**Patient preferences for treatment in steroid resistant ulcerative colitis – a discrete-choice experiment**

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**Background and aim:** Understanding treatment preferences in those patients who are not responding to corticosteroids for ulcerative colitis is important in informing treatment choices. This study aimed to assess the relative importance of treatment characteristics to patients by conducting a discrete-choice experiment.

**Methods:** Patients completed the questionnaire online. All data were collected between September and December 2020. Participants were shown 13 discrete-choice experiment tasks – a series of side-by-side comparison of competing, hypothetical treatment characteristics and asked to select a preferred treatment. Survey responses were analysed using descriptive statistics and regression analyses.

**Results:** 115 patients completed the study. Patient preferences were strongest for treatments with a lower chance of side effects, this attribute had the most influence on the choice of treatment patients preferred. The second most important attribute was improvement in maintaining remission. Conversely, route and frequency of administration were least important on the choice of treatment patients preferred. Respondents were willing to make trade-offs and accept treatment benefits to compensate them for receiving a treatment with a less desirable attribute level. Participants were willing to accept a larger benefit of 45% improvement in maintenance of remission to accept a treatment with higher probability of side-effects. The benefit required was smaller with 10.35% improvement in remission required to accept a treatment with lower probability of side-effects.

**Conclusion:** Quantifying preferences helps to identify and prioritise treatment characteristics that are important to patients. The results highlight the importance of careful discussion of side effects, including the magnitude of risk, using visualisation tools during a patient consultation to support decisions.

**Key words:** Ulcerative colitis; Patient preference; discrete-choice experiment

## **INTRODUCTION**

Ulcerative colitis is a chronic inflammatory condition that is characterised by debilitating symptoms of diarrhoea, rectal bleeding, abdominal pain, fatigue and impaired quality of life [1]. Patients experiencing a disease flare are often treated with corticosteroids [2]. However, just under 50% of patients do not respond fully to steroids [3]. This requires further treatment and individuals are therefore faced with selecting a treatment in partnership with their clinician.

Treatment options include small molecules, biologic agents, thiopurines or surgery [4]. Besides the effectiveness of treatments, other considerations may be important for patients when opting for treatment, including drug side effects, travel requirements to receive infusions and time off work [5]. Treatment efficacy may also incorporate time to response, steroid free remission, and longer-term treatment success. The best treatment option for patients will depend on how they value the characteristics of treatments. Understanding these patient values is important to choose a treatment that closely aligns with these values [6-8]. This information can then be used to facilitate the shared decision making process between the patient and clinician [9-11].

Discrete-choice experiments (DCEs) have been increasingly used to identify patient preferences in healthcare research [12]. DCEs quantify the strength of preferences for the features of hypothetical treatments that are most important to patients. While studies have applied the DCE method to different types of ulcerative colitis [13-17], evidence on preferences for the steroid resistant population is limited. Consequently, this study aimed to understand the relative importance of treatment characteristics to patients with steroid resistant ulcerative colitis by conducting a DCE.

## **METHODS**

### **Questionnaire design**

The first section of the survey contained the DCE tasks (see Supplementary Data 1). At the start of the DCE section, we described a steroid resistant ulcerative colitis scenario for participants to imagine. Then participants were shown a series of side-by-side comparisons of competing treatment profiles and were asked to select their preferred treatment profile (See Figure 1). The reason for describing an ulcerative colitis scenario for participants to imagine rather than asking patients to think about their own situation was to allow all patients to answer the DCE based on the same baseline. This context setting is important because the clinical presentation of ulcerative colitis varies from patient to patient and otherwise would have introduced significant heterogeneity.

The DCE was developed by reviewing the literature, conducting qualitative interviews with patients, and consulting patient representatives. We conducted thirty-three interviews with patients to identify key treatment characteristics patients consider important when selecting a treatment [18]. Thematic analysis of the qualitative interviews generated eight themes, which were ranked by the four patient and public involvement (PPI) group members. These members were patients with ulcerative colitis who have had medical or surgical treatments. Using a dot-voting technique[19], the patients were given sixteen dots to distribute across the eight themes, where themes with the most dots revealed the most desirable treatment characteristics. This process helped to convert and reduce the eight themes to attributes. Upon discussions with the PPI group, one theme around the need for regular monitoring was dropped as this was deemed the least important theme compared to the others. Themes that were similar or correlated were merged. For example, route and frequency of administration were merged to create one single attribute. Similarly, quality of life and inducing a treatment response were merged as the two attributes were highly correlated to create another single attribute. The final five attributes focused on effectiveness, remission, speed of response, treatment administration and safety of the treatment (Table 1). The levels for the attributes were selected to reflect plausible values from the published clinical trials literature. For the side-effects attribute we used the levels described in medical labels (i.e. very common, common) as patients are familiar with it.

