Patient preferences in Italy: health state utilities associated with attributes of weekly injection devices for treatment of type 2 diabetes

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Objectives: Several glucagon-like peptide-1 receptor agonists are administered as weekly injections for treatment of type 2 diabetes (T2D). These medications vary in their injection processes, and a recent study in the UK found that these differences had an impact on patient preference and health state utilities. The purpose of this study was to replicate the UK study in Italy to examine preferences of an Italian patient sample, while allowing for comparison between utilities in the UK and Italy.

Materials and methods: Participants with T2D in Italy valued health states in time trade-off interviews. All health states had the same description of T2D, but differed in description of the treatment process. As in the original UK study, the first health state described an oral treatment regimen, while additional health states added a weekly injection. The injection health states differed in three injection-related attributes: requirements for reconstituting the medication, waiting during medication preparation, and needle handling.

Results: Interviews were completed by 238 patients (58.8% male; mean age = 60.2 years; 118 from Milan, 120 from Rome). The oral treatment health state had a mean (SD) utility of 0.90 (0.10). The injection health states had significantly (p < 0.0001) lower utilities, which ranged from 0.87 (requirements for reconstitution, waiting, and handling) to 0.89 (weekly injection with none of these requirements). Differences in health state utility scores suggest that each administration requirement was associated with a disutility (ie, negative utility difference): −0.006 (reconstitution), −0.006 (needle handling), −0.011 (reconstitution, needle handling), and −0.022 (reconstitution, waiting, needle handling).

Conclusion: Disutilities associated with the injection device characteristics were similar to those reported with the UK sample. Results suggest that injection device attributes may be important to some patients with T2D, and it may be useful for clinicians to consider these attributes when choosing medication for patients initiating these weekly treatments.

Keywords: utility, type 2 diabetes, injection process, GLP-1 receptor agonist, time trade-off, weekly injection, Italy, injection device

Introduction

Health state utility assessments can provide insight into patient preferences among treatment options. Choice-based preference elicitation methods including time trade-off (TTO) and standard gamble procedures yield utility values representing the strength of preference for health states.1,2 These utilities may be used to calculate quality-adjusted life years (QALYs) in cost-utility analyses (CUAs) examining and comparing the value of pharmaceutical treatments.3 Although CUAs are often used to guide decisions on health care resource allocation in a wide range of geographical regions, the utilities used...
as inputs in these models are usually estimated in northern Europe, most frequently the United Kingdom.

Italy is one example of a country where CUAs often use utilities derived from preferences of patients or general population respondents in other countries including the UK. However, little is known about the extent to which health state preferences in Italy may differ from those in the UK. Some studies have derived utilities using the EQ-5D in multiple countries including Italy. While these studies allow for comparisons across countries, the preferences and resulting utilities are not directly comparable because the EQ-5D values are based on patients’ ratings of their own unique health, and there could be substantial variation in patient sample characteristics across countries. No studies were located that provide the opportunity for comparing utility values for the same health states in Italy and other countries.

The current study addresses this issue by replicating a UK health state utility valuation in a sample of Italian patients, using the same methods and health states so that results would be directly comparable between countries. This replication was designed to provide insight into whether UK patient preferences may be generalizable to a southern European country such as Italy. The UK study completed in 2015 focused on patient preferences related to treatment process attributes of weekly glucagon-like peptide-1 (GLP-1) receptor agonists, which are injectable medications that are often recommended as part of combination therapy for type 2 diabetes (T2D) when oral medication does not result in sufficient glycemic control.

Despite similarities in efficacy and safety, the weekly GLP-1 receptor agonists differ in terms of injection treatment process attributes. Therefore, this study focused on preferences and utility differences associated with three treatment-related attributes that vary among these treatments: 1) requirements for reconstitution (ie, mixing the medication) prior to injection, 2) waiting time required prior to the injection, and 3) whether patients are required to handle the injection needle. The first two attributes were hypothesized to be important to patients who prioritize ease of administration, while the third attribute is considered to be an important safety issue. The UK study found that these treatment process attributes had a quantifiable impact on patient preference.

