT Only ATB - WT ATB_FMT	3.9997	9.2371	23	0.4330	0.9978
WT Only ATB - HD ATB_FMT	-1.7233	9.2566	23	-0.1862	1.0000
HD Only ATB - WT ATB_FMT	14.3599	7.9504	23	1.8062	0.4814
HD Only ATB - HD ATB_FMT	8.6370	7.9730	23	1.0833	0.8830
WT ATB_FMT - HD ATB_FMT	-5.7229	8.7315	23	-0.6554	0.9851

5.4 Food Intake

Table 4: Post-hoc results for genotype * Treatment in Food Intake

contrast	estimate	SE	$\mathrm{d}\mathrm{f}$	t.ratio	p.value
WT No treatment - HD No treatment	-0.0973	0.0345	22	-2.8239	0.0906
WT No treatment - WT Only ATB	-0.1906	0.0366	292	-5.2061	0.0000
WT No treatment - HD Only ATB	-0.0659	0.0345	22	-1.9130	0.4208
WT No treatment - WT ATB_FMT	-0.1767	0.0347	22	-5.0869	0.0005

estimate	SE	df	t.ratio	p.value
-0.0519	0.0366	22	-1.4181	0.7162
-0.0933	0.0345	22	-2.7089	0.1134
0.0314	0.0304	292	1.0328	0.9066
-0.0794	0.0324	22	-2.4472	0.1835
0.0454	0.0345	22	1.3168	0.7729
0.1247	0.0345	22	3.6199	0.0166
0.0139	0.0347	22	0.4008	0.9985
0.1387	0.0366	22	3.7879	0.0113
-0.1108	0.0324	22	-3.4144	0.0262
0.0140	0.0345	22	0.4058	0.9984
	-0.0519 -0.0933 0.0314 -0.0794 0.0454 0.1247 0.0139 0.1387 -0.1108	-0.0519	-0.0519 0.0366 22 -0.0933 0.0345 22 0.0314 0.0304 292 -0.0794 0.0324 22 0.0454 0.0345 22 0.1247 0.0345 22 0.0139 0.0347 22 0.1387 0.0366 22 -0.1108 0.0324 22	-0.0519 0.0366 22 -1.4181 -0.0933 0.0345 22 -2.7089 0.0314 0.0304 292 1.0328 -0.0794 0.0324 22 -2.4472 0.0454 0.0345 22 1.3168 0.1247 0.0345 22 3.6199 0.0139 0.0347 22 0.4008 0.1387 0.0366 22 3.7879 -0.1108 0.0324 22 -3.4144

Table 4: Post-hoc results for genotype * Treatment in Food Intake (continued)

5.5 Water Intake

WT ATB FMT - HD ATB FMT

 $0.1248 \quad 0.0347$

22

3.5921

0.0177

Table 5: Post-hoc results for Treatment given genotype in Water Intake

contrast	genotype	estimate	SE	df	t.ratio	p.value
No treatment - Only ATB	WT	-0.3721	0.1511	264	-2.4622	0.0383
No treatment - ATB_FMT	WT	-0.2737	0.1434	22	-1.9092	0.1598
Only ATB - ATB $_$ FMT	WT	0.0984	0.1434	22	0.6862	0.7739
No treatment - Only ATB	$_{ m HD}$	-0.2293	0.0974	264	-2.3526	0.0505
No treatment - ATB_FMT	$_{ m HD}$	-0.3113	0.1372	22	-2.2691	0.0817
Only ATB - ATB $_$ FMT	HD	-0.0820	0.1372	22	-0.5974	0.8229

5.6 CFC Extinction



```
color = ifelse(that_cell,"red","black"))%>%
column_spec(1,color = ifelse(that_cell,"red","black"))
```

Table 6: Post-hoc results for genotype given Treatment in CFC Extinction

contrast	Treatment	estimate	SE	df	t.ratio	p.value
WT - HD	No treatment Only ATB ATB_FMT	0.000	4.0707 4.0536 4.2334	41	0.4131	0.6817

5.7 Ymaze

Table 7: Post-hoc results for genotype given Treatment*Arm in Ymaze

contrast	Treatment	Arm	estimate	SE	df	t.ratio	p.value
WT - HD	No treatment	Novel	17.1598	8.2060	39	2.0911	0.0431
WT - HD	Only ATB	Novel	2.8606	7.7449	39	0.3694	0.7139
WT - HD	ATB_FMT	Novel	14.4345	7.5293	39	1.9171	0.0626
WT - HD	No treatment	Familiar	16.3095	8.2060	39	1.9875	0.0539
WT - HD	Only ATB	Familiar	2.0103	7.7449	39	0.2596	0.7966
WT - HD	ATB_FMT	Familiar	13.5842	7.5293	39	1.8042	0.0789

