

# Exploration of the Safety of Droperidol Use in the ER

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FDA

Use droperidol to manage  
agitation for ER patients

# Tran's Hypothesis

droperidol is **safe** to use at the dosages administered in the ER

# Absolute Risk of Arrhythmia

$$AR = \frac{\text{\# of patients with an arrhythmia after receiving droperidol}}{\text{\# of patients that received droperidol}} \cdot 100$$

**Point estimate:**

$$\widehat{AR} = 6.9 \%$$

**95% confidence interval:**

$$(5.0 \%, 8.9\%)$$

About 7% of the patients that received droperidol in the ER had an arrhythmia

What factors are associated  
with an *increased* risk of  
arrhythmia?

# Possible predictors of arrhythmia

Variables measured before or at time of administering droperidol

Legal Sex

Age

Indication

EKG obtained before?

QTc (before)

Administered Dose Amount (mg)

# Possible predictors of arrhythmia

Variables measured before or at time **within 30 minutes** of administering droperidol

Legal Sex

Age

Indication

EKG obtained before?

QTc (before)

Administered Dose Amount (mg)

**Total Dose (mg)**

# Possible predictors of arrhythmia

Variables measured before or ~~at time~~ **within 30 minutes** of administering droperidol

Legal Sex

Age

Indication

EKG obtained before?

QTc (before)

**redundant**

QTc (before)

Administered Dose Amount (mg)

**Total Dose (mg)**



# Possible predictors of arrhythmia

Variables measured before or at time **within 30 minutes** of administering droperidol

Legal Sex

Age

Indication

~~EKG obtained before?~~

QTc (before)

Administered Dose Amount (mg)

**Total Dose (mg)**

**Similar, but different**

**Total Dose (mg)**

# Possible predictors of arrhythmia

Variables measured before or at time **within 30 minutes** of administering droperidol

Legal Sex

Age

Indication

~~EKG obtained before?~~

QTc (before)

~~Administered Dose Amount (mg)~~

**Total Dose (mg)**

# Logistic Regression

## Possible Predictors

Legal Sex

Age

Indication

QTc (before)

Total Dose (mg)

# Logistic Regression

## Possible Predictors

Legal Sex

Age

Indication

~~QTc (before)~~



75.7% of measurements  
missing

Total Dose (mg)

# Logistic Regression

## Possible Predictors

Legal Sex

Age

Indication

Total Dose (mg)

