Analysis of MST-41 Anti-Fibrotic Therapy

Peter Higgins

Anti-fibrotic Therapy Data

Background

- Intestinal Fibrosis leads to surgery in 70% of Crohn's disease
- Intestinal Myofibroblasts are stimulated to generate scar by:
 - injury/ wounding
 - matrix stiffness

Background 2

- Matrix Stiffness Activates Myofibroblasts
- Activated Myofibroblasts Synthesize Stiff Matrix
- Positive Feedback Loop -> More Fibrosis
 - This is Bad
 - * It gets worse
 - · It can get even worse

Hypothesis and Aims

Hypothesis

• Determine whether Anti-fibrotic drug can reverse fibrosis

Aims

- Determine whether Colon Weight is reduced by AFD
- Determine whether Gene Expression of Collagen 1, TGFb, CTGF is reduced by AFD
- Determine whether Protein expression of Collagen 1, aSMA, FN is reduced by AFD

Experimental Design

Experimental Design 1

- Get mice
- Strep cleanout

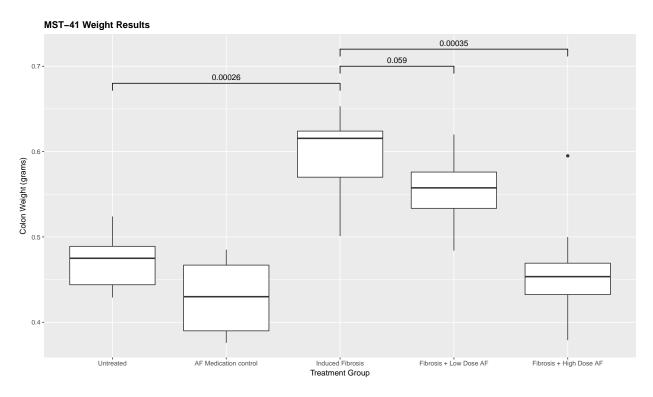
Experimental Design 2

- 1. Infect with Salmonella
- 2. Wait 8 days
- 3. Levo x 5 days in the water
- 4. Rest Day
- 5. Anti-fibrotic drug x 10 days
- 6. Image
- Ultrasound + SWV + Strain
- \bullet Photoacoustic
- 7. Sac