The DCE questions were generated using the NgeneTM software[20]. The combination of attributes and levels (three attributes with three levels each and two attributes with four levels each) produced 432 possible treatment profiles, so to create a manageable DCE questionnaire a computer software was used to select the treatment profiles to be included in the questionnaire (fractional factorial design). The software produced a D-optimal design which followed the principles of minimum overlap, orthogonality, and level balance[21]. Only one version of the survey was generated using NgeneTM which contained 12 unlabelled DCE questions. One additional dominant question that was logically better was also included to test whether participants understood the DCE task [22]. These 13 questions were displayed to the participants in a random order using the QualtricsTM survey platform. To create a realistic choice, participants were not given an opt-out option because treatment for ulcerative colitis is necessary to improve their length and quality of life.

Section two of the survey involved a ranking exercise where patients were asked to rank four commonly used treatments (adalimumab, infliximab, tofacitinib, and vedolizumab,) in order of preference from one to four. To aid this task, we provided comprehensive details of the treatments which included information on the effectiveness of the drug, speed of response to treatment, route of administration, side effects, and whether concomitant medication is needed [23]. These treatment descriptions were developed using published literature with clinical input from the study team and presented to participants in a randomised order to reduce question order bias (see Supplementary Table 2).

In section three, we gathered sociodemographic details and the respondents’ personal history and severity of ulcerative colitis. The survey included two validated instruments, the IBD-Control-8 questionnaire and the EQ-5D-5L questionnaire. The IBD-Control-8 questionnaire captured disease control and impact from the respondents’ perspective. It generated a summary score ranging from 0 representing worst control to 16 representing best control of disease and a score of 13 and above represents quiescent disease (cronbach’s α=0.86) [24]. The EQ-5D-5L instrument captured respondents’ overall quality of life, generating a summary score between -0.59 to 1, where higher scores represented better quality of life [25]. Section four contained feedback questions about the survey.

To increase face validity, the survey was piloted by three patient representatives who were not involved in the first stage of the DCE design to check if it was feasible for patients to complete the survey. Upon receiving the pilot feedback, we simplified the way the DCE was presented by providing detailed explanations of the attributes at the start of the DCE exercise. A series of screens displayed instructions providing detailed descriptions of the attributes, for example we explained risks qualitatively in the instruction screen (i.e. a drug that is 60% effective means that if 100 people had the same drug for ulcerative colitis, for 60 people the treatment would be effective but for 40 people treatment would not be effective) [26] but in the DCE tables risks were shown numerically (i.e. 60 in 100, 60%) as it was quicker for the participants to read. In addition to the pilot, a review of responses was undertaken after 50 respondents completed the study to assess problems, including comprehension and dropouts.

### **Participants**

The study population included adults aged at least 18 years who had a diagnosis of ulcerative colitis. Participants were primarily recruited through two National Health Service (NHS) hospitals, in an outpatient (non-hospitalised) setting. Staff working in inflammatory bowel disease (IBD) clinics advertised the study by sending potential participants invitation letters. The study was also advertised on social media to recruit further participants from across the UK. If individuals decided to take part, they were able to access the online survey via the QualtricsTM platform and complete the survey after providing informed consent. We hoped to recruit up to 300 survey participants on the basis of precedence where the literature shows that DCE sample size ranges between 100 to 300 participants [27]. However, the minimum sample size required was N≥83.3 (N= 500 x (4 [maximum number of levels])/(2 [# of alternatives] x 12 [ # of tasks]) to estimate a model using the rule of thumb approach [28].

### **Statistical Analysis**

Descriptive statistics were performed to analyse demographic data, IBD characteristics of the respondents and to rank the order of importance of medications. We performed conditional logistic regression models to analyse the DCE task data. All attributes were included as independent variables; attributes were first included as categorical variables, but after checking for linear relationships through visual inspection and model fit, speed and remission attributes were treated as continuous variables and effectiveness, administration and side effects as categorical variables in the model.