Headache

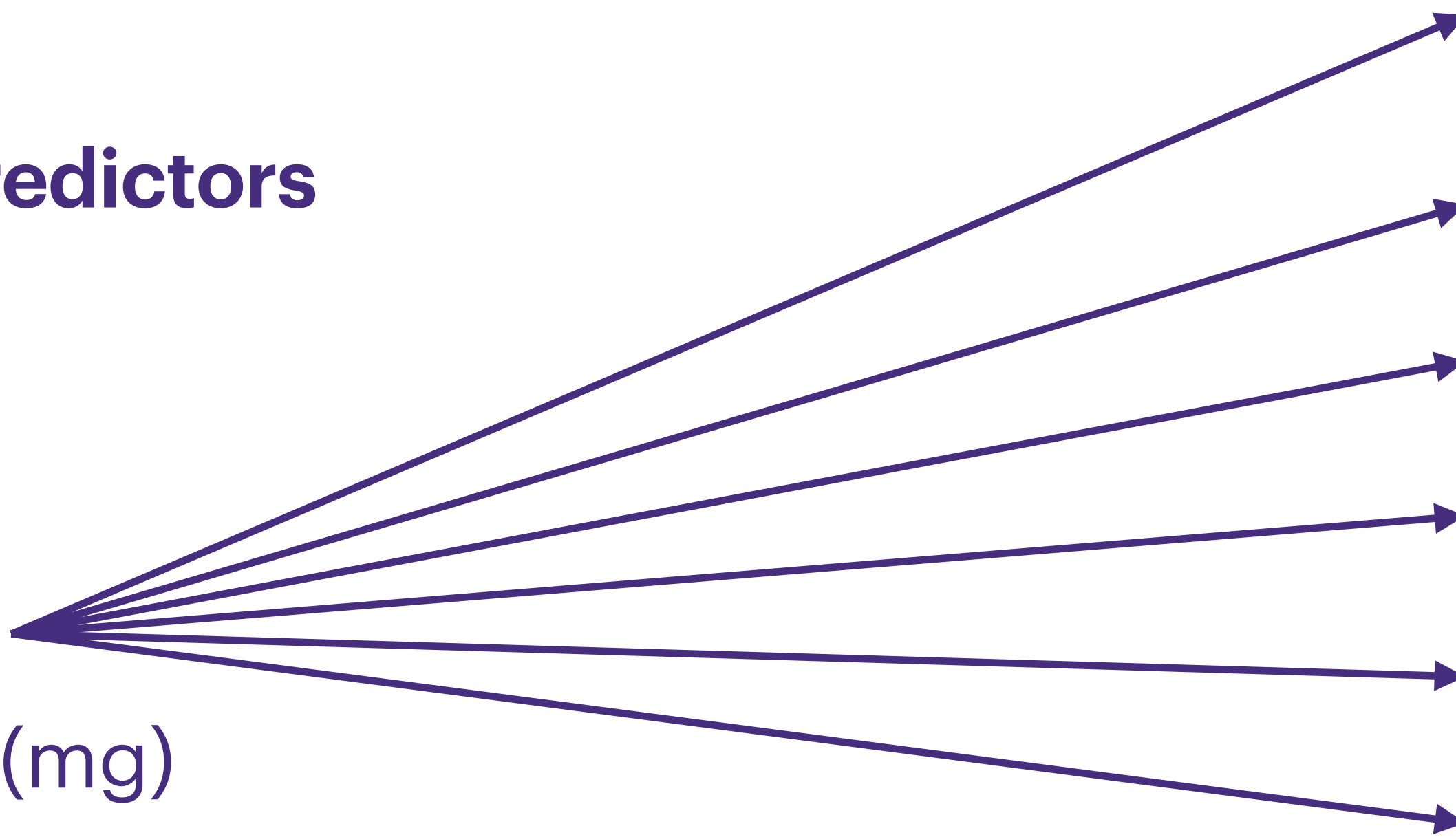
Abdominal Pain

Vomiting

Nausea

Agitation

Unknown



# Logistic Regression

## Correlation Matrix

	Legal Sex	Age	Total Dose	QTc (before)	Abdominal pain	Nausea	Agitation	Vomiting	Unknown	Headache
Legal Sex	1.00	0.07	0.09	0.14	0.01	-0.07	0.20	-0.08	-0.06	-0.19
Age	0.07	1.00	-0.01	0.12	0.00	-0.09	0.10	-0.09	0.00	-0.03
Total Dose	0.09	-0.01	1.00	0.16	-0.06	-0.29	0.51	-0.28	-0.12	-0.33
QTc (before)	0.14	0.12	-0.16	1.00	-0.10	0.03	0.03	0.03	0.09	-0.10
Abdominal pain	0.01	0.00	-0.06	-0.10	1.00	-0.04	-0.14	-0.07	-0.02	-0.05
Nausea	-0.07	-0.09	-0.29	0.03	-0.04	1.00	-0.64	1.00	-0.09	-0.15
Agitation	0.20	0.10	0.51	0.03	-0.14	-0.64	1.00	-0.64	-0.19	-0.51
Vomiting	-0.08	-0.09	-0.28	0.03	-0.07	1.00	-0.64	1.00	-0.09	-0.15
Unknown	-0.06	0.00	-0.12	0.09	-0.02	-0.09	-0.19	-0.09	1.00	-0.07
Headache	-0.19	-0.03	-0.33	-0.10	-0.05	-0.15	-0.51	-0.15	-0.07	1.00

removed nausea

# Logistic Regression

## Possible Predictors

Legal Sex

Age

Headache

Abdominal Pain

Vomiting

Agitation

Unknown

Total Dose (mg)



### **warning**

glm.fit: fitted probabilities  
numerically 0 or 1 occurred

# Logistic Regression

## Possible Predictors

Legal Sex

Age

Headache

No arrhythmias

Abdominal Pain

Vomiting

Agitation

Unknown

No arrhythmias

Total Dose (mg)

### warning

glm.fit: fitted probabilities  
numerically 0 or 1 occurred



# Logistic Regression

## Possible Predictors

Legal Sex

Age

~~Headache~~

Abdominal Pain

Vomiting

Agitation

~~Unknown~~

Total Dose (mg)