Experimental Design Diagram

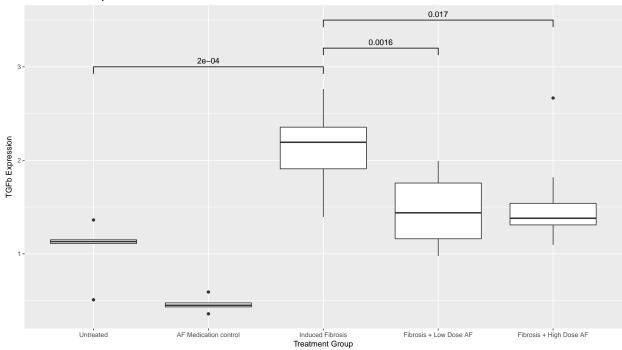
Results

Weight Result



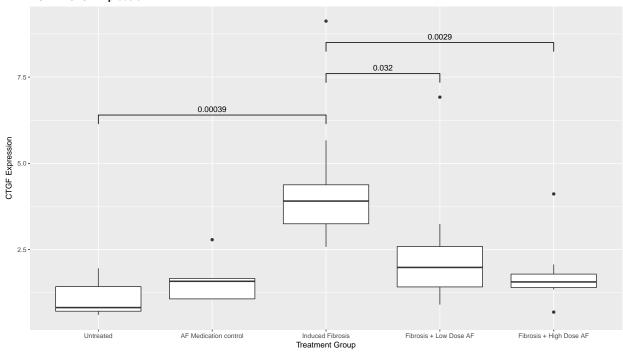
TGFb Result

MST-41 TGFb Expression



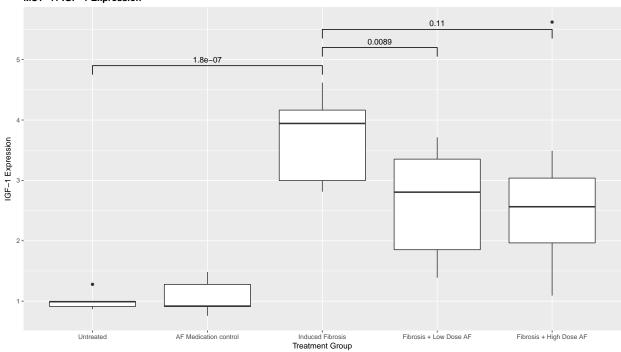
CTGF Result

MST-41 CTGF Expression



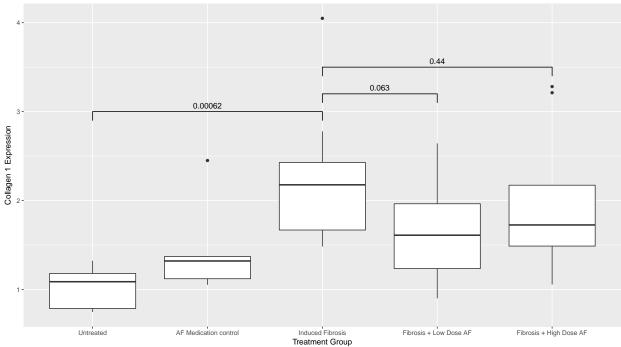
IGF-1 Result

MST-41 IGF-1 Expression



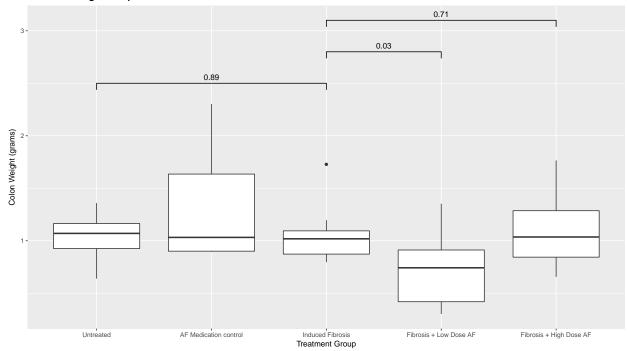
Collagen 1 Result

MST-41 Collagen 1 Expression



Collagen 3 Result

MST-41 Collagen 3 Expression



Conclusions

Conclusions

- Antifibrotic Drug Works in Human Intestinal Organoids
- Antifibrotic Drug Works in Salmonella Mouse Model
- Antifibrotic Drug Works in TNBS Rat Model
 - This is Amazing
 - * Clinical Trials to Follow
 - · Sure Hope the Ultrasound Works

Examples of other stuff you can do with Rmd

R Markdown

- This is an R Markdown document. Markdown is a simple formatting syntax for authoring HTML, PDF, MS Word, and now **PowerPoint** documents. For more details on using R Markdown see
- When you click the *Knit* button a document will be generated that includes both content as well as the output of any embedded R code chunks within the document.

Block Quotes

A friend once said:

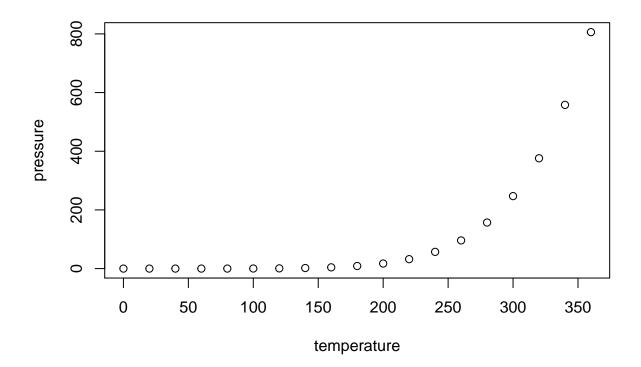
It's always better to give than to receive. Totes magotes, my friend, he said. I don't know why he said this.

You can embed an R code chunk like this:

summary(cars)

```
dist
##
        speed
##
            : 4.0
                               2.00
                    Min.
    1st Qu.:12.0
                    1st Qu.: 26.00
##
                    Median : 36.00
    Median:15.0
##
                            : 42.98
##
    Mean
            :15.4
                    Mean
    3rd Qu.:19.0
                    3rd Qu.: 56.00
            :25.0
##
    Max.
                    Max.
                            :120.00
```

Including Plots



And Tables

Treatment Arm	Colon Weight (gm)
Control	17.2
S.typhi	23.7
S.typhi+ AFD	17.9

Inline R code

Inline R Code

There were 50 cars studied

R Code Chunks

```
summary(cars$dist)
```

```
## Min. 1st Qu. Median Mean 3rd Qu. Max.
## 2.00 26.00 36.00 42.98 56.00 120.00
```

summary(cars\$speed)

```
## Min. 1st Qu. Median Mean 3rd Qu. Max.
## 4.0 12.0 15.0 15.4 19.0 25.0
```

Add Links

Links

Use a plain http address or add a link to a phrase:

Miscellaneous

• Manual Line breaks, end line with 2 or more spaces Roses are red, Violets are blue.

 $superscript^2\\$

 $\frac{\text{strikethrough}}{\text{strikethrough}}$

Horizontal Line break

Horizontal Rule / Page Break

Three or more asterisks or dashes:

and continue

Use of the echo parameter

- Note that the echo = FALSE parameter was added to the previous plotting code chunk to prevent printing of the R code that generated the plot.
- \bullet If you want to see the R code and the output, like in the previous summary table, then set the parameter to echo=TRUE