Parameter estimates from the conditional model were also used to calculate minimum acceptable benefit and to calculate the change in probability of uptake from a baseline scenario where all attributes are set to their worst level and then improved each attribute one at a time [29] (see Supplementary Data 2 for an example calculation). All statistical analyses were conducted using STATATM v16.

### **Ethical Considerations**

The study was reviewed and approved by the NHS Research Ethics Committee – East Midlands Derby (19/EM/0011) and the Health Research Authority approved the research (IRAS ID: 255616).

## **RESULTS**

### **Study Population**

Seven hundred and twenty invitation letters were sent to eligible participants. Of the invited participants, 166 visited the QualtricsTM survey platform, 115 completed the survey and 5 people declined to complete the survey. Based on the respondents who visited the survey, the response rate was 69% however, based on the total invites sent, the response rate is 16% (see figure 5). All data were collected between September and December 2020. Table 2 presents the socio-demographic characteristics of the respondents and the clinical and treatment characteristics of the sample including current and previous medical treatments. Median age was 41 years and 52% of the responders were female. Our study sample is representative of the UK ulcerative colitis population in terms of gender and age [30]. The majority of patients were white, employed and educated to secondary school level or above. The median time from diagnosis was 10 years. The median IBD-Control-8 score was 13. Fifty-five percent of patients reported quiescent disease with a score of 13 and over. The median quality of life score was 0.77, which is lower than the UK general population mean quality of life score of 0.85 [31]. The majority of patients had previously received steroids (93%), thiopurines (70%) and biological therapies or tofacitinib (70%) - most commonly infliximab (51%).

### **Patient Preferences Weights**

All attributes had a significant influence on the choice of treatment patients preferred. Figure 2 shows the relative changes that make respondents more likely (positive coefficient) or less likely (negative coefficient) to take a treatment. A treatment having a lower likelihood of side effects (i.e. a change from very common to very rare (ß 2.937, p<0.001) or very common to uncommon (ß 2.260, p<0.001) or very common to common (ß 1.417, p<0.001)) strongly increased the likelihood of the respondent choosing a treatment. Similarly, higher levels of the maintenance of remission attribute strongly influenced respondents’ choice of treatment (1% increase in remission: ß 0.065, p<0.001). When induction of response was higher 50% or 60% (relative to 40%), respondents were also more likely to choose the treatment. However, patients were less likely to choose a treatment that takes longer to improve their symptoms (change per week: ß -0.145, p<0.001). Orally administered medication, taken daily (ß 0.848, p<0.001) and injections at home every 8 weeks (ß0.541, p<0.001) were preferred to infusions every 8 weeks at hospital. Notably, a change from infusion at hospital every 8 weeks to injection, every 2 weeks at home, was not significant (ß -0.029, p= 0.85) suggesting that patients found these two ways of administering the treatment comparable (see Supplementary Table 1).

### **Overall Rank Ordering of Attribute Importance**

To compare the attributes that had the most or least importance on the choice of treatments patients preferred we calculated the change in predicted probabilities from a baseline scenario compared to a new scenario. In the baseline scenario the treatment profile consisted of the worst possible levels for all the attributes, and in each new scenario, only one attribute level was improved at a time to find out how important that attribute was when selecting a treatment. Figure 3 shows the change in uptake rates of the improved treatment profiles. Larger changes in uptake rates indicate more favourable treatment attributes. A treatment with very rare side effects was ranked very favourably with a change in uptake rate of 90% (Figure 3). Comparing the change in probabilities across these scenarios, the attributes can be compared in terms of priority, with side effect attributes having the most influence on the choice of treatment patients preferred followed by a treatment with a 70% rate for maintaining remission. Conversely, route and frequency of administration were least important on the choice of treatment patients preferred.

### **Minimum Acceptable Benefit**

A minimum acceptable benefit was calculated to find the minimum change in benefit required by patients to compensate them for receiving a treatment with a less desirable attribute level, such as the chance of experiencing side effects. Table 3 shows the percentage point increase in maintenance of remission required to compensate participants for the worsening levels of the chance of getting side effects. Participants would require a larger 45% improvement in maintenance of remission to accept the worst change in risk of side effects, an increase from very rare to very common. When the increase in risk of side effect is smaller, from very rare to common, the participants are willing to accept 23% improvement in maintenance of remission. Even smaller 10.35% improvement in remission was required to choose to a treatment with change in risk from very rare to uncommon side-effect. Participants preferred route of administration was a pill but to switch from taking a pill to another mode of administration participants would require a larger improvement in maintenance of remission, specifically, a 13.39% increase in remission to administer an injection every two weeks or a 12.95% increase in remission to have an infusion every 8 weeks at hospital or a 4.69% increase in remission to administer an injection every eight weeks.