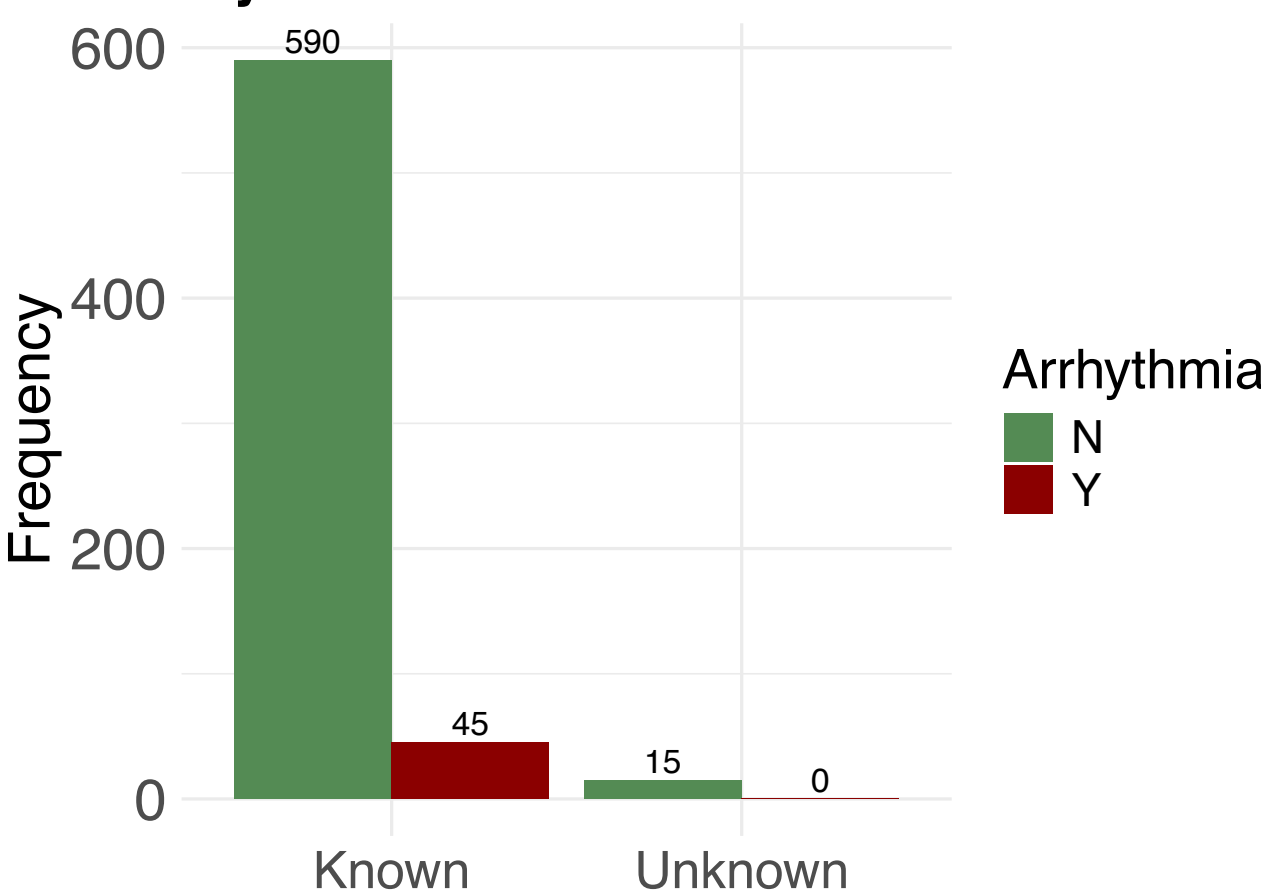
Agitation associated with  
*increased* risk of arrhythmia

	$\hat{\beta}$	Std. Error	p-value
Intercept	-4.733	0.885	$8.838 \times 10^{-8}$
Sex (Male)	0.104	0.344	0.762
Sex (X)	-13.424	1333.927	0.992
Age	0.007	.010	0.508
Total Dose	0.002	.082	0.982
Abdominal Pain	-12.195	799.084	0.989
Agitation	2.055	0.723	0.004
Vomiting	1.203	0.721	0.095

