Educating Patients on Unnecessary Antibiotics: Personalizing potential harm aids patient understanding

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Abstract

Introduction: Antibiotic resistance is a public health emergency fueled by inappropriate antibiotic use. Public education campaigns often focus on global antibiotic resistance or societal harm of antibiotic misuse. There has been little research into what messages have the greatest impact on patient preferences for non-indicated antibiotics in ambulatory clinics.

Methods: We administered a survey at a primary care clinic in Baltimore, MD. 250 participants rated 18 statements about potential harm from antibiotics on how each statement changed their likelihood to request antibiotics for an upper respiratory tract infection (URI). Statements focused on potential harm either to the individual, to contacts of the individual, to society, and related or not to antibiotic resistance. Initial and final likelihood of requesting antibiotics was measured, and the impact of the statements in each category were compared using general linear models and Wilcoxon rank sum or Kruskal-Wallis tests.

Results: All statements decreased patient likelihood to request antibiotics. Statements about harm to the individual or contacts of the individual decreased participant likelihood to request antibiotics significantly more than statements about societal harm of antibiotic misuse. Statements not discussing antibiotic resistance decreased participant likelihood of requesting antibiotics significantly more than statements discussing antibiotic resistance. Overall likelihood to request antibiotics decreased after the survey by 2.2 points on an 11-point Likert scale (p<0.001).

Conclusion: When dissuading patients from requesting non-indicated antibiotics, providers and public health campaigns should focus on potential harm of non-indicated antibiotics to the individual rather than societal harm or antibiotic resistance.
Keywords
Antibiotic Resistance; Antibiotic Stewardship; Antibiotic Prescription; Ambulatory care

Introduction:
While progress has been made in antibiotic stewardship, antibiotic resistance is increasing at an alarming rate. Antibiotic-resistant organisms cause more than 2.8 million infections and 35,000 deaths per year in the United States [1]. In 2015, approximately 47 million antibiotic courses were prescribed for infections not needing antibiotics [1].

Many interventions aimed at improving antibiotic use in ambulatory settings have focused on clinician behavior. However, clinicians cite perceived patient desire for antibiotics, impact of patient satisfaction scores, time demands, or lack of patient understanding about negative impacts of harmful antibiotic use as several reasons they may prescribe antibiotics for conditions where they are not indicated, such as for upper respiratory infections (URIs) [2] [3]. Studies have shown that when physicians perceive that patients expect antibiotics, the physician is ten times more likely to prescribe an antibiotic [4] [2].

To decrease unnecessary antibiotic prescriptions, clinicians may benefit from an improved understanding of how to communicate with patients about antibiotics. Patients may not appreciate the negative impact of taking non-indicated antibiotics [3], and it is unclear what information clinicians could provide to patients that has the greatest impact on patient desire for antibiotics. There is limited data on the effectiveness of specific language meant to dissuade patients’ desire and requests for antibiotics for non-indicated conditions. While public health campaigns have often focused on global antibiotic resistance, little research has directly compared this approach with more personal messaging [5] [6].

The purpose of this study was to determine which statements most deter requests for antibiotics for non-indicated conditions by effectively communicating the risks associated with antibiotics. We hypothesized that statements about potential harm to the individual patient or contacts of the individual patient from unnecessary antibiotics would decrease participant desire for antibiotics more than statements about societal harm from non-indicated antibiotics.

Methods:
Survey Instrument:

We developed a survey with 18 statements in 4 categories. Participants rated each statement based on the impact on patient likelihood of requesting antibiotics for a URI. Each statement described a potential harm caused by antibiotics. Eight statements described potential harm to the individual taking the antibiotics, four statements described potential harm to people close to the individual taking the antibiotics, and six statements described potential harm to society. In addition, eight of the statements discussed harm caused by antibiotic resistance while ten of the statements did not discuss antibiotic resistance. To increase readability, statements were edited to utilize short sentences, words with few syllables, and clear
sentence structure. The average Flesch-Kincaid readability score was grade 10.6. The survey was piloted with ten adults, five female and five male, with ages ranging from 19-58 and education level ranging from high school completion to advanced degree completion. Statements were refined for clarity and were narrowed from 24 to 18 in number based on pilot feedback, with the deletion of questions that were reported to be the most confusing or repetitive (See Appendix).