### **Ranking Treatments**

When participants were asked to rank the four biological treatments in order of preference, where a ranking of ‘1’ equated to most preferred treatment and ‘4’ equated to least preferred, the most preferred treatments were infliximab (38%) and tofacitinib (38%), followed by vedolizumab (17%) and adalimumab (6%) (Figure 4). The least preferred medication was adalimumab (54%), followed by vedolizumab (26%), infliximab (11%) and tofacitinib (9%). A subgroup (n=35) of biologic-naive patients were analysed separately and compared to those who were on biologics. For the biologic-naive subgroup, the most preferred treatment ordering changed, with tofacitinib ranked first (46%), followed by infliximab (37%), vedolizumab (11%) and adalimumab (6%). The least preferred treatment was adalimumab (60%), vedolizumab (26%), infliximab (9%), and tofacitinib (6%). This ordering was similar to the full sample. Moreover, comparing the ranking treatments with modelled predictions of the DCE tasks showed that infliximab (46%) was the most preferred treatment followed by tofacitinib (42%), vedolizumab (6%) and adalimumab (5%) (see Supplementary Table 3). While both methods yielded similar treatment preference rankings, the proportion of shares of treatment were different.

### **Understanding and engagement**

All participants completed all the choice tasks and for the remainder of the survey missing data was low and did not exceed 5%. The majority of (97%) survey participants answered the dominant choice question (where one treatment profile was logically better) correctly. Only four participants failed this internal validity test and the modelling results did not change when the four participants were excluded from the analysis (results not reported), which suggest that participants were able to understand the DCE tasks and the probabilities presented in them (see Supplementary Table 4). Moreover, participants’ feedback about the survey showed that the majority of respondents understood the DCE (92%) and ranking tasks (91%) and agreed that the information provided in the DCE task was appropriate (70%) (see Supplementary Table 4).

## **DISCUSSION**

This study was conducted to elicit patient preferences in the treatment of steroid resistant ulcerative colitis. In this cohort of patients with ulcerative colitis, choices of treatment were most influenced by fewer side effects compared to other attributes in their decision making. Improvement in maintaining remission was also valued but less than treatments with fewer side effects. It is possible that the conduct of the study during the COVID-19 pandemic influenced patients prioritising side effects over symptom improvement because of heightened fears about infections from hospital attendance or as a result of immunosuppressive medication. The result highlights the importance of careful discussion of side effects, including the magnitude of risk, for example using visualisation tools during a patient consultation to support decisions.

Infliximab and tofacitinib were rated as the most preferred treatments by this sample of patients. In the UK infliximab and adalimumab are the most widely used biologics. Tofacitinib is more recently available and prescribing is less frequent [32]. The findings suggest that the place of tofacitinib earlier in the treatment pathway for steroid resistant ulcerative colitis needs evaluation. This will be affected by factors including specific toxicities that were not part of the DCE such as thromboembolic disease and herpes zoster infection.

This is the first DCE study to focus on patients with steroid resistant ulcerative colitis conducted in the UK, though other studies have used DCEs to quantify patient preferences for IBD treatments [13-15, 33, 34]. Our finding, that treatments with fewer side effects are most important to patients, contrasts with the study by Bewtra, where patients with IBD were prepared to accept relatively high risks of lymphoma to avoid a disease relapse over the next 5 years [35]. Almario et al. also found that efficacy was the most important attribute for patients with ulcerative colitis, but that side effects were the key priority for patients with Crohn’s disease [15]. Boeri et al. reported that symptom improvement and risk of malignancy were the two most important attributes. Similar to our study, Boeri et al. reported that patients preferred oral modes of administration compared to subcutaneous or intravenous administration [14]. Our results also showed that less frequent administration of subcutaneous injects were preferred. Feedback from a patient panel indicated that frequent injections could be intrusive or a reminder of their illness.

