Quantifying the Effectiveness of Dr. Bredesen's ReCODE Protocol in Alzheimer's Disease

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> Alzheimer's Disease currently has no cure, although there are various drugs to treat the symptoms of cognitive decline. However, Dr. Dale Bredesen claims to have discovered a way to prevent and reverse Alzheimer's Disease, with a protocol of interventions that he termed "ReCODE". To test the effectiveness of the ReCODE protocol, researchers collected patient reviews from the books off Amazon, rated the reviews on a 1-5 scale of patient cognitive change, and analyzed the results using a Chi-square test for goodness of fit. Statistical analysis rejected the null hypothesis and accepted the alternative hypothesis that the ReCODE protocol was effective in the treatment of Alzheimer's disease.

1 Introduction

For many decades, Alzheimer's Disease has been assumed 2 to be a fatal disease with inevitable decline in memory 3 and cognition until the patient's death. Alzheimer's is a 4 currently incurable neurodegenerative disease that causes 5 memory loss, and mental decline that impairs daily life. 6 In the brain, Alzheimer's is characterized by amyloid beta 7 plaques and tau neurofibril tangles [1]. To combat the se-8 vere illness, patients often choose to use lecanemab and 9 aducanumab-two drugs that have the ability to temporar-10 ily delay the progression of Alzheimer's [1]. However, 11 rather than use drugs to restrain the cognitive disease, Dr. 12 Bredesen explains in detail about the interventions in Re-13 CODE in two books: The End of Alzheimer's and The 14 End of Alzheimer's Program [3, 4, 2]. In these books, 15 Dr. Bredesen explains that there are three major causes 16 of Alzheimer's: inflammation, loss of trophic support, and 17 exposure to toxic compounds [3]. Instead of solely focus-18 ing on amyloid beta, the ReCODE protocol addresses and 19 underlines essential information about the roles of nutri-20 tion, exercise, sleep, stress management, supplements, and 21 hormones in the escalation of Alzheimer's [2, 3]. 22

Dr. Bredesen outlines specific tests for risk factors, including: ApoE4 status, obesity and diabetes, cardiovascular diseases such as hypertension, heart attack or stroke, systemic inflammation as manifested in chronic pain, fatigue, GI problems and mood problems, diets that are rich

in animal proteins and starch, lack of antioxidant supple-28 ments, chronic stress, and low activity level [3]. Examples 29 of certain courses of actions to take are stated in the Re-30 31 CODE protocol. They include the intake of supplements, such as vitamin D, and daily fasting, which would induce 32 ketosis, both of which have shown to be effective in im-33 proving the patient's cognitive functions [10]. Dr. Bre-34 desen concludes that the ReCODE protocol is successful 35 and more importantly, impactful, as it was able to reverse 36 the worsening of cognitive functions in more than 200 pa-37 tients [10]. 38

Nevertheless, many people voiced concerns about the ef-39 fectiveness of the method due to its technicality while oth-40 ers reinforced its credibility through positive reviews. In 41 this research endeavor, we assessed the effectiveness of 42 the ReCODE program by quantifying the experience of 43 patients through their reviews of the books on Amazon. 44 The reviews were carefully selected for its credibility in 45 participation, processed through a subjective rating system 46 and further analyzed using a Chi-square test. Our results 47 demonstrate a successful rejection of the null hypothesis, 48 confirming the effectiveness of the ReCODE program. 49

50 2 Methodology

In order to collect credible data and reach conclusions for our research we collected data from the two books and gathered different opinions on how well the ReCODE pro-

gram itself works [3, 4]. In order to find out the effec-54 tiveness of the program, we looked for patient reviews that 55 contained specific information on how the reader's symp-56 toms improved or did not improve, not those about how 57 well the books were written or feedback on the program's 58 hypothetical logic.

First, we collected reviews of the two books, book 1 re-60 ferred to as The End of Alzheimer's and book 2 as The End 61 of Alzheimer's Program on Amazon. The first book had 62 ~5000 ratings, but only ~1000 reviews and the second book 63 had ~2000 ratings and ~100 reviews [3, 4]. To efficiently 64 collect data, each researcher divided the work to gather the 65 viable reviews of each book onto a spreadsheet which all 66 the researchers could access. Our collection criteria was 67 that the patient had to have tried the protocol, or describe 68 the person who tried the protocol. After collecting all pa-69 tient ratings of both books, researchers rated each review 70 on a scale of 1 to 5 uninfluenced by the original patient's 71 star rating: 1 being significant deterioration of symptoms, 2 72 being mild deterioration of symptoms, 3 being no change, 73 4 being mild improvement of symptoms, and 5 being sig-74 nificant improvement of symptoms as indicated in the cus-75 tomer review. Multiple researchers rated each patient re-76 view so that the average of the ratings allowed for an unbi-77 ased scoring of the reviews. 78

Researchers then did statistical analysis to quantify the 79 effectiveness of the ReCODE program through a Chi-80 Square goodness of fit test on the null hypothesis to reach 81 a conclusion on the effectiveness of the program based on 82 patients who followed the book's protocol and described a 83 change in their symptoms of cognitive decline. 84

3 **Results** 85

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Different researchers had different 1-5 ratings on the pa-86 tients level of improvement, and as a result the average 87 rating contained decimal values. Out of the ~1000 writ-88 ten reviews for book 1, there were 34 viable reviews that 89 fulfilled our selection criteria [3]. Out of the ~100 written 90 reviews for book 2, there were 24 viable reviews that ful-91 filled our selection criteria [4]. Across both books, there 92 were 58 total viable reviews collected and 9 rating values 93 for our rating scale from 1-5 on patient improvement. The 94 histogram below displays our collected data. 95

