fMRI Biomarker for Deep Brain Stimulation

Code ▼

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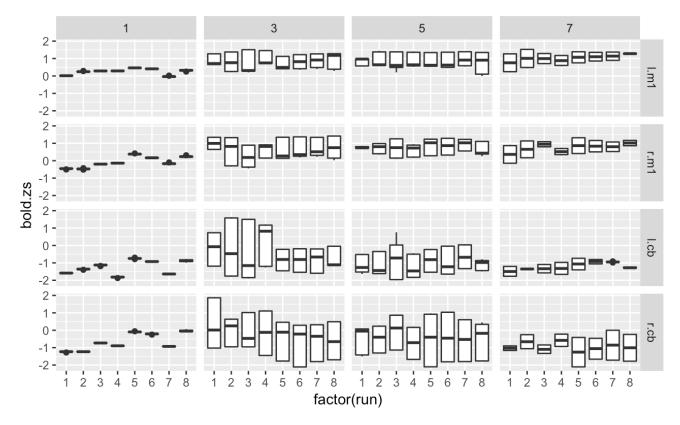
```
library(dplyr)
library(ggplot2)
library(tidyr)
library(readr)
library(feather)
library(readxl)
library(rpart)
```

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```
data.frg <-
 data.frq %>%
 mutate(group = "frequency") %>%
  gather(run, bold, run1:run6)
data.vlt <-
  data.vlt %>%
 rename(voltage = volt) %>%
 mutate(freq = 130L) %>%
 mutate(group = "voltage") %>%
 gather(run, bold, run1:run8)
data <-
 bind rows(
   data.frq,
   select (
     data.vlt, .dots =
       c("subj", "area", "run", "dbs0n0ff", "block", "contact", "voltage", "freq
", "group", "bold")
    )
  ) %>%
  select(group, everything()) %>%
 rename(dbs = dbsOnOff) %>%
 mutate(dbs = if_else(dbs == 2, 0, 1)) %>%
 mutate(run = as.integer(substr(run, 4, 4))) %>%
 mutate at(vars(group:run), funs(as.factor)) %>%
 mutate(area.full = as.factor(ifelse(area == "1", "1.m1", ifelse(area == "2", "r
.ml", ifelse(area == "3", "l.cb", ifelse(area == "4", "r.cb", NA)))))) %>%
 mutate(area.full = factor(area.full, c("l.m1", "r.m1", "l.cb", "r.cb"))) %>%
 mutate(subj.alt = as.factor(ifelse(subj == "76", "s3", ifelse(subj == "635", "s
1", ifelse(subj == "985", "s2", NA))))) %>%
 group by(group, subj, run) %>%
 mutate(bold.zs = (bold - mean(bold)) / sd(bold)) %>%
 ungroup() %>%
 mutate(area.rela = ifelse(subj.alt %in% c("s1", "s3"), ifelse(area.full == "l.m
1", "ipsi.m1", ifelse(area.full == "r.m1", "contra.m1", ifelse(area.full == "l.cb
", "ipsi.cb", ifelse(area.full == "r.cb", "contra.cb", NA)))), ifelse(area.full =
= "l.m1", "contra.m1", ifelse(area.full == "r.m1", "ipsi.m1", ifelse(area.full ==
"l.cb", "contra.cb", ifelse(area.full == "r.cb", "ipsi.cb", NA))))))
data <-
 data %>%
  select(group, subj, subj.alt, area, area.full, area.rela, dbs:bold, bold.zs)
rm(data.frq, data.vlt)
```

```
ggplot(
  data %>% filter(group == "voltage", subj.alt == "s1"),
  aes(
    x = factor(run),
    y = bold.zs
)
) +
  geom_boxplot() +
  facet_grid(
    area.full ~ voltage
)
```