In sum, the current study had three goals. First, TTO utility elicitation interviews were conducted to provide insight into patient preferences among weekly GLP-1 receptor agonists injection device attributes in Italy. Second, the interviews were designed to yield utility scores that could be useful in economic modeling of weekly GLP-1 receptor agonists in Italy. Third, by enabling comparison of current results to those of the original UK utility valuation study, the current replication can provide insight into potential geographic differences in treatment preference and utility scores.

Materials and methods
Overview of study design
This vignette-based study elicited patient preferences and estimated health state utilities for injection process attributes of weekly GLP-1 receptor agonists in a sample of patients with T2D in Italy (Milan and Rome) in March and April 2017. This was a replication of a previous utility valuation study conducted in the UK in 2015. Other than the translation of study materials into Italian, health states and utility elicitation methods were the same as those previously described for the UK study. Health state descriptions (often called “health states” or vignettes) drafted for the original UK study were valued by patients in TTO interviews with a 20-year time horizon. These methods were based on those of two previous studies that were conducted to estimate utilities associated with attributes of injectable treatments for T2D. The utilities reported in these previous publications have been used in multiple published CUAs of treatments for T2D.

Utilities were estimated using health state descriptions because the vignette-based approach is well-suited for isolating the impact of specific treatment-related attributes on utility. Although generic preference-based measures such as the EQ-5D are often recommended for estimating utilities, these standardized instruments do not specifically assess the impact of treatment process or treatment convenience. Therefore, they are unlikely to be sensitive to differences in the injection device attributes that were the focus of this study.

Health state development
The health states that were valued in this Italian study were a translated version of health states developed for the previous UK study. For the UK study, the health states were drafted and refined in five steps. First, health states from two previously published studies were adapted for the introductory description of T2D appearing at the beginning of every health state. Second, a literature search was performed to identify attributes of injection devices that were likely to be important to patients and differentiate the weekly GLP-1 receptor agonists. Third, interviews with four clinicians (all with MD degrees and research/clinical experience with T2D) were conducted to finalize the list of injection device attributes and inform the development of...
had type 1 diabetes, latent autoimmune diabetes in adults
informed consent. Participants were not eligible if they
5) be a resident of Italy; and 6) be willing and able to give
understand the assessments as judged by the investigator;
treatment for diabetes, and/or age of diagnosis; 4) be able
of diagnosis, current or previous non-insulin medication
years old; 3) have T2D as indicated by patient report
medical professional; 2) be between the ages of 30 and
targets were used to ensure that no particular demographic
group was over-represented: maximum 60% of either men
or women, maximum 10% unemployed, maximum 50%
retired (the maximum for retired participants was relatively
high because the population of patients with T2D tends to
be older on average). A total of 450 patients were screened
to assess whether they met the inclusion criteria, and 157 of
these did not meet criteria. Of the 293 who were eligible, 241
were scheduled, and 240 attended their interviews. Two of
the 240 participants had difficulty understanding the utility
interview procedures and health states, and were therefore
unable to provide valid data. Thus, a total of 238 (120 Rome,
118 Milan) valid interviews were completed.

Participants
To be eligible for inclusion in the study, participants were
required to 1) be diagnosed with diabetes by a recognized
medical professional; 2) be between the ages of 30 and
75 years old; 3) have T2D as indicated by patient report
diagnosis, current or previous non-insulin medication
treatment for diabetes, and/or age of diagnosis; 4) be able
to understand the assessments as judged by the investigator;
5) be a resident of Italy; and 6) be willing and able to give
informed consent. Participants were not eligible if they
had type 1 diabetes, latent autoimmune diabetes in adults
(LADA), cognitive impairment, hearing difficulty, or severe
pathology that could interfere with their ability to complete
the interview. All participants were required to verify their
diagnosis of T2D in one of two ways. Participants who
were currently receiving medication treatment for T2D were
required to bring proof of this medication to the interviews
(eg, medication packaging or a letter from a doctor). Partici-
pants who were not receiving medication were required
to describe their symptom history, diagnosis process, and
disease management strategies in a way that clearly indicated
that they were honestly reporting their diagnosis.