On our rating scale of 1-5, a rating of 3 marks the mid-96 dle ground of no symptom change. If the ReCODE proto-97 col was ineffective, the ratings would be equally distributed 98 across all star levels since there would be no association 99 with cognitive improvement. On the other hand, partici-100 pants whose cognitive decline improved from using the 101 ReCODE protocol received a rating greater than 3. Out 102

of the 58 reviews collected, 43 of our ratings were higher 103 than 3, meaning approximately 74.13% of the participants' 104 cognitive decline benefited from the ReCODE protocol. 105



Figure 1: Histogram of Observed Rating Values

We performed a Chi-square goodness of fit test to evalu-106 ate whether the observed distribution significantly deviates 107 from the expected equal distribution of ratings. The hy-108 potheses were set as follows: 109

Null Hypothesis (H_0) : People with cognitive decline 110 who tried the ReCODE protocol showed no significant 111 improvement in their symptoms, and the data will have 112 equally distributed ratings. 113

Alternative Hypothesis (H_A) : People with cognitive de-114 cline who tried the ReCODE protocol showed significant 115 improvement in their symptoms and the ratings do not fol-116 low a uniform distribution. 117

The data included 9 groups, and with an expected equal 118 distribution, the expected frequency for each group is the 119 total number of reviews divided by 9: 120

Expected =
$$\frac{11+5+3+15+2+7+10+5+0}{9} = 7.25$$
 (1)

The Chi-square number is calculated with the formula 121 where O_i is the observed frequency in group *i* and E_i is the 122 expected frequency in group *i*: 123

$$\chi^2 = \frac{\sum (O_i - E_i)^2}{E_i} \tag{2}$$

The degrees of freedom are calculated as: 124

$$df = Number of Groups - 1 = 8$$
 (3)

Using the above formulas, the Chi-square calculated 125 statistic is about 26.22, and with its corresponding df of 8, 126 the p-value is approximately 0.000965. Since the p-value 127 is smaller than our alpha level of 0.05, we reject the null 128 hypothesis and accept the alternative hypothesis that peo-129 ple with cognitive decline who tried the ReCODE protocol 130 showed significant improvement in their symptoms, and the 131 ratings do not follow a uniform distribution. 132

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4 Discussion

Using our analysis, which resulted in a p-value less than the
alpha level of 0.05, we were able to reject the null hypothesis and accept the alternative hypothesis that the ReCODE
protocol was effective in the improvement of Alzheimer's
symptoms such as cognitive decline.

The ReCODE program specifically, involves the use of a 139 use of a plant-rich ketogenic diet that prioritizes low mer-140 cury, wild-caught seafood, and pastured eggs, a long daily 141 fast; an exercise program comprised of both aerobic and 142 anaerobic exercise combined with the suggestion to avoid 143 prolonged periods of sitting; 7-8 h of quality, restorative 144 sleep; a stress management program emphasizing regu-145 lar deep breathing breaks with meditation; regular brain 146 training, maintenance of social connectivity; an avoidance 147 of toxins along with instructions to upregulate detoxifica-148 tion [3]. Daily supplements include probiotics, omega-3s, 149 multivitamin, and antioxidants [3]. Many reviews com-150 mented on their decreased, and some reversed, cognitive 151 decline after following Dr. Bredesen's program, includ-152 ing clearer thinking and better memory. Some described 153 elevated moods or a decrease in depression after follow-154 ing the program. Others also mentioned improved physical 155 functioning and sleep. For those who experienced signifi-156 cant improvements, many had followed Dr. Bredesen's Re-157 CODE diet, cutting out processed foods, eating vegetables, 158 fresh fruits and minimal carbs to stay in ketosis. Others 159 also mentioned taking supplements and exercise. 160

Although we were able to support our alternative hypoth-161 esis, there are some limitations to this study. The method 162 of sampling used has aspects of convenience sampling due 163 to the accessibility of Amazon book reviews. Because 164 the data was collected from Amazon reviews of Dr. Bre-165 desen's book, this excluded those who may have been rec-166 ommended the ReCODE program by a health professional 167 or those who followed the ReCODE program but did not 168 make a review or buy the book off of Amazon. 169

Dr. Bredesen's approach to preventing Alzheimer's, 170 slowing cognitive decline, and reversing neurodegenerative 171 diseases is a multifaceted approach that focuses on over-172 all health instead of solely focusing on associated factors 173 such as amyloid beta [3]. The treatment suggestion of fol-174 lowing a plant rich ketogenic diet is similar to the Mediter-175 ranean diet, which research has shown to be associated with 176 decreased cognitive decline [6, 9]. Both diets emphasize 177 eating healthy fats and limiting processed food, although 178 a ketogenic diet is more restrictive towards carbohydrates 179 [5]. Other aspects of the ReCODE protocol, such as aero-180 bic and anaerobic exercise as well as meditation are shown 181 to be associated with Alzheimer's prevention [8, 7]. The 182 ReCODE protocol utilizes lifestyle factor changes to pre-183 vent and combat Alzheimer's disease progression. 184

185 **5** Conclusion

In The End of Alzheimer's and The End of Alzheimer's Pro-186 gram, Dr. Bredesen outlines his ReCODE protocol in the 187 prevention and treatment of Alzheimer's disease. Based on 188 the statistical analysis, we reject the null hypothesis and 189 accept the alternative hypothesis that the reCODE proto-190 col was effective in the improvement of Alzheimer's symp-191 toms such as cognitive decline. In future studies examining 192 the effectiveness of the ReCODE protocol on cognitive de-193 cline, research should further investigate the relationship 194 between the specific aspects of the ReCODE protocol and 195 Alzheimer's disease. 196

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