```
std.err <-
  function(x) sd(x)/sqrt(length(x))
data.voltage <-
  data %>%
  filter(
    group == "voltage"
CalculateAreaMeasures <- function(data) {</pre>
  data.test.tmp <-
    data %>%
    group_by(area.rela, contact, voltage, run, block) %>%
    summarise(
      bold.zs.avg = mean(bold.zs),
      dbs.val = unique(dbs)
    )
  data.test.tmp <-</pre>
    data.test.tmp %>%
    mutate(
      res =
        (lag(bold.zs.avg) - bold.zs.avg) /
        mean(c(lag(bold.zs.avg), bold.zs.avg), na.rm = TRUE) *
    ) %>%
    filter(dbs.val == 0)
  data.test.tmp <-
    data.test.tmp %>%
    group by(area.rela, contact, voltage, run) %>%
    mutate(pair = 1:n()) %>%
    select(area.rela, contact, voltage, run, pair, res)
  data.test.tmp <-
    data.test.tmp %>%
    group by(area.rela, contact, voltage) %>%
    summarise(
      res.avg = mean(res, na.rm = TRUE),
      res.sem = std.err((res))
    ) %>%
    ungroup()
}
out <- list()
subj.contact.voltage.unqs <-</pre>
  data %>% select(subj.alt, contact, voltage) %>% unique()
for (i in 1:nrow(subj.contact.voltage.unqs)) {
  subj.alt.no <-</pre>
    subj.contact.voltage.ungs[i,]$subj.alt
```

```
contact.no <-
    subj.contact.voltage.unqs[i,]$contact
  voltage.no <-
    subj.contact.voltage.unqs[i,]$voltage
 data.voltage.no <-
    data.voltage %>%
    filter(
      subj.alt == subj.alt.no &
        contact == contact.no &
        voltage == voltage.no
 data.voltage.no.res <-
    CalculateAreaMeasures(data.voltage.no) %>%
    mutate(
     subj.alt = subj.alt.no
    )
 out[[i]] <- data.voltage.no.res</pre>
}
result <-
 bind rows(out) %>%
  select(subj.alt, area.rela:res.sem) %>%
 group by(subj.alt, contact, voltage) %>%
 mutate(
   res.avg.wins = ifelse(area.rela != "ipsi.m1" & res.avg < 0, abs(mean(res.avg)
), res.avg),
    ipsi.ml.weighted = (0.9 * (res.avg.wins[area.rela == "ipsi.ml"])),
    other.weighted = (0.1 * (mean(c(res.avg.wins[area.rela == "ipsi.cb"], res.avg
.wins[area.rela == "contra.cb"], res.avg.wins[area.rela == "contra.ml"])))),
    res.overall =
      ipsi.ml.weighted / other.weighted
  arrange(subj.alt, desc(res.overall)); result %>%
  ungroup()
```

subj.alt <fctr></fctr>	area.rela <chr></chr>	contact <fctr></fctr>	voltage <fctr></fctr>	res.avg <dbl></dbl>	res.sem <dbl></dbl>	res.avg.wins <dbl></dbl>	ips
s1	contra.cb	2	3	-1.492424171	2.68586407	0.372365767	
s1	contra.m1	2	3	0.124078968	0.25987457	0.124078968	
s1	ipsi.cb	2	3	0.669021111	0.70813147	0.669021111	
s1	ipsi.m1	2	3	2.188787158	1.58336077	2.188787158	
s1	contra.cb	1	1	0.460450739	1.07330687	0.460450739	
s1	contra.m1	1	1	1.048837796	1.21138233	1.048837796	

subj.alt <fctr></fctr>	area.rela <chr></chr>	contact <fctr></fctr>	voltage <fctr></fctr>	res.avg <dbl></dbl>	res.sem <dbl></dbl>	res.avg.wins ips <dbl></dbl>
s1	ipsi.cb	1	1	0.087214912	0.14367365	0.087214912
s1	ipsi.m1	1	1	2.936753351	3.65902335	2.936753351
s1	contra.cb	3	5	0.023934234	0.04168758	0.023934234
s1	contra.m1	3	5	0.978259652	0.34377631	0.978259652
1-10 of 84 rows 1-8 of 10 columns				Previous 1	2 3 4	5 6 9 Next