Participants were recruited in compliance with Italian
privacy law (Legislative Decree no 196 of 30 June 2003
[Codice in materia di protezione dei dati personali, the
“Privacy Code”]). Patients were recruited from a database of
patients who participated in previous studies and expressed
interest in being contacted for future studies. Each patient
was originally contacted and added to the database by a
network of recruiters who work for a market research com-
pany. For the current study, participants were contacted via
telephone or e-mail, depending on their preferred contact
method listed in the database. In addition to strictly adher-
ing to the inclusion criteria described earlier, several recruitment
targets were used to ensure that no particular demographic
group was over-represented: maximum 60% of either men
or women, maximum 10% unemployed, maximum 50%
retired (the maximum for retired participants was relatively
high because the population of patients with T2D tends to
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interview procedures and health states, and were therefore
unable to provide valid data. Thus, a total of 238 (120 Rome,
118 Milan) valid interviews were completed.

Translation
Study materials were translated into Italian for use in this
study. The translation process for study materials other than
the health states (eg, device display page, standardized inter-
view guide, consent form, demographic form) consisted of
the following steps. First, the initial translation from English
to Italian was conducted by a native Italian speaker. Then,
an additional translator who is a native Italian speaker and
was not involved in the forward translation reviewed the
translation and performed editing and proofreading of the
Italian version in conjunction with trained translation project
managers. Finally, Italian members of the study team and data
collection staff reviewed all Italian documents and recommended final edits to ensure that the materials would be clear and comprehensible to Italian-speaking study participants.

Translation of the health states followed the same methodology, but with an additional back-translation step. This back translation was performed by a native English speaker fluent in Italian, and it was reviewed by an additional translator who is a native Italian speaker (but who was not involved in the forward translation) in conjunction with trained translation project managers.

**Utility interview procedures and scoring**

Health state utilities were elicited in a valuation study conducted in March and April 2017 in two locations in Italy: Milan and Rome. All participants provided written informed consent. The study protocol was approved by an independent institutional review board (Ethical & Independent Review Services; Study Number 17011-01). Each one-on-one interview was conducted in a quiet private room by one of five Italian-speaking interviewers. The TTO utility assessment followed a semi-structured interview script in order to standardize the utility assessment procedures. The principal investigator (who also directed the original UK study on which this study was based) trained the interviewers and observed each interviewer multiple times to ensure that procedures were followed consistently.

Utility interview procedures were the same as those previously described for the UK study. In each interview, the basic health state describing T2D treated with oral medication (health state A) was presented first. Then, the device display page was presented to introduce the injection device attributes. After participants indicated that they understood these injection device attributes, the six health states describing treatment with injectable medication (B to G) were presented in random order. Then, participants ranked the health states from most preferable to least preferable.

After the introductory ranking task was completed, participants valued the health states in a TTO task with a 20-year time horizon and 1-year (ie, 5%) trading increments. Following commonly used TTO procedures, participants were asked which of two options they would prefer: 20 years in the health state being rated or a shorter amount of time in full health. The amount of time in full health alternated between longer and shorter time periods (ie, 20 years, 0 years [ie, dead], 19 years, 1 year, 18 years, 2 years, 17 years, 3 years...). The utility value of each health state was calculated as $u = x/y$ based on the point of indifference between $y$ years in the health state being evaluated (ie, 20 years) and $x$ years in full health (followed by dead).

**EQ-5D-3L**

The EQ-5D-3L was administered to characterize the sample in terms of overall health status. The EQ-5D is a self-administered, generic, preference-weighted measure of health status. The first section consists of five dimensions to assess health-related quality of life (mobility, self-care, usual activities, pain/discomfort, and anxiety/depression). These five dimensions are scored based on preference weightings to obtain an “index score” that is often used as a utility in economic modeling. The second section consists of a visual analog scale on which respondents rate their current health, with anchors of 0 (worst imaginable health state) and 100 (best imaginable health state). The EQ-5D-3L index score was computed using published Italian tariffs.