5.8 Brain Weight



Table 8: Post-hoc results for for genotype given Treatment in Brain weight

contrast	Treatment	estimate	SE	df	t.ratio	p.value
WT - HD	No treatment Only ATB ATB_FMT	0.0510	0.0202	23	2.5239	0.0032 0.0190 0.0039

5.9 Propel Brake Ratio- Week 14

Table 9: Post-hoc results for genotype given Treatment in Propel Brake Ratio

contrast	Treatment	estimate	SE	df	t.ratio	p.value
	No treatment Only ATB		000-		-3.0043 0.7281	
WT - HD	$\overline{\text{ATB}}_{-}\overline{\text{FMT}}$	-0.9988	0.4918	45	-2.0310	0.0482

5.10 FITC- Week 20

Table 10: Post-hoc results for genotype given Treatment in FITC

contrast	Treatment	estimate	SE	df	t.ratio	p.value
WT - HD	No treatment Only ATB ATB FMT	-0.2904	0.2277	12	-2.5616 -1.2756 -3.8653	0.2262



5.11 CecumWeight- Week 14

Table 11: Post-hoc results for Treatment given genotype in Cecum Weight

contrast	genotype	estimate	SE	df	t.ratio	p.value
No treatment - Only ATB	WT	0.0300	0.0350	16	0.8560	0.6747
No treatment - ATB_FMT	WT	0.0850	0.0379	16	2.2455	0.0936
Only ATB - ATB_FMT	WT	0.0550	0.0379	16	1.4530	0.3388
No treatment - Only ATB	$^{ m HD}$	-0.0193	0.0363	16	-0.5317	0.8571
No treatment - ATB_FMT	$^{ m HD}$	0.0477	0.0410	16	1.1626	0.4914
Only ATB - ATB_FMT	HD	0.0670	0.0400	16	1.6768	0.2442

5.12 Colon Length- Week 20

Table 12: Post-hoc results for genotype given Treatment in Colon length

contrast	Treatment	estimate	SE	df	t.ratio	p.value
WT - HD	No treatment	1.2244	0.4248	23	2.8821	0.0084
WT - HD	Only ATB	0.4483	0.3918	22	1.1443	0.2648
WT - HD	ATB_FMT	0.8804	0.3882	23	2.2675	0.0331

5.13 Acetate- Week 20

```
x<-emmeans(Acetate_w20.lmm, pairwise ~ genotype*Treatment)
df<-data.frame(x$contrasts)
that_cell<-df$p.value<0.05</pre>
```



Table 13: Post-hoc results for genotype Treatment in Acetate

contrast	estimate	SE	df	t.ratio	p.value
WT No treatment - HD No treatment	9.5017	18.6410	12	0.5097	0.9948
WT No treatment - WT Only ATB	10.3542	16.6730	12	0.6210	0.9872
WT No treatment - HD Only ATB	-37.8161	17.4868	12	-2.1625	0.3206
WT No treatment - WT ATB_FMT	20.1300	16.6730	12	1.2073	0.8254
WT No treatment - $HD ATB_FMT$	-10.0521	17.4868	12	-0.5748	0.9909
HD No treatment - WT Only ATB	0.8525	18.6410	12	0.0457	1.0000
HD No treatment - HD Only ATB	-47.3178	19.3723	12	-2.4425	0.2161
HD No treatment - WT ATB_FMT	10.6283	18.6410	12	0.5702	0.9913
HD No treatment - HD ATB_FMT	-19.5538	19.3723	12	-1.0094	0.9061
WT Only ATB - HD Only ATB	-48.1703	17.4868	12	-2.7547	0.1342
WT Only ATB - WT ATB_FMT	9.7758	16.6730	12	0.5863	0.9901
WT Only ATB - HD ATB_FMT	-20.4063	17.4868	12	-1.1670	0.8439
HD Only ATB - WT ATB_FMT	57.9461	17.4868	12	3.3137	0.0539
HD Only ATB - HD ATB_FMT	27.7640	18.2644	12	1.5201	0.6593
WT ATB_FMT - HD ATB_FMT	-30.1821	17.4868	12	-1.7260	0.5414

5.14 IL17E- Week 20

Table 14: Post-hoc results for Treatment given genotype in IL17E

contrast	genotype	estimate	SE	df	t.ratio	p.value
No treatment - Only ATB	WT	0.0321	0.1801	14	0.1782	0.9827
No treatment - ATB_FMT	WT	-0.4044	0.1819	14	-2.2229	0.1018
Only ATB - ATB $_$ FMT	WT	-0.4365	0.1877	14	-2.3252	0.0850
No treatment - Only ATB	$_{ m HD}$	0.2024	0.2292	14	0.8833	0.6592
No treatment - ATB_FMT	$_{ m HD}$	0.1352	0.2339	14	0.5783	0.8337
Only ATB - ATB_FMT	HD	-0.0672	0.1952	14	-0.3442	0.9371