```
# https://plot.ly/r/3d-scatter-plots/
# http://www.statmethods.net/graphs/scatterplot.html
plot.data <-
    result %>%
    group_by(subj.alt, contact, voltage) %>%
    summarise(res = unique(res.overall)) %>%
    ungroup() %>%
    filter(!(subj.alt == "s3" & contact == "3" & voltage == "5"))
library(plotly)
```

```
Attaching package: 'plotly'

The following object is masked from 'package:ggplot2':

last_plot

The following object is masked from 'package:stats':

filter

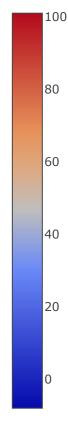
The following object is masked from 'package:graphics':

layout
```

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```
p <- plot_ly(</pre>
  plot.data,
  x = \sim contact, y = \sim voltage, z = \sim res,
  marker = list(color = ~res, colorscale = c('#FFE1A1', '#683531'), showscale = T
RUE)) %>%
  add_markers() %>%
  layout(scene = list(xaxis = list(title = 'Contact'),
                       yaxis = list(title = 'Voltage'),
                       zaxis = list(title = 'Effectiveness')),
         annotations = list(
           x = 1.13,
           y = 1.05,
           text = 'Weighted Adjusted Ipsi M1 Activation',
           xref = 'paper',
           yref = 'paper',
            showarrow = FALSE
         ))
р
```

Weighted Adjusted Ipsi M1 Activation



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```
# # Create a shareable link to your chart
# # Set up API credentials: https://plot.ly/r/getting-started
# chart link = plotly POST(p, filename="scatter3d/colorscale")
# chart_link
                                                                            Hide
# http://www.statmethods.net/stats/regression.html
fit <-
 lm(bold.zs ~ group + area.rela + dbs + contact + voltage + freq, data = data)
summary(fit)
Call:
lm(formula = bold.zs ~ group + area.rela + dbs + contact + voltage +
   freq, data = data)
Residuals:
   Min
            10 Median
                           30
                                  Max
-3.5012 -0.4139 -0.0182 0.3796 2.4053
Coefficients: (1 not defined because of singularities)
                   Estimate Std. Error t value Pr(>|t|)
                  -1.111089 0.010644 -104.387 < 2e-16 ***
(Intercept)
groupvoltage
                   0.290294 0.006958 41.722 < 2e-16 ***
area.relacontra.m1 1.219311 0.005439 224.176 < 2e-16 ***
area.relaipsi.cb -0.160875 0.005439 -29.578 < 2e-16 ***
area.relaipsi.m1
                  1.530060 0.005439 281.309 < 2e-16 ***
dbs1
                   0.001181 0.003847
                                        0.307 0.75884
contact2
                  -0.015744 0.005565 -2.829 0.00467 **
                  -0.228379 0.005470 -41.754 < 2e-16 ***
contact3
                  0.366700 0.008777 41.778 < 2e-16 ***
voltage3
voltage5
                  0.180250 0.008777 20.536 < 2e-16 ***
voltage7
                   0.391127
                            0.009751 40.111 < 2e-16 ***
                                   NA
freq130
                         NA
                                            NA
                                                    NΑ
freq180
                   0.323256
                            0.008159 39.621 < 2e-16 ***
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
Residual standard error: 0.6568 on 116628 degrees of freedom
Multiple R-squared: 0.5686, Adjusted R-squared: 0.5685
F-statistic: 1.397e+04 on 11 and 116628 DF, p-value: < 2.2e-16
```

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```
coefficients(fit)
```

```
(Intercept)
                        groupvoltage area.relacontra.ml
                                                           area.relaipsi.cb
     -1.11108854
                          0.29029410
                                              1.21931118
                                                                -0.16087506
area.relaipsi.ml
                                dbs1
                                                contact2
                                                                    contact3
      1.53005981
                          0.00118103
                                             -0.01574431
                                                                 -0.22837915
        voltage3
                            voltage5
                                                voltage7
                                                                     freq130
      0.36669974
                          0.18024999
                                              0.39112665
                                                                          NA
         freq180
      0.32325625
```

```
confint(fit, level=0.95)
```

```
2.5 %
                                      97.5 %
                   -1.131950511 -1.090226576
(Intercept)
                    0.276656848 0.303931362
groupvoltage
area.relacontra.ml 1.208650680 1.229971681
area.relaipsi.cb
                   -0.171535564 -0.150214563
area.relaipsi.m1
                    1.519399309 1.540720310
dbs1
                   -0.006358944 0.008721004
contact2
                   -0.026652510 -0.004836110
contact3
                   -0.239099526 -0.217658769
voltage3
                    0.349496468 0.383903016
voltage5
                    0.163046718
                                 0.197453267
voltage7
                    0.372014464
                                 0.410238840
freq130
                             NA
                                          NA
                    0.307265499 0.339247000
freq180
```