This study has several strengths. The DCE was conducted to a high standard following published best practice guidelines [22]. The study included an internal validity check to test if participants understood the DCE task. Only four people failed the internal validity DCE question and the findings did not change when the four participants were excluded from the analysis. Also, the majority of participants reported that they understood the survey questions and the information provided in the DCE tasks were deemed sufficient. Additionally, the survey was developed by conducting qualitative interviews with patients [18] and accompanied by iterative rounds of piloting, with the help of patient and clinical representatives to ensure that information was accurate and easily understandable.

This study also has some limitations. First, while every effort was made to objectively describe the four treatments, interpretation of the ranking of treatments should take into consideration the uncertainty in the evidence base. While there is a significant amount of data assessing treatment safety and effectiveness there are several uncertainties in the evidence base [36]. The majority of the evidence comes from placebo-controlled efficacy trials with only limited head-to-head trials and some real-world evidence from registries [36]. This is further complicated by different definitions of outcomes, which are not always measured with the same instrument or scale[36]. Thus, developing an accurate treatment profile was challenging. Nevertheless, to ensure rigour and objectivity, the descriptions were compiled with key statistics which were calculated using systematic reviews and with input from experienced consultant gastroenterologists. Due to the significant heterogeneity that exists in the literature it is possible that different researchers could use different statistics, and until the gaps in the evidence base is met by well conducted head-to-head trials this problem is likely to persist.

Second, our intention was to recruit 300 participants, however, due to the pandemic we were unable to achieve this target as staff at NHS trusts were working under extreme pressures and the data collection period was shortened. Nevertheless, the sample size in our study was larger than the minimum sample required according to the rule of thumb approach to estimate a model and was sufficient to run the regression analyses and find statistically significant results. A larger sample size would also have enabled better subgroup analyses. We estimated models to assess preference heterogeneity using interaction terms for gender, ethnicity, age groups but we did not find statistically significant differences, probably due to the small sample size in the subgroups. The response rate in this study was also low. Possible reasons include the inability to recruit from a face to face clinic attendance with the switch to remote clinics during the CoViD-19 pandemic. It is be possible that the respondents filling the survey differ from non-respondents but we do not have data on those who did not complete the survey and therefore firm conclusions cannot be drawn.

A third challenge is surrounding the content of the DCE. The task involved trading hypothetical treatment profiles which were defined by five attributes. The selection of attributes contained a trade-off, the more attributes we include in the DCE task, the higher the likelihood that the task will become unwieldy contributing to poor quality of data[37]. Consequently, a few attributes were excluded from the DCE, for example the time burden involved with treatment monitoring or the option to have or avoid surgery. It is unclear the impact of excluding these attributes from the DCE and how respondents would behave differently when these factors are introduced in the real-world. The study was planned and undertaken before the availability of ustekinumab for ulcerative colitis in the UK and ustekinumab was therefore not included in the ranking exercise. Whilst this might have affected the uptake rates for particular agents, the drug attributes prioritised by patients are still applicable in terms of efficacy and toxicity.

Understanding patient preferences has implications for decision making. By quantifying the importance of different treatment characteristics, this study helps to facilitate discussions between patients and clinicians when reviewing treatment options. Results highlight the importance of clearly communicating risks of treatment using visualisation tools to have more informed discussions during a patient consultation. Also, selecting treatments that are effective but fast acting, and providing patients the opportunity to select how they would like to administer the treatments should be encouraged where possible. Improving communication between clinicians and patients about different treatment characteristics could lead to better therapeutic decision making. Treatments that align closely to patient preferences are likely to increase adherence to treatment, improve shared decision making and improve patient satisfaction [9-11]. Non-adherence to treatment not only impairs the patient recovery but also costs valuable resources [38]. Prioritising treatment characteristics that patients value when considering treatments could lead to better shared decision making and better outcomes. Future research could evaluate the trade-offs involved with choosing surgery vs medical therapies, facilitators for prescribing tofacitinib and improving ways to provide personalised treatments.