**Statistical analysis procedures**

Statistical analyses were completed using SAS version 9.4 (SAS Institute, Cary, NC, USA). Descriptive statistics were calculated for continuous variables (means and standard deviations) and categorical variables (frequencies and percentages). Demographic subgroups were compared with chi-square analyses (for categorical variables) and independent $t$-tests (for continuous variables). Pairwise comparisons between health state utilities were performed using Student’s $t$-tests. For each injection process attribute, disutilities (ie, decreases or differences in utility) were calculated by subtracting mean utility scores. Independent $t$-tests were also used to compare utilities between demographic subgroups (eg, gender, age), as well as between the current Italian sample and the previous UK sample.

**Results**

**Sample description**

Interviews were completed by a total of 238 participants with T2D, including 120 in Rome and 118 in Milan (Table 1). At the time of their interviews, the majority of respondents were treated with oral medication for T2D ($n = 170, 71.4\%$). Fewer patients were receiving injectable medication ($n = 18, 7.6\%$), combined oral/injectable treatment regimens ($n = 40, 16.8\%$), or no medication ($n = 10, 4.2\%$). Of the 238 participants, 228 brought proof of current medication to the interviews. The other 10 participants who were not taking medication described their symptoms, diagnosis, and treatment at a level of detail indicating that they were accurately reporting their diagnoses. The most commonly reported comorbid health conditions were hypertension (46\%); diabetic retinopathy (11\%); anxiety (13\%); heart attack or heart disease (11\%); and arthritis (10\%).
Independent t-tests revealed that there were no statistically significant differences in utility or disutility between men and women; between older and younger respondents (categorized by median split); or among treatment regimen groups (analysis of variance [ANOVA] comparing no medication, oral only, and injectable with or without oral medication).

The EQ-5D-3L mean index score was 0.91 (SD = 0.12) when computed with published Italian EQ-5D tariffs. This mean is slightly higher than most scores published previously for patients with T2D without complications, likely because the Italian tariffs yield higher index scores than the more commonly used UK tariffs.

### Health state utilities

When ranking the seven health states, respondents generally placed them in a logical order. Health states describing more inconveniences associated with the injection device were typically ranked as less preferable. Rankings ranged from 1 (most preferable health state) to 7 (least preferable health state). The oral-only health state (A) was always ranked as most preferable (mean ranking of 1.00), followed by G (2.07), F (3.57), E (3.65), D (5.18), C (5.60), and B (6.94).

The utility scores followed a similar pattern to the ranking, with injection-related inconveniences associated with lower utility (Table 2). The mean utility for health state A (representing diabetes with oral treatment and no injection)
was 0.900. The addition of a weekly injection in health state G reduced utility to 0.890. The addition of treatment process inconveniences associated with the weekly injection decreased utility further. The lowest utility value was for health state B, which described all three injection-related inconveniences: reconstitution, waiting, and needle handling (0.868). All participants rated every health state as better than dead, and therefore, there were no negative utility values.

### Disutilities and comparisons among health states

Disutility scores (ie, decreases in utility) are provided relative to health state A describing oral-only treatment and relative to health state G describing oral plus weekly injectable treatment (Table 2). The disutility of each injection process attribute (ie, reconstitution, waiting, needle handling) was calculated by subtracting health state G from health states B to F. For example, the disutility of the needle handling requirement was calculated as the difference between G and F (−0.006), two health states that were identical except for the addition of needle handling. Each individual injection attribute had a relatively small disutility (eg, disutility = −0.006). However, larger disutilities were found for health states that included multiple injection-related inconveniences, such as health state B representing the combination of reconstitution, waiting, and needle handling requirements (disutility = −0.022).