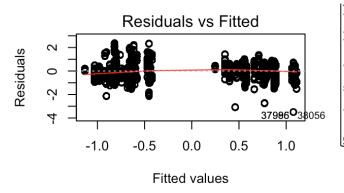
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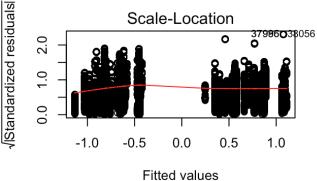
```
# fitted(fit)
# residuals(fit)
anova(fit)
```

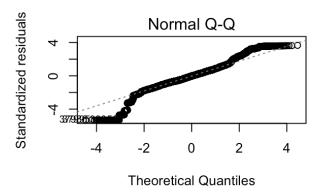
```
Analysis of Variance Table
Response: bold.zs
              Df Sum Sq Mean Sq
                                    F value Pr(>F)
group
               1
                       0
                             0.0
                                     0.0000 1.0000
                  63528 21176.0 49095.0171 <2e-16 ***
area.rela
               3
dbs
                      0
                             0.0
                                     0.0943 0.7588
               1
contact
               2
                    874
                           436.8 1012.5845 <2e-16 ***
voltage
               3
                   1220
                           406.8
                                   943.0367 <2e-16 ***
freq
               1
                    677
                           677.1 1569.8616 <2e-16 ***
Residuals 116628
                  50305
                             0.4
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
```

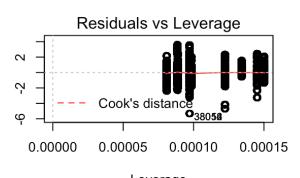
```
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```

```
# vcov(fit)
# influence(fit)
layout(matrix(c(1,2,3,4),2,2)) # optional 4 graphs/page
plot(fit)
```









Leverage

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```
# http://www.statmethods.net/advstats/cart.html
# http://blog.revolutionanalytics.com/2013/06/plotting-classification-and-regress
ion-trees-with-plotrpart.html
# http://www.milbo.org/rpart-plot/prp.pdf
# grow tree
fit <-
 rpart(
    bold.zs ~ group + area.rela + dbs + contact + voltage + freq,
    method = "anova",
    control = rpart.control(cp = 0.003),
    data = data
# fit <- rpart(Mileage~Price + Country + Reliability + Type,</pre>
     method="anova", data=cu.summary)
printcp(fit) # display the results
```