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**Data Availability:** The data underlying this article will be shared on reasonable request to the corresponding author

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## **Tables**

#### **Table 1:** Treatment attribute descriptions and levels

|  |  |
| --- | --- |
| **Treatment Attributes** | **Levels** |
| How effective the drug is at treating your symptoms: The drugs may improve or settle your symptoms (for example in reducing stool frequency and bleeding, or returning these to normal), improve your quality of life and make you feel better. | * 40% * 50% * 60% |
| Speed of response to treatment: Some drugs take longer than others to take effect. | * 6 weeks * 8 weeks * 14 weeks |
| Chance of your symptoms remaining improved after 12 months: After your initial symptoms improve, the drugs can help to control your symptoms over time. However, there is also a possibility that you may lose the improvement and develop a flare of your symptoms. | * 35% * 50% * 70% |
| Route and frequency of administration: How and where the medication would be taken is different according to which drug you take. | * a pill taken daily at home * a self-administered injection under the skin, administered every 2 weeks at home * a self-administered injection under the skin, administered every 8 weeks at home * an intravenous infusion (drip) administered every 8 weeks at hospital |
| Chance of experiencing side effects: Drugs can cause unwanted side effects. Common side effects include nausea, headache, skin rashes and mild infections. These often settle without treatment, can be easily treated, or are reversed if the drug is stopped. In rare cases, the drugs may cause severe side effects over a longer period of time. These include more severe infections (including tuberculosis and viral infections including the shingles virus), some cancers including, lymphoma (lymph gland cancer), blood clots in the leg (deep vein thrombosis) or lung (pulmonary embolism), and nervous system problems. The chance of experiencing severe side effects is very rare for all treatments. | * very common (may affect more than 10 in 100 people) * common (may affect up to 10 in 100 people) * uncommon (may affect 1 in 100 people or fewer) * Very rare (may affect up to 1 in 10,000 people) |

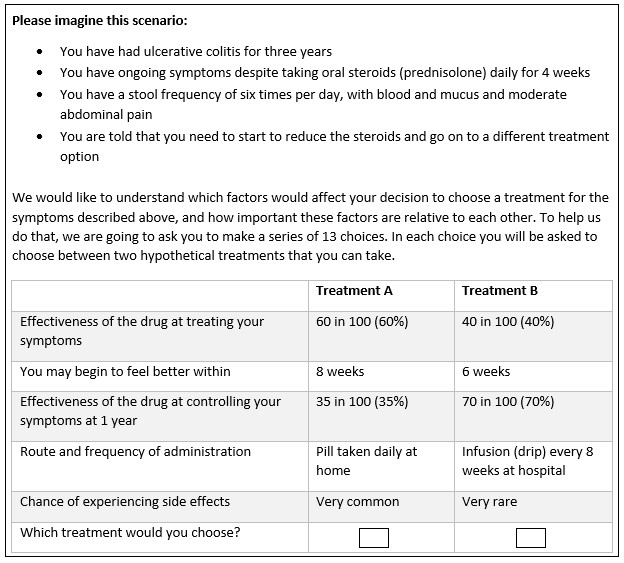
#### **Table 2:** Patient characteristics and ulcerative colitis medication

|  |  |  |
| --- | --- | --- |
|  | No. | % |
| **Sex** |  |  |
| Female | 60 | 52% |
| Male | 55 | 48% |
| **Age by category** | |  |
| 18 to 29 | 17 | 15% |
| 30 to 39 | 32 | 28% |
| 40 to 49 | 24 | 21% |
| 50 to 59 | 18 | 16% |
| 60 to 69 | 12 | 10% |
| 70+ | 12 | 10% |
| *Median age (range)* | 41 (18-84) | |
| **Activity** |  |  |
| Employed | 82 | 71% |
| Retired | 17 | 15% |
| Homemaker | 6 | 5% |
| Unemployed | 5 | 4% |
| Student | 4 | 3% |
| Volunteer work | 1 | 1% |
| **Ethnicity** |  |  |
| White | 109 | 95% |
| Asian | 5 | 4% |
| Mixed | 1 | 1% |
| Black | 0 | 0% |
| **Education** |  |  |
| Primary | 1 | 1% |
| GCSE | 21 | 18% |
| A-Level | 37 | 32% |
| Degree | 53 | 46% |
| Prefer not to say | 3 | 3% |
| **Marital status** |  |  |
| Married/Partner | 89 | 77% |
| Single | 21 | 18% |
| Divorced/Separated | 3 | 3% |
| Widowed | 1 | 1% |
| Prefer not to say | 1 | 1% |
| **Ulcerative colitis severity** | **Median** | **range** |
| Duration with UC, years | 10 | 1 - 51 |
| Quality of life- EQ-5D-5L utility score | 0.77 | -0.25 - 1 |
| IBD-Control-8 score | 13 | 0 - 16 |
| Poor control of disease (IBD-Control-8 score≤12)- n (%) | 51 (44.7%) | |
| Good control of disease (IBD-Control-8 score≥13)- n (%) | 63 (55.3%) | |
| **Medication** | **Current use** | **Ever usea** |
| **n (%)** | **n (%)** |
| Adalimumab | 6 (5.3) | 32 (27.8) |
| Infliximab | 11 (9.7) | 59 (51.3) |
| Golimumab | 0 (0.0) | 4 (3.5) |
| Steroids | 4 (3.5) | 107 (93.0) |
| Tacrolimus | 1 (0.9) | 0 (0.0) |
| Tofacitinib | 3 (2.7) | 8 (7.0) |
| Thiopurines | 7 (6.2) | 81 (70.4) |
| Ustekinumab | 1 (0.9) | 3 (2.6) |
| Vedolizumab | 19 (16.8) | 34 (29.6) |
| Combination treatments | 29 (25.7) | - |
| 5-ASA | 24 (21.2) | - |
| None | 8 (7.1) | 0 (0) |
| a Patients reported the use of multiple treatments, so the % does not add up to 100. | | |