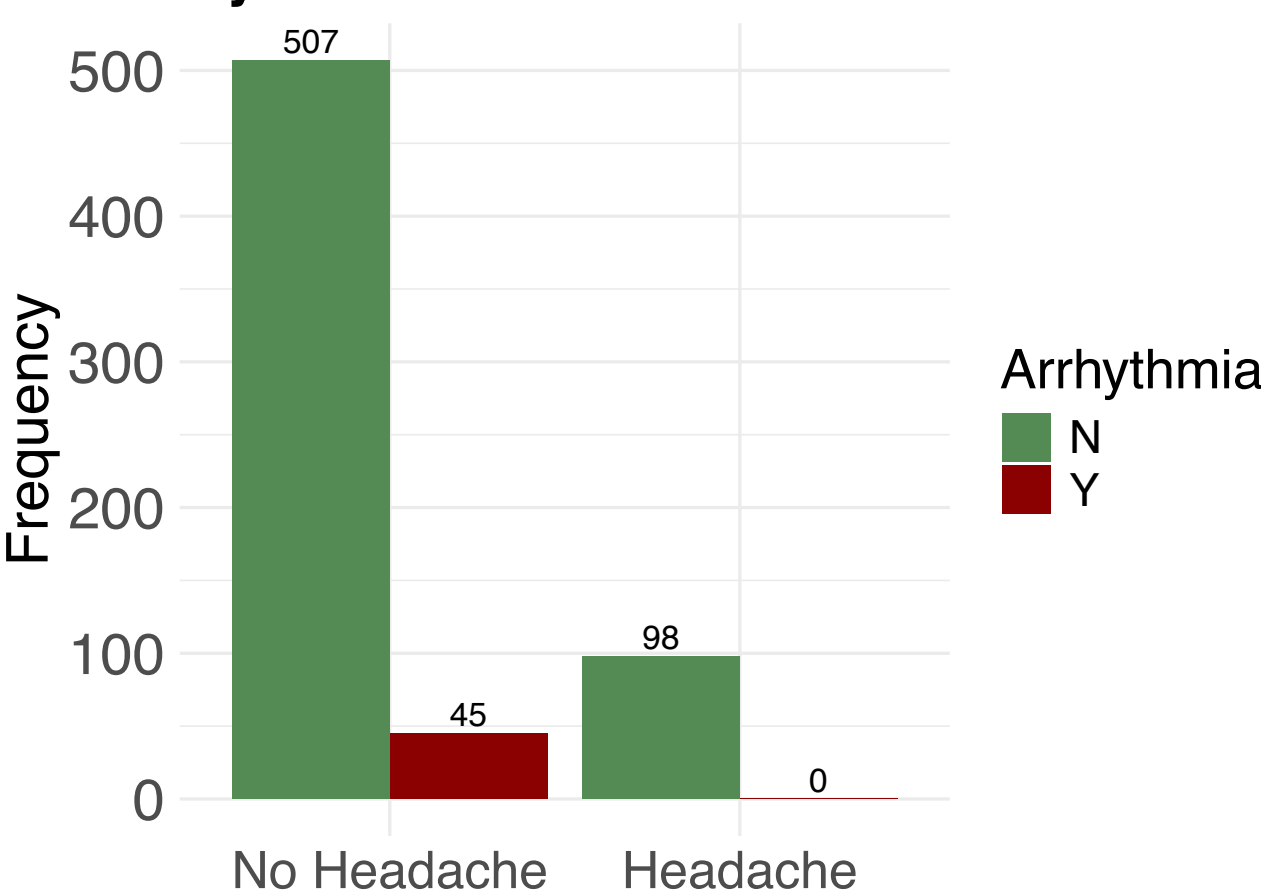
# Fisher's Exact Test

## Test of association for low frequencies

Arrhythmia vs. Unknown Indication