Standardized residuals

```
Regression tree:
rpart(formula = bold.zs ~ group + area.rela + dbs + contact +
   voltage + freq, data = data, method = "anova", control = rpart.control(cp = 0
.003))
Variables actually used in tree construction:
[1] area.rela contact freq
Root node error: 116604/116640 = 0.99969
n = 116640
        CP nsplit rel error xerror
                                        xstd
               0 1.00000 1.00003 0.0028884
1 0.5295092
2 0.0206381
                    0.47049 0.47050 0.0026948
3 0.0149736
                2 0.44985 0.44987 0.0027459
4 0.0120743
                3
                    0.43488 0.43492 0.0025123
               4 0.42280 0.42285 0.0025424
5 0.0096233
6 0.0043959
                6 0.40356 0.40362 0.0024508
7 0.0036837
               7 0.39916 0.39923 0.0024328
8 0.0035013
               8 0.39548 0.39556 0.0024223
9 0.0030000
                    0.38848 0.38857 0.0024361
             10
```

```
plotcp(fit) # visualize cross-validation results
summary(fit) # detailed summary of splits
```

```
Call:
rpart(formula = bold.zs ~ group + area.rela + dbs + contact +
    voltage + freq, data = data, method = "anova", control = rpart.control(cp = 0
.003))
 n = 116640
           CP nsplit rel error
                                  xerror
                                                xstd
1 0.529509184
                   0 1.0000000 1.0000268 0.002888431
2 0.020638139
                   1 0.4704908 0.4705037 0.002694793
3 0.014973560
                   2 0.4498527 0.4498723 0.002745938
4 0.012074315
                   3 0.4348791 0.4349208 0.002512326
                  4 0.4228048 0.4228533 0.002542409
5 0.009623287
6 0.004395947
                   6 0.4035582 0.4036229 0.002450821
7 0.003683676
                  7 0.3991623 0.3992347 0.002432814
                  8 0.3954786 0.3955552 0.002422309
8 0.003501347
9 0.003000000
                 10 0.3884759 0.3885661 0.002436089
Variable importance
area.rela
          voltage
                      contact
                                   freq
                                            group
       88
                  5
                            5
                                                1
Node number 1: 116640 observations,
                                       complexity param=0.5295092
  mean=3.988198e-18, MSE=0.9996914
  left son=2 (58320 obs) right son=3 (58320 obs)
  Primary splits:
      area.rela splits as LRLR, improve=5.295092e-01, (0 missing)
                splits as LRLR, improve=6.329797e-03, (0 missing)
      voltage
      contact
                splits as RRL,
                                improve=6.291888e-03, (0 missing)
                                 improve=3.266464e-03, (0 missing)
      freq
                splits as LRR,
      dbs
                splits as LR,
                                 improve=3.486433e-07, (0 missing)
Node number 2: 58320 observations,
                                     complexity param=0.02063814
  mean=-0.7275615, MSE=0.6003057
  left son=4 (22680 obs) right son=5 (35640 obs)
  Primary splits:
                splits as RRL, improve=0.0687375500, (0 missing)
      contact
                splits as LRLR, improve=0.0184741400, (0 missing)
      voltage
                splits as LRR, improve=0.0122965000, (0 missing)
      area.rela splits as R-L-, improve=0.0107781700, (0 missing)
                                 improve=0.0001607752, (0 missing)
      group
                splits as LR,
  Surrogate splits:
      voltage splits as RRRL, agree=0.648, adj=0.095, (0 split)
Node number 3: 58320 observations,
                                    complexity param=0.01207432
  mean=0.7275615, MSE=0.3403855
  left son=6 (29160 obs) right son=7 (29160 obs)
  Primary splits:
      area.rela splits as -L-R, improve=0.070923040, (0 missing)
```