Student’s t-tests revealed that the utility of the oral-only health state (health state A) was statistically significantly greater than the utility of every other health state (all \( p < 0.0001 \)). In addition, the injection health state without any of the three inconveniences (health state G) was compared to each of the other injection health states (B to F). The utility of G was found to be significantly greater than the utility of all health states that included reconstitution, waiting, and/or mixing (all \( p < 0.0001 \)).

### Geographical comparisons

Independent t-tests found no significant differences in mean utility or disutility between the UK sample in the original study and the Italian sample in this replication study (Tables 3 and 4). Utilities and disutilities were also compared between the two cities in Italy, and the relationships among health states were almost identical in the two cities. There were no significant differences between the two cities in disutility associated with injection-device attributes (ie, the difference score when subtracting the utility of health state G from the utility of the other injection health states), and disutility in disutility between the two cities was 0.00 for all health states (ie, identical disutilities in Milan and Rome to two decimal places). However, mean utility values for each health state were greater in Milan than in Rome (all \( p < 0.01 \)). The mean utility in Milan was greater than Rome by a magnitude of 0.04 for health state A and a magnitude of 0.05 for all other health states. For example, utilities of health states B, D, and G were 0.84, 0.85, and 0.87 in Rome, compared with 0.89, 0.91, and 0.92, respectively, in Milan. Thus, the pattern of mean scores was the same in the two cities, indicating similar preference among injection device characteristics, but the lower utility scores in Rome suggest greater willingness to trade time regardless of health state.

### Discussion

In this replication study in Italy, patient preferences for attributes of weekly GLP-1 receptor agonist injections were consistent with those in the original UK study. As in the UK study, each individual injection inconvenience (reconstitution, waiting, needle handling) had a relatively small impact on utility (−0.006 to −0.011), but health states representing

<table>
<thead>
<tr>
<th>Health statea utilities</th>
<th>Italy (n = 238)</th>
<th>UK (n = 209)</th>
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<th>t-test</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>A. Oral treatment only</strong></td>
<td>0.900 (0.097)</td>
<td>0.888 (0.120)</td>
<td>0.012 (0.108)</td>
<td>1.132</td>
</tr>
<tr>
<td><strong>Health states with oral and injectable treatment</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>B. Reconstitution, waiting, needle handling</td>
<td>0.868 (0.109)</td>
<td>0.858 (0.165)</td>
<td>0.010 (0.138)</td>
<td>0.779</td>
</tr>
<tr>
<td>C. Reconstitution, waiting</td>
<td>0.872 (0.109)</td>
<td>0.863 (0.161)</td>
<td>0.009 (0.136)</td>
<td>0.681</td>
</tr>
<tr>
<td>D. Reconstitution, needle handling</td>
<td>0.879 (0.102)</td>
<td>0.868 (0.159)</td>
<td>0.011 (0.132)</td>
<td>0.885</td>
</tr>
<tr>
<td>E. Reconstitution</td>
<td>0.884 (0.100)</td>
<td>0.874 (0.157)</td>
<td>0.010 (0.130)</td>
<td>0.804</td>
</tr>
<tr>
<td>F. Needle handling</td>
<td>0.884 (0.101)</td>
<td>0.874 (0.156)</td>
<td>0.010 (0.130)</td>
<td>0.781</td>
</tr>
<tr>
<td>G. No inconveniences</td>
<td>0.890 (0.100)</td>
<td>0.878 (0.156)</td>
<td>0.012 (0.129)</td>
<td>0.989</td>
</tr>
</tbody>
</table>

Notes: UK data adapted with permission from Matza LS, Boye KS, Stewart KD, Davies EW, Paczkowski R. Health state utilities associated with attributes of weekly injection devices for treatment of type 2 diabetes. BMC Health Serv Res. 2017;17:774. Health states B to G include health state A, plus treatment with a weekly injection. Independent t-tests were used.