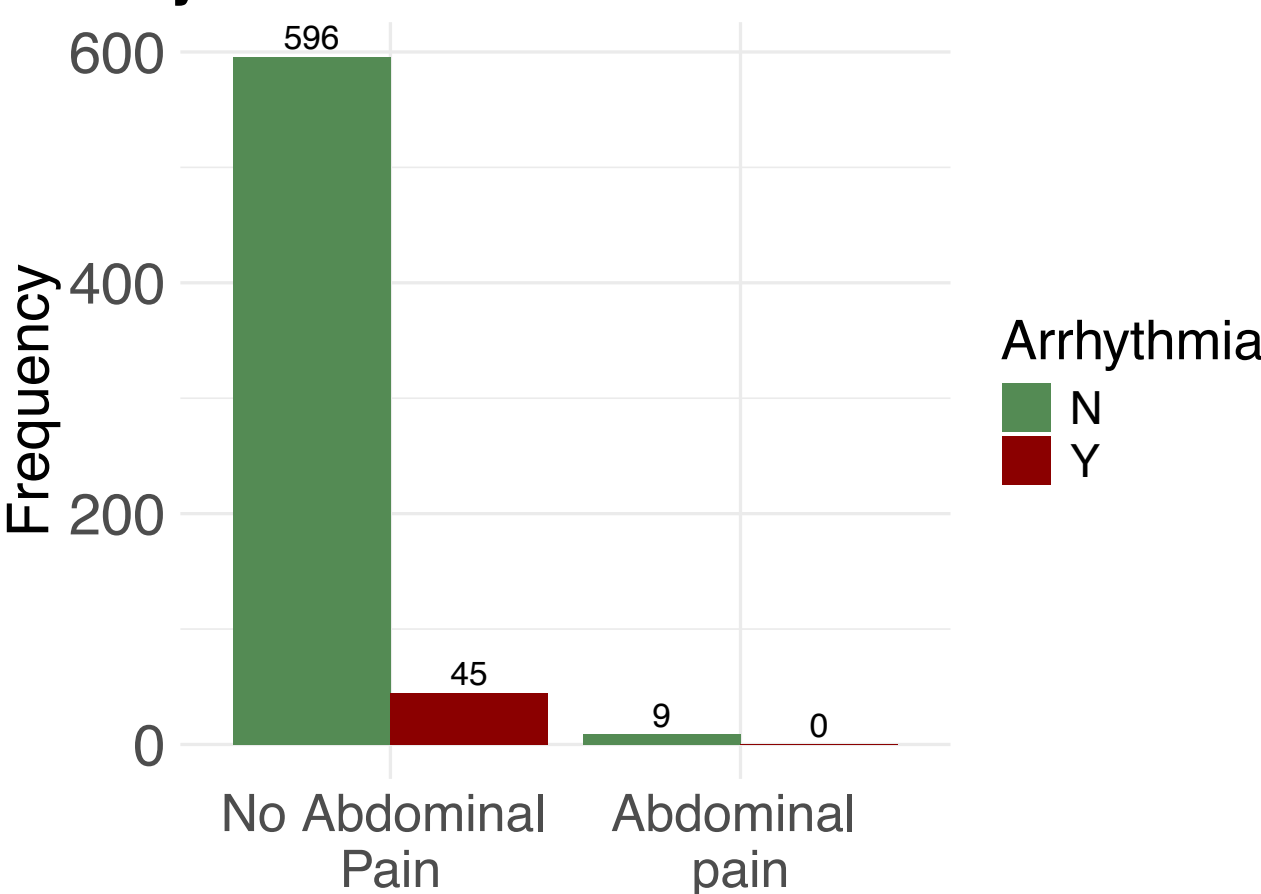


Arrhythmia vs. Headache Indication

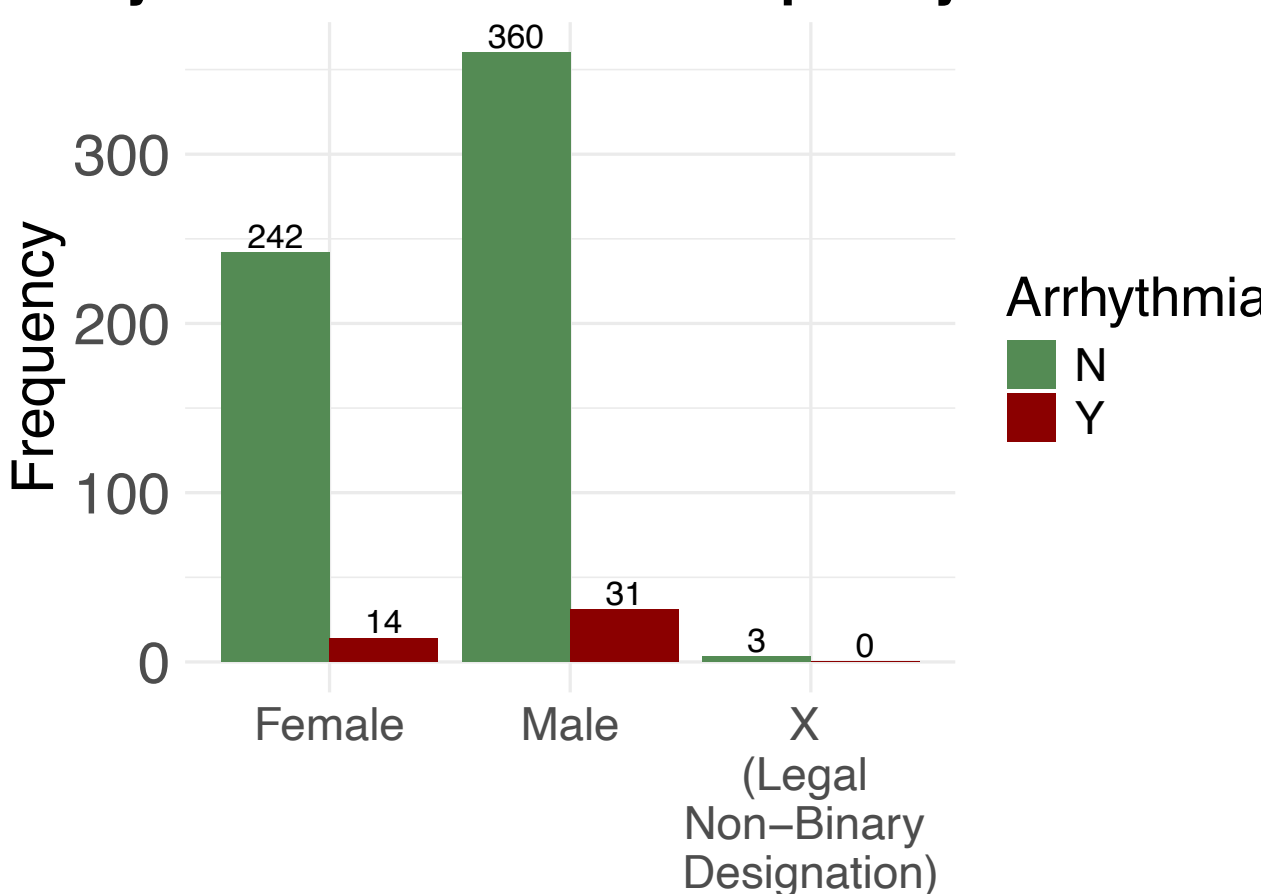


Variable	Odds Ratio	95% CI	p-value
Unknown	0	(0, 3.8)	0.6154
Headache	0	(0, 0.5)	0.0008163
Abdominal pain	0	(0, 6.9)	1
Sex (X)	0	(0, 33.0)	1