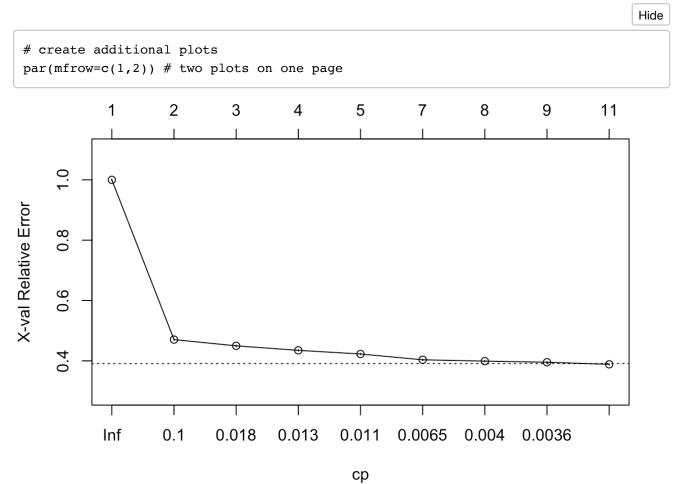
```
splits as LRR,
                                improve=0.031089810, (0 missing)
      contact
      voltage
                splits as LRLR, improve=0.008498923, (0 missing)
                splits as LLR,
                                improve=0.005027549, (0 missing)
      freq
      group
                splits as RL,
                                 improve=0.000283544, (0 missing)
Node number 4: 22680 observations,
                                     complexity param=0.01497356
  mean=-0.9822038, MSE=0.4796253
  left son=8 (16200 obs) right son=9 (6480 obs)
  Primary splits:
      voltage
               splits as -LLR, improve=1.605068e-01, (0 missing)
      area.rela splits as R-L-, improve=2.969807e-03, (0 missing)
                splits as LLR, improve=4.780479e-04, (0 missing)
                                improve=3.864680e-04, (0 missing)
      group
                splits as RL,
      dbs
                splits as LR,
                                improve=4.739182e-08, (0 missing)
Node number 5: 35640 observations,
                                     complexity param=0.009623287
  mean=-0.5655164, MSE=0.6095801
  left son=10 (4860 obs) right son=11 (30780 obs)
  Primary splits:
      freq
                splits as LRR, improve=0.042530210, (0 missing)
               splits as LRRR, improve=0.030376280, (0 missing)
      voltage
      area.rela splits as R-L-, improve=0.018995820, (0 missing)
      group
                splits as LR,
                                improve=0.007825542, (0 missing)
               splits as RL-, improve=0.002721304, (0 missing)
      contact
Node number 6: 29160 observations
  mean=0.5721872, MSE=0.2480955
Node number 7: 29160 observations,
                                    complexity param=0.004395947
  mean=0.8829358, MSE=0.3843932
  left son=14 (8100 obs) right son=15 (21060 obs)
  Primary splits:
      contact splits as LRR, improve=4.573015e-02, (0 missing)
      voltage splits as RRLL, improve=1.502599e-02, (0 missing)
      freq
              splits as LLR, improve=1.478618e-03, (0 missing)
                              improve=2.985965e-04, (0 missing)
      group
              splits as RL,
      dbs
              splits as LR,
                               improve=6.113844e-06, (0 missing)
  Surrogate splits:
      voltage splits as LRRR, agree=0.796, adj=0.267, (0 split)
Node number 8: 16200 observations
  mean=-1.157684, MSE=0.1499303
Node number 9: 6480 observations
 mean=-0.5435037, MSE=1.034422
Node number 10: 4860 observations
  mean=-0.9707266, MSE=0.4960903
```

```
Node number 11: 30780 observations,
                                      complexity param=0.009623287
  mean=-0.5015359, MSE=0.5974805
  left son=22 (17280 obs) right son=23 (13500 obs)
  Primary splits:
               splits as LRLL, improve=0.071789450, (0 missing)
      voltage
      area.rela splits as R-L-, improve=0.020276010, (0 missing)
      contact
               splits as RL-, improve=0.004737381, (0 missing)
                                improve=0.004197774, (0 missing)
      group
                splits as RL,
               splits as -LR, improve=0.004197774, (0 missing)
      freq
  Surrogate splits:
      group splits as RL, agree=0.719, adj=0.36, (0 split)
      freq splits as -LR, agree=0.719, adj=0.36, (0 split)
Node number 14: 8100 observations
  mean=0.6691515, MSE=0.4748433
Node number 15: 21060 observations
  mean=0.9651606, MSE=0.3252655
Node number 22: 17280 observations, complexity param=0.003683676
  mean=-0.6845932, MSE=0.441233
  left son=44 (8640 obs) right son=45 (8640 obs)
  Primary splits:
      area.rela splits as R-L-, improve=5.633562e-02, (0 missing)
      voltage splits as L-RR, improve=4.620928e-02, (0 missing)
               splits as RL-, improve=3.511804e-02, (0 missing)
      contact
      dbs
                splits as LR,
                                improve=4.892559e-07, (0 missing)
Node number 23: 13500 observations
  mean=-0.2672224, MSE=0.6996816
Node number 44: 8640 observations
 mean=-0.8422547, MSE=0.3597722
Node number 45: 8640 observations,
                                    complexity param=0.003501347
 mean=-0.5269318, MSE=0.4729796
  left son=90 (2160 obs) right son=91 (6480 obs)
  Primary splits:
      voltage splits as L-RR, improve=6.577038e-02, (0 missing)
      contact splits as RL-, improve=2.276113e-02, (0 missing)
              splits as LR, improve=8.546691e-07, (0 missing)
      dbs
Node number 90: 2160 observations
  mean=-0.8324218, MSE=0.1822746
Node number 91: 6480 observations,
                                   complexity param=0.003501347
 mean=-0.4251017, MSE=0.5284039
  left son=182 (4320 obs) right son=183 (2160 obs)
  Primary splits:
```

```
contact splits as RL-, improve=1.599764e-01, (0 missing)
voltage splits as --RL, improve=5.910179e-03, (0 missing)
dbs splits as LR, improve=7.994244e-07, (0 missing)

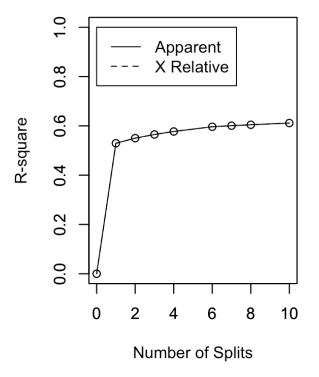
Node number 182: 4320 observations
mean=-0.6306889, MSE=0.2629851

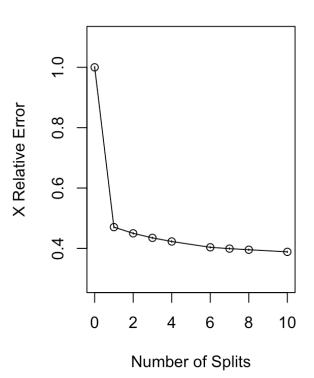
Node number 183: 2160 observations
mean=-0.01392737, MSE=0.805645
```



