# EFFECTS OF THE BETA-ADRENERGIC ANTAGONIST PROPRANOLOL ON ADAPTIVE AND PROBLEM BEHAVIOR AND RELATIONSHIP WITH HEART RATE VARIABILITY IN PATIENTS WITH AUTISM SPECTRUM DISORDER

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## INTRODUCTION

- Autism spectrum disorder (ASD) is a neurodevelopmental disorder that
  affects many individuals in the United States. ASD is characterized by
  persistent deficits in social communication and social interaction across
  multiple contexts.
- Children with ASD have high anxiety levels that affects their social communication and social interaction abilities. As such, it is important to find ways to help increase socially appropriate behavior in ASD and maximize their social communication and interaction.
- Research indicates that propranolol, a pharmaceutical drug that blocks
  the brain and body's use of norepinephrine both centrally and
  peripherally, reducing noradrenergic system activity. As a result of
  noradrenergic blockade, propranolol decreases blood pressure and
  reduces anxiety
- A previous study from our team found that a single dose of propranolol significantly increased conversational reciprocity in a sample of patients with ASD.
- Here, we examined the effects of serial doses of propranolol for 12 weeks on the effects of social severity and anxiety and fears in patients with ASD.
- Further, we examined the effects of propranolol on heart rate variability (HRV).
- We hypothesized that propranolol would decrease both social severity and anxiety and fears while also increasing HRV.

# **METHODS**

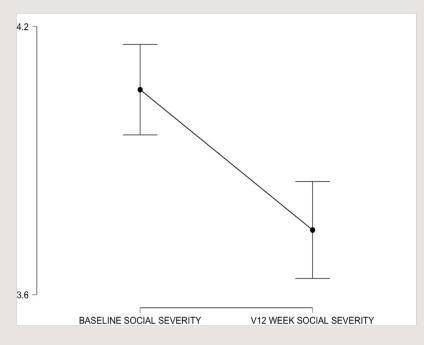
- Participants: 51 individuals diagnosed with ASD (M age = 14.02, SD = 4.7, range = 7-23, 10 females).
- Propranolol Administration:
- Participants received propranolol for 12 weeks in an unblinded manner. The drug was titrated slowly to ensure it was tolerated well. Patients aged 7-14 were titrated according to weight, and those 15-24 were titrated up to 100mg as follows:
- Week 1: 40 mg propranolol (1 capsule, nightly)
- Week 2: 80 mg propranolol (2 40mg capsules, morning & night)
- Weeks 3 12: 100 mg propranolol (3 capsules, 40 mg/morning, 20mg/afternoon, & 40mg/night)

### RESULTS

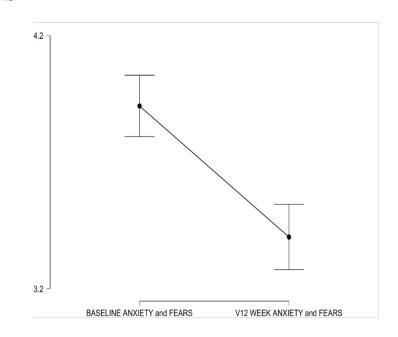
- Preliminary results indicate that anxiety reduced significantly, and social severity decreased significantly after 12 weeks of propranolol (See Figures 1 & 2).
- Heart rate variability significantly increased after 12-weeks of taking propranolol.

# **GRAPHS**

# BASELINE SOCIAL SEVERITY - 12 WEEK SOCIAL SEVERITY BASELINE ANXIETY and FEARS - 12 WEEK ANXIETY and FEARS



**Figure 1.** There was a significant decrease in social interaction severity scores between baseline (M = 4.06, SD = 0.76) compared to 12-week social interaction severity scores (M = 3.75, SD = 0.82), t(46) = 4.47, p < 0.001. See Figure 1.



**Figure 2.** There was a significant decrease in anxiety and fear severity scores between baseline (M = 3.92, SD = 0.89) compared to 12-week anxiety and fear severity scores (M = 3.40, SD = 1.01), t(46) = 5.40, p < 0.001. See Figure 2.