Arrhythmia vs. Abdominal Pain Indication



Arrhythmia Occurrence Grouped by Patient Sex



Headache associated with *decreased* risk of arrhythmia

# Inference using QTc measurements

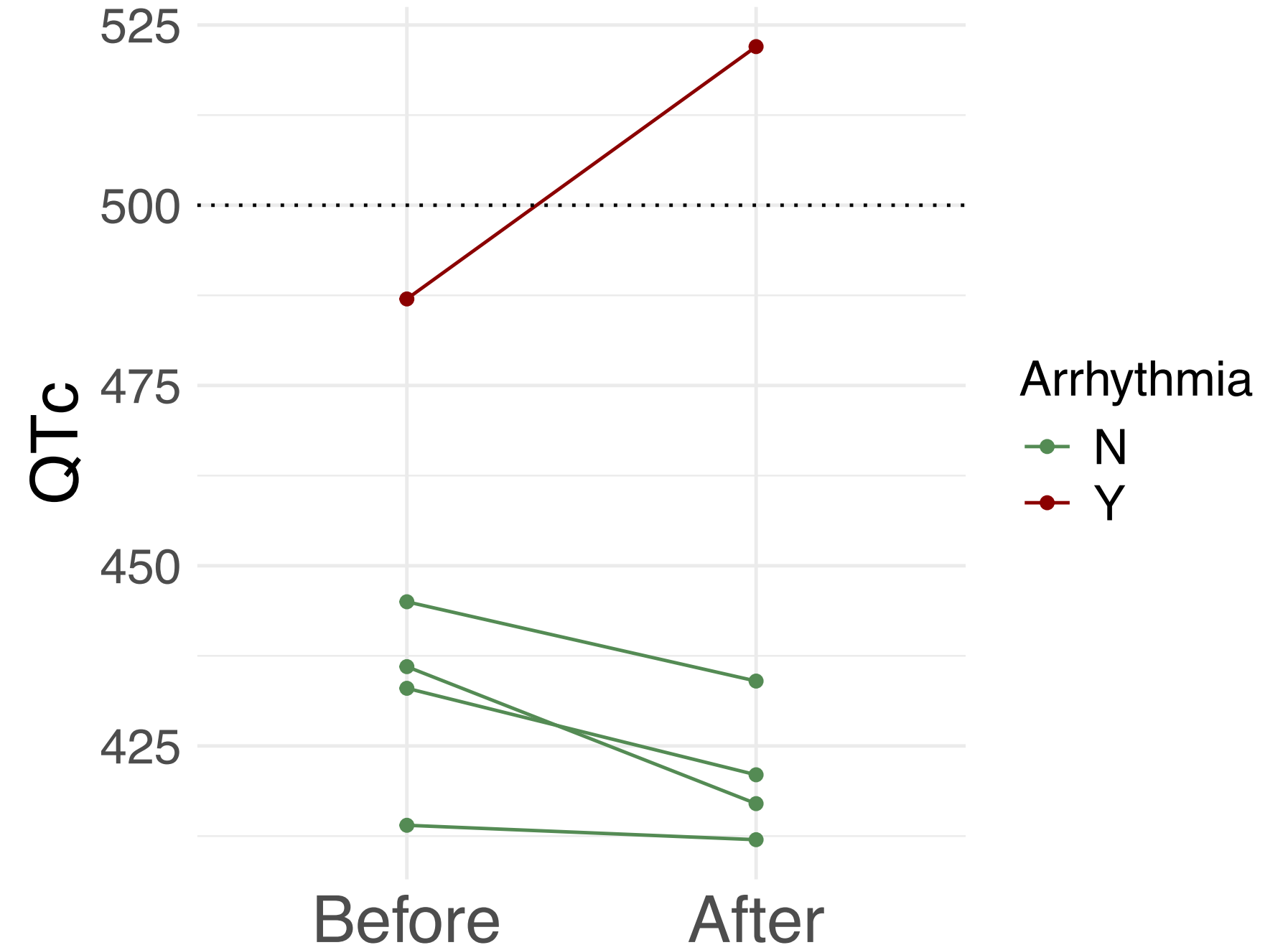
## among agitated patients

- Higher QTc  $\longrightarrow$  higher risk of arrhythmia

### Note:

- Unfortunately, don't have many paired observations for QTc(before) and QTc(after)
- EKG reading to measure QTc requires agitated patients to be calm, so QTc (before) measurements may be biased toward lower values

Paired QTc Measurements for 5 Agitated Patients



# Decision Trees

- **Imbalanced data:** 7% had an arrhythmia

- **Predictors:**

Legal Sex

Age

Headache

Abdominal Pain

Vomiting

Agitation

Unknown

Total Dose (mg)