Participation was requested by visual signage. Surveys were delivered in random question order either on paper (N=40) or an electronic tablet using the Qualtrics survey platform (N=210), based on participant preference. Participants read each statement and then rated on a 11-point Likert scale how that statement changed their likelihood to request antibiotics for a URI. A “0” meant the statement made the patient much less likely to request antibiotics and “10” meant the statement made them much more likely to request antibiotics. We utilized an 11 point Likert scale to allow for a neutral choice (5), and since we anticipated most responses would fall on the “less likely” side of neutral, 5 options on either side of neutral provided more distinction between choices than 3 or 4 options. We wished to keep a 0-10 scale with neutral as 5 as this is the way people are used to seeing surveys. However to report the data we transposed the neutral “5” value to a Delta Mean of “0,” the prior value of “0” to a delta mean of “−5,” and the prior score of “10” to “5.” Delta means more clearly display that the most effective statements had a large negative delta mean, while less effective statements had a less negative delta mean. Before and after reading the 18 statements, participants were asked about their overall likelihood of requesting antibiotics for a URI.

Population:
Our survey was available to adults in the waiting room of an outpatient internal medicine and obstetrics and gynecology practice in Baltimore, MD on weekdays between July 20, 2018 to July 27, 2018. Participants were not required to be patients of the clinic (for example, they could have been a family member waiting with a clinic patient). Participants received a $10 gift card upon completion of the survey.

Statistical Analysis
Many of the survey response data were not normally distributed. Therefore, results were summarized using both non-parametric (medians, interquartile ranges [IQRs]) and parametric (means, standard deviations [SDs]) statistics. Group comparisons were made using Wilcoxon rank sum or Kruskal-Wallis tests. General linear models regression was used to estimate the least squares means of the change in the likelihood of requesting antibiotics following the 18 statements. Statistical analysis was performed using SAS version 9.4 (SAS Institute, Inc., Cary, NC). All tests were two-sided, and significance was set at p<0.05.

Results:
Of 1150 total adult patients in clinic and an unknown number of family members or others accompanying these patients during the 6-day course of the survey administration, 250
adults took the survey. Researchers were not allowed to approach any patients, and the 250 participants approached only after seeing signage. These 250 people were out of an unknown number of the 1,150 patients in clinic who actually saw and read the signage. Most participants were women (N= 184, 73.6%), most had completed at least some college or trade school (N=140, 56%), and most were African American (N=152, 61%) (Table 1).

The initial likelihood of requesting antibiotics for a URI like illness was a raw mean score of 5.3 (Slightly more likely to request antibiotics than not, as “5” meant “I may or may not request antibiotics”). After participants completed the survey, this likelihood decreased to a raw mean of 3.1 (“3 meant “I probably will not request antibiotics”, p<0.001) for a mean delta of −1.9. All statements reduced participant likelihood of requesting antibiotics (Table 2). The most impactful statement was “Taking antibiotics can hurt your body’s natural defenses. This makes it easier for you to get another infection” with a change from neutral (5) in mean likelihood of requesting antibiotics (mean delta) of −2.56 while the least effective statement was “Antibiotic resistance costs the U.S. between $20-35 billion each year” with a delta mean of −1.35.

Statements about both individual harm and harm to contacts of an individual had a significantly greater reduction in likelihood of requesting antibiotics (delta mean −2.30 and −2.18 respectively) than statements about harm to society (delta mean −1.80; p<0.001 for both individual harm and harm to contacts of an individual compared with society) (Table 3). Statements about individual harm (delta mean −2.30) and harm to others close to an individual (delta mean −2.18) did not significantly differ in their impact on likelihood of requesting antibiotics (p=0.11). Statements discussing antibiotic resistance (delta mean −1.91) led to a smaller reduction in likelihood to request antibiotics than statements not discussing antibiotic resistance (delta mean −2.26) (p<0.001). This difference was seen in each category of statement; that is, within the individual harm category, statements discussing antibiotic resistance were less effective than statements not discussing antibiotic resistance.