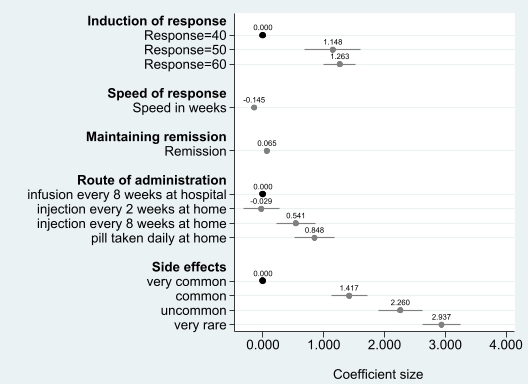
#### **Table 3:** Minimum acceptable benefit

|  |  |
| --- | --- |
| **Attribute level change** | **Percentage improvement of maintenance of remission required to compensate for the attribute level change (95% CI)** |
| **Side effects:** change from very rare to uncommon | 10.35 % (4.92 - 15.78) |
| **Side effects:** change from very rare to common | 23.22% (18.28 - 28.163) |
| **Side effects:** change from very rare to very common | 44.86% (38.08 - 51.64) |
| **Administration:** change from daily pill to injection every 8 weeks | 4.69% (-.71 - 10.09) |
| **Administration:** change from daily pill to injection every 2 weeks | 13.39% (8.50 - 18.28) |
| **Administration:** change from daily pill to infusion every 8 weeks | 12.95% (7.96 - 17.93) |

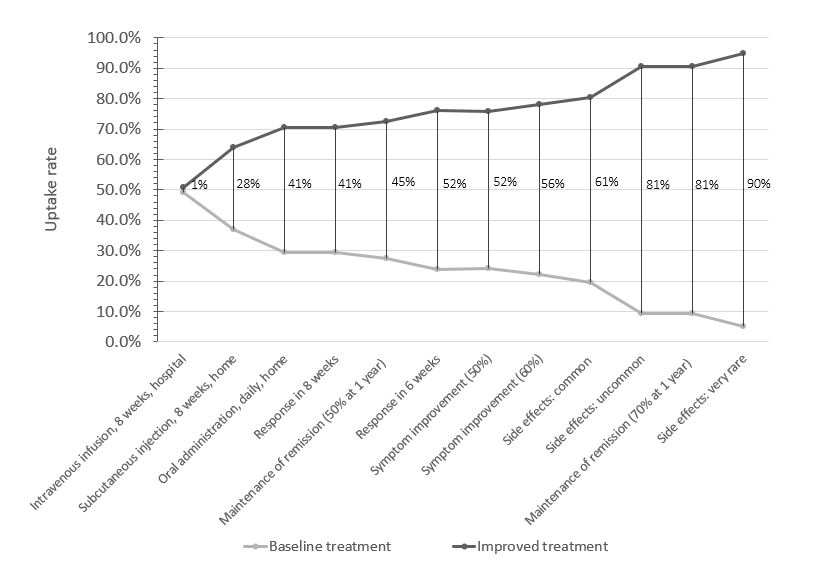
#### **Figure 1:** Example of a discrete-choice question