6 Information about the used R session

sessionInfo()

```
## R version 4.1.0 (2021-05-18)
## Platform: x86_64-apple-darwin17.0 (64-bit)
## Running under: macOS Big Sur 10.16
##
## Matrix products: default
           /Library/Frameworks/R.framework/Versions/4.1/Resources/lib/libRblas.dylib
## LAPACK: /Library/Frameworks/R.framework/Versions/4.1/Resources/lib/libRlapack.dylib
##
## locale:
## [1] en_AU.UTF-8/en_AU.UTF-8/en_AU.UTF-8/C/en_AU.UTF-8/en_AU.UTF-8
##
## attached base packages:
                 graphics grDevices utils
## [1] stats
                                                datasets methods
                                                                     base
## other attached packages:
                           MASS_7.3-54
   [1] tibble_3.1.6
                                               kableExtra_1.3.4
                                                                   patchwork_1.1.1
##
   [5] rstatix_0.7.0
                           lmerTest_3.1-3
                                               lme4_1.1-27.1
                                                                   Matrix_1.3-4
##
  [9] viridis_0.6.1
                           viridisLite_0.4.0
                                               ordinal_2019.12-10 tidyr_1.1.4
## [13] emmeans_1.7.0
                           jtools_2.1.4
                                               TSA_1.3
                                                                   xtable_1.8-4
                                                                   nlme_3.1-153
## [17] sjPlot_2.8.9
                           dotwhisker_0.7.4
                                               stargazer_5.2.2
## [21] car_3.0-11
                           carData_3.0-4
                                               ggpubr_0.4.0
                                                                   ggplot2_3.3.5
## [25] magrittr_2.0.1
                           readxl_1.3.1
                                               dplyr_1.0.7
                                                                   knitr_1.36
##
## loaded via a namespace (and not attached):
   [1] TH.data_1.1-0
                            minqa_1.2.4
                                                 colorspace_2.0-2
##
   [4] ggsignif_0.6.3
                             ellipsis_0.3.2
                                                 rio_0.5.27
  [7] sjlabelled_1.1.8
                             estimability_1.3
                                                 ggstance_0.3.5
                                                 farver_2.1.0
## [10] parameters_0.14.0
                            rstudioapi_0.13
## [13] fansi_0.5.0
                            mvtnorm_1.1-2
                                                 xm12_1.3.2
                             splines_4.1.0
## [16] codetools 0.2-18
                                                 leaps_3.1
## [19] sjmisc_2.8.7
                            nloptr_1.2.2.2
                                                 ggeffects_1.1.1
## [22] broom_0.7.9
                             effectsize_0.5
                                                 compiler_4.1.0
## [25] httr_1.4.2
                             sjstats_0.18.1
                                                 backports_1.2.1
## [28] assertthat_0.2.1
                             fastmap_1.1.0
                                                 htmltools_0.5.2
## [31] tools_4.1.0
                             coda_0.19-4
                                                 gtable_0.3.0
## [34] glue_1.5.0
                             Rcpp_1.0.7
                                                 cellranger_1.1.0
## [37] vctrs_0.3.8
                             svglite_2.0.0
                                                 insight_0.14.4
## [40] xfun_0.26
                             stringr_1.4.0
                                                 openxlsx_4.2.4
## [43] rvest_1.0.1
                             lifecycle_1.0.1
                                                 zoo_1.8-9
## [46] scales_1.1.1
                                                 sandwich_3.0-1
                             hms_1.1.1
## [49] yaml_2.2.1
                             curl_4.3.2
                                                 gridExtra_2.3
## [52] pander_0.6.4
                             stringi_1.7.5
                                                 ucminf_1.1-4
## [55] bayestestR_0.11.0
                            boot_1.3-28
                                                 zip_2.2.0
## [58] rlang_0.4.12
                             pkgconfig_2.0.3
                                                 systemfonts_1.0.2
## [61] evaluate_0.14
                             lattice_0.20-45
                                                 purrr_0.3.4
## [64] labeling_0.4.2
                             tidyselect_1.1.1
                                                 R6_2.5.1
## [67] generics_0.1.0
                            multcomp_1.4-17
                                                 DBI 1.1.1
## [70] pillar_1.6.4
                            haven_2.4.3
                                                 foreign_0.8-81
```



##	[73]	withr_2.4.2	mgcv_1.8-37	survival_3.2-13
##	[76]	datawizard_0.2.1	abind_1.4-5	performance_0.8.0
##	[79]	modelr_0.1.8	crayon_1.4.2	utf8_1.2.2
##	[82]	rmarkdown_2.11	locfit_1.5-9.4	grid_4.1.0
##	[85]	data.table_1.14.2	forcats_0.5.1	digest_0.6.28
##	[88]	webshot_0.5.2	numDeriv_2016.8-1.1	munsell_0.5.0