Abbreviation: TTO, time trade-off.
simultaneous multiple inconveniences were associated with greater disutility (up to −0.022 for all three inconveniences combined in health state B). The mean disutility scores for the injection device attributes estimated in this Italian sample were nearly identical to those in the UK sample. These disutilities are similar in magnitude to utility differences reported for other treatment process variables, such as route of administration and variations in the injection process, which typically range from 0.01 to 0.05.26,35,36 These small differences in utility can have an impact on the outcome of a cost-utility model, particularly when modeling large groups of patients over a long time period. Thus, the current utility values may be useful for representing patient preferences in economic models of weekly GLP-1 receptor agonists for T2D in Italy. In addition, the utilities could potentially be applicable in models of other medications that are administered with similar injection devices.

As with results of the UK study,11 researchers working on cost-utility models comparing weekly GLP-1 receptor agonists should focus primarily on differences among health states B through G (the column titled “Disutilities: Difference from Health State G” in Table 2). Because these six health states are identical except for differences in the three injection attributes, any differences in utility can be attributed specifically to these three attributes (ie, reconstitution, waiting, needle handling). For example, a model could compare cost-effectiveness in Italy of a medication requiring needle handling to a medication without this requirement. In this situation, the utility difference between health states F and G would be relevant, and the corresponding disutility of −0.006 may be applied to the treatment arm representing the injection pen with the needle handling requirement.

Despite the similarity in mean disutility scores between Italy and the UK, an interesting geographical difference within Italy did emerge. The key results of the study are the disutilities associated with each of the three injection-related attributes, and these disutility values were nearly identical in the Milan and Rome groups (and also very similar to results from the original UK sample). However, while the pattern of differences among health states was the same across the two Italian cities, the mean utility of each individual health state was greater in Milan than in Rome by a consistent magnitude for all health states (ie, difference of 0.05 for all injection health states). These results indicate that while preferences among injection-related attributes were consistent across the cities, there may be geographical differences in willingness to trade time in a TTO task. On average, respondents in Rome were willing to trade more time than respondents in Milan, resulting in lower utilities in Rome. While this difference in trading style had no apparent impact on the disutilities of the treatment-related attributes, which were consistent across the two cities, it may be useful for future research to examine demographic or geographic differences in willingness to trade time.

Table 4 t-tests comparing TTO disutility scores between countries (Italy vs UK) (N = 447)

<table>
<thead>
<tr>
<th>Health state disutilities</th>
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<tbody>
<tr>
<td>Disutilities relative to health state A</td>
<td>Mean (SD)</td>
<td>Mean (SD)</td>
<td>Mean (SD)</td>
<td>t-statistic</td>
</tr>
<tr>
<td>A vs B: Oral treatment only vs reconstitution, waiting, needle handling</td>
<td>−0.032 (0.059)</td>
<td>−0.030 (0.073)</td>
<td>−0.001 (0.066)</td>
<td>−0.212</td>
</tr>
<tr>
<td>A vs C: Oral treatment only vs reconstitution, waiting</td>
<td>−0.027 (0.058)</td>
<td>−0.025 (0.066)</td>
<td>−0.003 (0.062)</td>
<td>−0.474</td>
</tr>
<tr>
<td>A vs D: Oral treatment only vs reconstitution, needle handling</td>
<td>−0.020 (0.040)</td>
<td>−0.020 (0.063)</td>
<td>−0.000 (0.052)</td>
<td>−0.082</td>
</tr>
<tr>
<td>A vs E: Oral treatment only vs reconstitution</td>
<td>−0.015 (0.032)</td>
<td>−0.014 (0.058)</td>
<td>−0.002 (0.046)</td>
<td>−0.343</td>
</tr>
<tr>
<td>A vs F: Oral treatment only vs needle handling</td>
<td>−0.016 (0.033)</td>
<td>−0.014 (0.058)</td>
<td>−0.002 (0.047)</td>
<td>−0.411</td>
</tr>
<tr>
<td>A vs G: Oral treatment only vs no inconveniences</td>
<td>−0.009 (0.025)</td>
<td>−0.010 (0.056)</td>
<td>0.001 (0.042)</td>
<td>0.167</td>
</tr>
<tr>
<td>Disutilities relative to health state G</td>
<td>Mean (SD)</td>
<td>Mean (SD)</td>
<td>Mean (SD)</td>
<td>t-statistic</td>
</tr>
<tr>
<td>G vs B: No inconveniences vs reconstitution, waiting, needle handling</td>
<td>−0.022 (0.052)</td>
<td>−0.020 (0.042)</td>
<td>−0.002 (0.048)</td>
<td>−0.455</td>
</tr>
<tr>
<td>G vs C: No inconveniences vs reconstitution, waiting</td>
<td>−0.018 (0.052)</td>
<td>−0.014 (0.032)</td>
<td>−0.003 (0.044)</td>
<td>−0.865</td>
</tr>
<tr>
<td>G vs D: No inconveniences vs reconstitution, needle handling</td>
<td>−0.011 (0.030)</td>
<td>−0.010 (0.027)</td>
<td>−0.001 (0.029)</td>
<td>−0.412</td>
</tr>
<tr>
<td>G vs E: No inconveniences vs reconstitution</td>
<td>−0.006 (0.021)</td>
<td>−0.004 (0.016)</td>
<td>−0.002 (0.019)</td>
<td>−1.281</td>
</tr>
<tr>
<td>G vs F: No inconveniences vs needle handling</td>
<td>−0.006 (0.021)</td>
<td>−0.004 (0.015)</td>
<td>−0.003 (0.018)</td>
<td>−1.512</td>
</tr>
</tbody>
</table>