#### **Figure 2:** Conditional logit model results

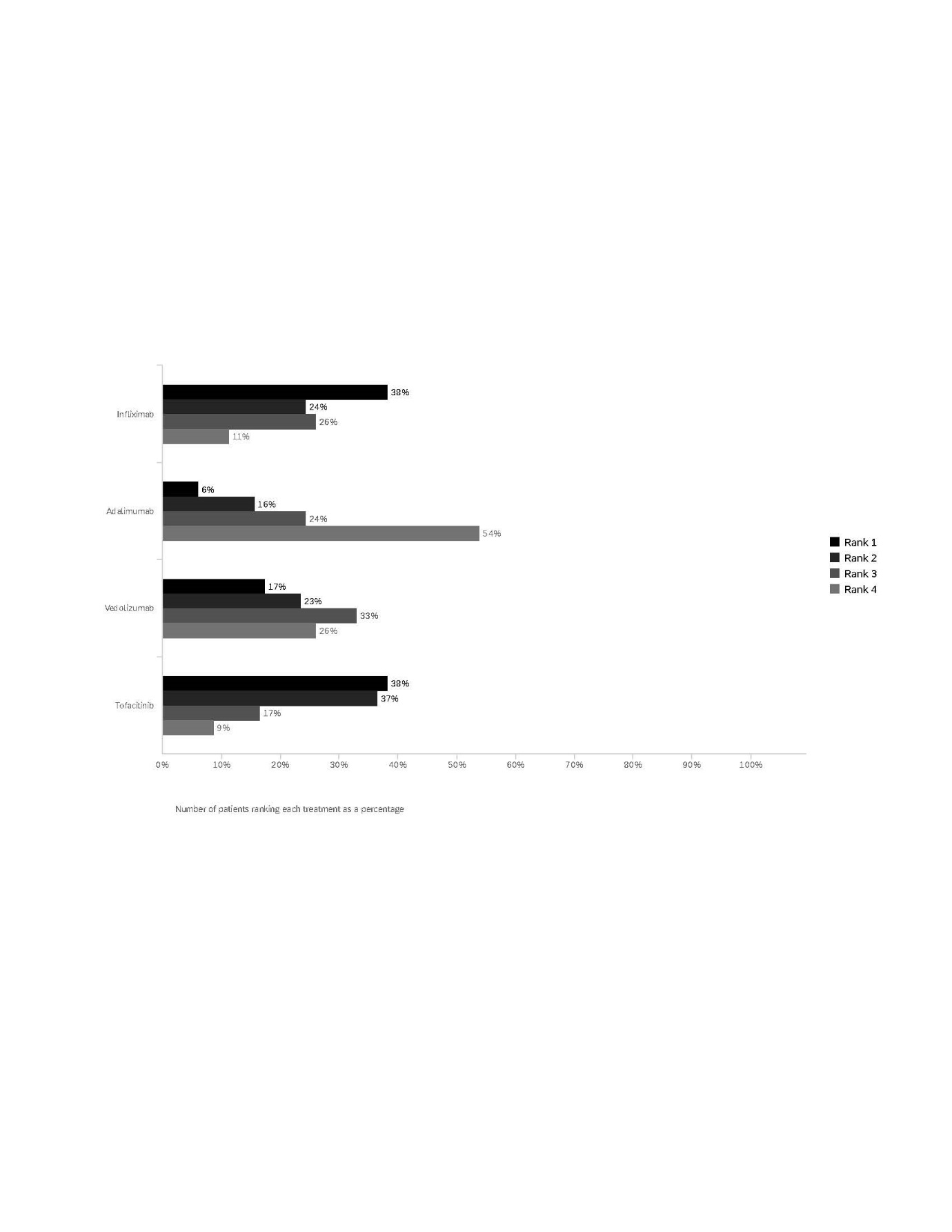


#### **Figure 3:** Overall rank ordering of attribute importance



Note: Each vertical line represents the change in uptake rate of two treatments profiles. The baseline treatment profile was constant where induction of response was 40%, speed of response was 14 weeks, remission rate was 35%, mode of administration was injection every 2 weeks at home, with very common side effects. One attribute level was improved from the base level in each improved treatment profile.

#### **Figure 4:** Biological therapies ranked in order of preference



**Figure 5:** Study flow chart detailing the recruitment of participants.