## Training Data

70% of patients  
*with* arrhythmia

70% of patients  
*without* arrhythmia

## Testing Data

30% of patients  
*with* arrhythmia

30% of patients  
*without* arrhythmia

# Decision Trees

## Without Balancing



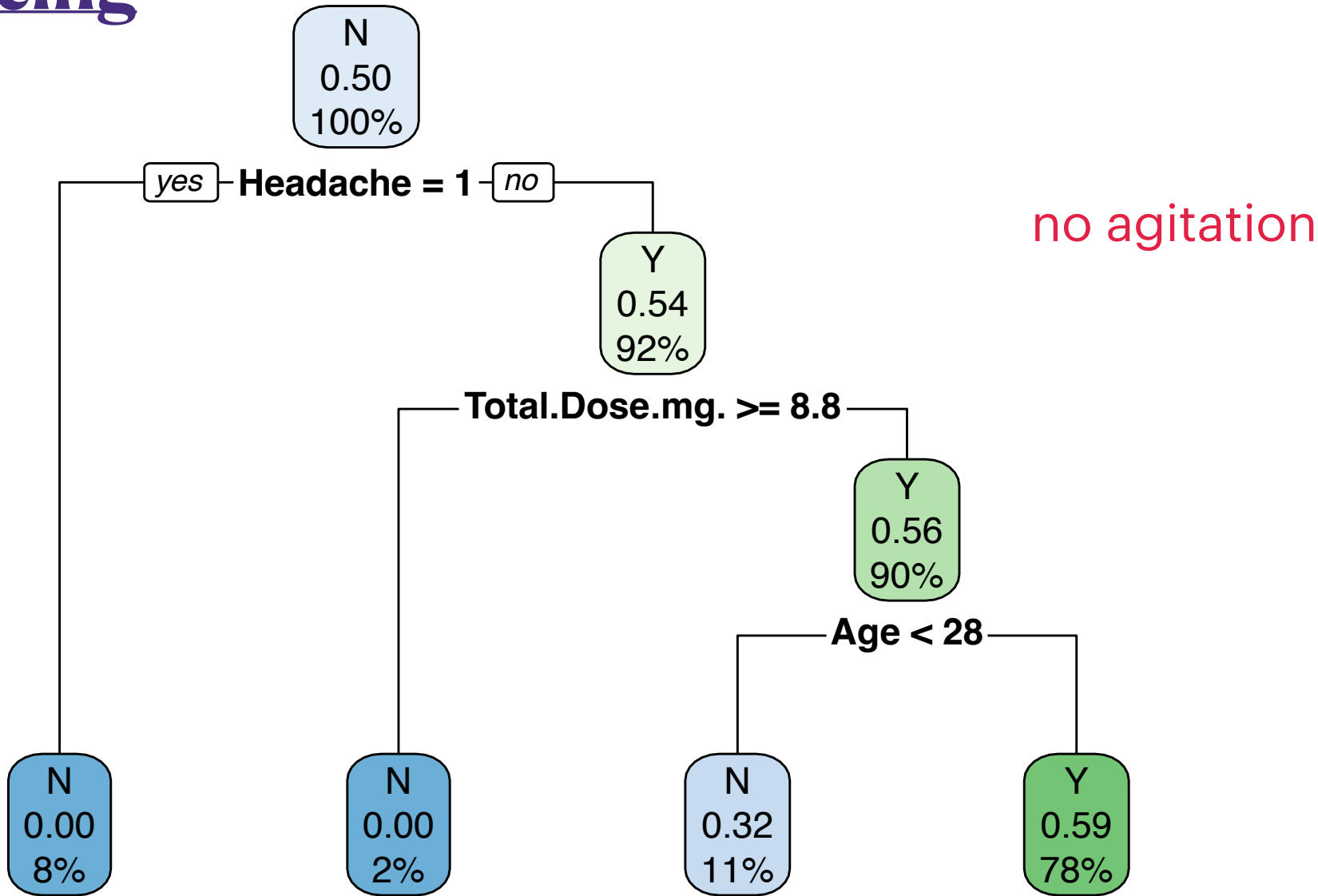
**total misclassification error:**  
6.7%

**arrhythmia misclassification error:**  
100%

**contingency table:**

predictions	N	Y
N	181	13
Y	0	0

## With Balancing



**total misclassification error:**  
64.4%

**arrhythmia misclassification error:**  
23.1%

**contingency table:**

predictions	N	Y
N	66	3
Y	115	10

# Conclusion

- Agitation associated with *increased* risk of arrhythmia when administered droperidol
- Headache associated with *decreased* risk of arrhythmia when administered droperidol

## Next Steps

- For now, perhaps limit use of droperidol for agitated patients
- Repeat QTc inference w/ paired data
  - Obtain EKG measurements, *both* before and after, administering droperidol to agitated patients to isolate effect of droperidol (probably impossible)
- Collect similar data for patients in the ER that do not receive droperidol to compare arrhythmia occurrence with and without droperidol use