Notes: UK data adapted with permission from Maza LS, Boye KS, Stewart KD, Davies EW, Paczkowski R. Health state utilities associated with attributes of weekly injection devices for treatment of type 2 diabetes. BMC Health Serv Res. 2017;17:774;11 Health state G includes health state A, plus treatment with a weekly injection; ‘disutilities computed for “A. Oral only” as x – A; and for “G. No inconveniences (Dulaglutide)” as x – G. Independent t-tests were used.

Abbreviation: TTO, time trade-off.
Limitations
An important limitation of this research is that disutilities of the injection process attributes are based on patients’ interpretations of hypothetical health state vignettes, rather than personal experience with each injection device. Vignette-based utilities should be interpreted and used with caution because this methodology could be subject to biases. For example, preferences expressed during the TTO task could be influenced by a focusing effect, causing respondents to attend closely to small differences among health states and potentially exaggerating differences in utility.

Furthermore, comparability of utilities estimated via vignette-based methods to utilities derived from more standardized approaches such as the EQ-5D is uncertain. Some health technology assessment guidelines, particularly the guide issued by National Institute for Health and Care Excellence, indicate a preference for generic preference-based utility measures such as the EQ-5D to maximize “consistency across appraisals.” This guide adds that utilities estimated using other methods may be useful when the EQ-5D is not “appropriate.” Estimation of utilities associated with treatment process is one situation when the EQ-5D is not likely to be appropriate because generic instruments such as the EQ-5D may not be sensitive to specific treatment-related attributes. The vignette approach is a useful alternative for assessment of treatment process utilities because vignettes can be designed to differ based on specific treatment process attributes. This is likely why the great majority of studies on “process utilities” use vignette-based methods despite the challenges in comparing results to generic measures such as the EQ-5D.

Conclusion
Despite limitations, findings add to the growing body of literature suggesting that the treatment process has a measurable impact on patient preference and health state utility, particularly among patients with T2D. The current study builds on this previous research by quantifying patient preferences for some key injection-related attributes of weekly GLP-1 receptor agonists. In addition to providing further insight into patient preferences, the resulting utility scores may be useful in economic modeling of these treatments. The similarity between the results from this Italian replication and the original UK study supports confidence in these findings, indicating that patient preferences were consistent despite geographical, cultural, and linguistic differences. In a larger sense, this consistency between the two geographic locations provides initial support for using UK utility values as inputs in cost-utility models in Italy. Additional utility elicitation research in Italy and other locations is needed to further examine generalizability of patient preferences and utilities across geographic regions.

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Disclosure
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