rsq.rpart(fit) # visualize cross-validation results

```
Regression tree:
rpart(formula = bold.zs ~ group + area.rela + dbs + contact +
    voltage + freq, data = data, method = "anova", control = rpart.control(cp = 0
.003))
Variables actually used in tree construction:
[1] area.rela contact
Root node error: 116604/116640 = 0.99969
n = 116640
         CP nsplit rel error xerror
                                           xstd
1 0.5295092
                     1.00000 1.00003 0.0028884
2 0.0206381
                     0.47049 0.47050 0.0026948
3 0.0149736
                     0.44985 0.44987 0.0027459
4 0.0120743
                 3
                     0.43488 0.43492 0.0025123
5 0.0096233
                     0.42280 0.42285 0.0025424
6 0.0043959
                     0.40356 0.40362 0.0024508
7 0.0036837
                 7
                     0.39916 0.39923 0.0024328
8 0.0035013
                     0.39548 0.39556 0.0024223
                 8
9 0.0030000
                     0.38848 0.38857 0.0024361
                10
```





```
# # plot tree
# plot(fit, uniform=TRUE,
# main="Regression Tree for DBS Effect on Bold")
# text(fit, use.n=TRUE, all=TRUE, cex=.8)
# # create attractive postcript plot of tree
# post(fit, file = "c:/tree2.ps",
# title = "Regression Tree for Mileage ")
rpart.plot(fit, type = 3, box.palette="RdYlGn", tweak = 1.2)
```