Each demographic group rated individual harm statements as most effective, followed by statements about harm to others and then harm to society (Table 3). Within each category, statements had slightly different rankings, such as women rating statements involving the harm of antibiotic misuse on a pregnant woman’s child as more impactful than men did. For men, the most impactful statement was “Taking antibiotics can hurt your body’s natural defenses. This makes it easier for you to get another infection,” (delta mean −2.30) and the second most impactful was “By changing your normal gut bacteria, antibiotics can cause allergies, asthma, and stomach problems” (delta mean −2.16).

**Discussion:**

The results suggest that providers should focus on individual harm or harm to others close to an individual, and avoid discussing societal harm. Further, providers may wish to avoid focusing on antibiotic resistance, as discussing antibiotic resistance did not decrease participant desire for antibiotics as much as other statements. Patients likely care the most about harm that they can both understand and apply to their own life.
All statements decreased participant likelihood of requesting antibiotics, and participants were less likely to request antibiotics after completing the questionnaire. Clinicians are more likely to prescribe a non-indicated antibiotic when they perceive that patients expect one [2] [4]. Clinician communication with patients about antibiotics can increase patient awareness of antibiotic resistance and understanding of appropriate indications for an antibiotic [7]. Improving patient expectations around antibiotic use may decrease patient requests for antibiotics. Our study provides evidence that using specific statements may impact patient desire for antibiotics and provides guidance as to which of these statements are the most impactful.

Public education campaigns have frequently focused on the spread of antibiotic resistance [5] [6]. Our data suggest that this approach may not be the most effective messaging to convince an individual why they should not request non-indicated antibiotics. Antibiotic resistance is a complex topic that is poorly understood by the public [8]. For example, even though 92% of the American public agrees that inappropriate antibiotic use contributes to antibiotic resistance, only 30% believes that antibiotic resistance is a problem [3]. In administering our study, several study participants asked the study team member questions after the survey about whether antibiotic resistance was a positive or negative characteristic. Antibiotic resistance may be a complex topic to explain and to comprehend, so statements focusing on antibiotic resistance may not be as effective as statements focusing on other harms in talking to patients.

Statements focusing on harm to the individual had the greatest impact on patient likelihood to request antibiotics. It is possible that participants did not see how a global problem impacted them personally. While no research has directly measured patient consideration of individual harm from antibiotics in requesting antibiotics, recent work has shown that even when patients understand that antibiotics only treat bacteria and not viruses, they still believe that antibiotics are so low-risk for causing side effects that they may believe that the antibiotics are more likely to help than hurt symptoms of a URI [8]. In assessing evidence to this effect, recent public health campaigns have shifted towards personalizing the harm of antibiotic resistance such as by discussing possible harms such as *Clostridioides difficile*, allergic reactions, and antibiotic resistant infections [1]. Our study suggests that this type of information is impactful when provided in a series of short statements. Making antibiotic misuse personal may be the most effective strategy in convincing patients why they should not request non-indicated antibiotics [8].

In considering how best to dissuade a patient from non-indicated antibiotics, clinicians should keep in mind that different types of statements may have more of an impact on members of some demographic groups than others. For example, statements about the impact of pregnant women taking antibiotics on the development of babies had a greater impact on women than men. Level of educational attainment also impacted statement impact: those with a lower level of education were more likely to both initially desire antibiotics and be dissuaded by the statements than other groups. Providing education about potential harm may be most effective for those with lower levels of education.
Our quantitative examination into antibiotic resistance communication and motivations for patients to want antibiotics is novel. The relatively large study population included a high proportion of African American patients which matched the local community’s demographics and increases study applicability to similar populations. A larger multi-site study would increase generalizability of results.