#### **Paired Samples T-Test**

|                        |                               | t      | df | p      | Cohen's d |
|------------------------|-------------------------------|--------|----|--------|-----------|
| BL_Mean RR             | - 12 WEEK Mean RR             | -5.001 | 26 | < .001 | -0.962    |
| BL_Mean HR             | - 12 WEEK Mean HR             | 4.876  | 26 | < .001 | 0.938     |
| BL_Min HR              | - 12 WEEK Min HR              | 4.145  | 26 | < .001 | 0.798     |
| BL_Max HR              | - 12 WEEK Max HR              | 3.845  | 26 | < .001 | 0.740     |
| BL_STD HR              | - 12 WEEK STD HR              | 0.486  | 26 | 0.631  | 0.094     |
| BL_SDNN                | - 12 WEEK SDNN                | -2.590 | 26 | 0.016  | -0.498    |
| BL_RMSSD               | - 12 WEEK RMSSD               | -2.771 | 26 | 0.010  | -0.533    |
| BL_NN50                | - 12 WEEK NN50                | -2.692 | 26 | 0.012  | -0.518    |
| BL_pNN50               | - 12 WEEK pNN50               | -2.942 | 26 | 0.007  | -0.566    |
| BL_RR Triangular Index | - 12 WEEK RR Triangular Index | -1.946 | 26 | 0.062  | -0.375    |
| BL_TINN                | - 12 WEEK TINN                | -2.680 | 26 | 0.013  | -0.516    |
| BL_Stress Index        | - 12 WEEK Stress Index        | 2.816  | 26 | 0.009  | 0.542     |

**Table 1.** Paired Sample T between baseline and 12-week HRV parameters.

|                             | N         | Mean    | SD      | SE           |
|-----------------------------|-----------|---------|---------|--------------|
| BL_Mean RR                  | 32        | 667.401 | 106.498 | 18.826       |
| 12 WEEK Mean RR             | 28        | 787.040 | 132.136 | 24.971       |
| BL_Mean HR                  | 32        | 92.212  | 15.274  | 2.700        |
| 12 WEEK Mean HR             | 28        | 78.504  | 14.344  | 2.711        |
| BL_Min HR                   | 32        | 80.396  | 15.920  | 2.814        |
| 12 WEEK Min HR              | 28        | 69.151  | 13.222  | 2.499        |
| BL_Max HR                   | 32        | 106.660 | 14.957  | 2.644        |
| 12 WEEK Max HR              | 28        | 94.465  | 14.869  | 2.810        |
| BL_STD HR                   | 32        | 4.480   | 2.163   | 0.382        |
| 12 WEEK STD HR              | 28        | 4.264   | 1.813   | 0.343        |
| BL_SDNN                     | 32        | 34.549  | 19.500  | 3.447        |
| 12 WEEK SDNN                | 28        | 44.404  | 23.864  | 4.510        |
| BL_RMSSD                    | 32        | 29.763  | 21.065  | 3.724        |
| 12 WEEK RMSSD               | 28        | 46.479  | 33.651  | 6.359        |
| BL_NN50                     | 32        | 43.563  | 57.129  | 10.099       |
| 12 WEEK NN50                | 28        | 74.571  | 67.167  | 12.693       |
| BL_pNN50                    | 32        | 11.413  | 15.312  | 2.707        |
| 12 WEEK pNN50               | <b>28</b> | 22.429  | 21.590  | <b>4.080</b> |
| BL_RR Triangular Index      | 32        | 8.497   | 4.201   | 0.743        |
| 12 WEEK RR Triangular Index | 28        | 9.916   | 4.270   | 0.807        |
| BL_TINN                     | 32        | 176.781 | 95.022  | 16.798       |
| 12 WEEK TINN                | 28        | 219.107 | 96.430  | 18.224       |
| BL_Stress Index             | 32        | 16.901  | 9.916   | 1.753        |
| 12 WEEK Stress Index        | 28        | 12.186  | 6.933   | 1.310        |

Table 2. Means standard deviations and standard errors for HRV variables.

# GRAPHS CONT....

# **CONCLUSION**

- Preliminary analyses show that anxiety significantly decreased and social severity significantly decreased after taking propranolol for 12 weeks.
- The change in adaptive behavior and problem behavior is associated with the change in their heart rate variability.
- After taking propranolol for 12 weeks HRV significantly increased in participants.
- This indicates that participants are able to function at high levels which helps improve their quality of life and overall, makes it easier for them to function in their daily lives..