The study had several limitations. Generalizability is limited as it was conducted at a single site. Additionally, as not every potential patient could be included in the study, the participants were not randomly selected, and participants may have been motivated to varying degrees by the $10 gift card provided, there was the possibility of response bias. Researchers were not allowed to approach any potential participants about the study, and had to wait for patients to approach after seeing signage. The inability to offer the survey to every patient in clinic, or to determine how many of the 1,150 patients in clinic during survey administration saw and read the advertising signage excludes the denominator necessary to calculate a response rate. The survey’s reading level was relatively high, driven by the words “antibiotic” and “resistance.” While this could have impacted participant understanding of the statements, distributing this survey without including the words “antibiotic” and “resistance” would have not been reflective of public health educational campaigns to decrease antibiotic resistance [1], [9]. This study should be validated by administration of a similar survey at multiple sites, and could be offered or administered to every patient at participating sites to improve study validity.

**Conclusions:**

Patient expectation of an antibiotic increases physician prescription by as much as ten times [4], and while public education campaigns have demonstrated an ability to increase patient knowledge around appropriate antibiotic usage [5] [7], little research exists addressing how best to directly dissuade a patient who is requesting non-indicated antibiotics from their request. Prior public health campaigns addressing patients have focused on antibiotic resistance and large-scale societal harm. Our research supports the approach suggested by the CDC and other stakeholder organizations to shift outpatient clinic and overall discussion towards a more individual-oriented conversation focused on harm to the individual [1]. Our research provides a list of clear statements that clinicians can use in talking with patients about antibiotics. Clinicians and public health campaigns should focus on individual side-effects or harm to others close to the patient when dissuading patients from non-indicated antibiotics, and should not emphasize societal impacts or antibiotic resistance. Whenever possible, clinicians should use statements that are simple, understandable, and directly related to the patient.

**Supplementary Material**

Refer to Web version on PubMed Central for supplementary material.
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Main point:
Prescriber statements about personal harm to patients from non-indicated antibiotic use decreased participant desire for an antibiotic prescription to treat an upper respiratory tract infection more than statements about antibiotic resistance or possible societal harm of antibiotics.
Table 1:
Characteristics of 250 adult participants in a survey about harms of antibiotic use

| Characteristic               | Count (%)   |
|------------------------------|-------------|
| **Sex**                      |             |
| Male                         | 64 (25.6%)  |
| Female                       | 184 (73.6%) |
| Other                        | 2 (0.8%)    |
| **Age**                      |             |
| 18-24                        | 18 (7.2%)   |
| 25-34                        | 70 (28.0%)  |
| 35-44                        | 43 (17.2%)  |
| 45-54                        | 40 (16.0%)  |
| 55-64                        | 40 (16.0%)  |
| 65+                          | 39 (15.6%)  |
| **Education**                |             |
| High School Graduate or less | 54 (21.6%)  |
| Any college or trade school up to bachelor’s degree | 140 (56.0%) |
| Master’s Degree or Higher    | 56 (22.4%)  |
| **Race/Ethnicity**           |             |
| White (Non-Hispanic/Latino)  | 69 (27.6%)  |
| Black or African American    | 152 (60.8%) |
| All others                   | 29 (11.6%)  |
| **Living Situation**         |             |
| I live with family members or roommates | 190 (76.0%) |
| I live by myself             | 60 (24%)    |
### Table 2:
Change in Likelihood of Requesting Antibiotics for URI-Like Symptoms After Reading Each Statement About Potential Harm of Non-Indicated Antibiotics

| Statement                                                                 | Delta Mean Likelihood (Standard Deviation) | Median (Interquartile range) | Statement Category | Discusses Antibiotic Resistance? |
|---------------------------------------------------------------------------|-------------------------------------------|-------------------------------|--------------------|----------------------------------|
| “Taking antibiotics can hurt your body's natural defenses. This makes it easier for you to get another infection” [1] [9]. | −2.56 (2.14)                              | −3 (−5 to −1)                | Individual         | No                               |
| “Antibiotics can change how babies grow inside pregnant women” [10] [11]. | −2.54 (2.28)                              | −3 (−5 to 0)                 | Others close to you | No                               |
| “By changing your normal gut bacteria, antibiotics can cause allergies, asthma, and stomach problems” [10] [12] [13]. | −2.54 (2.21)                              | −3 (−5 to −1)                | Individual         | No                               |
| “Antibiotics can make you more likely to be obese (fat)” [14].            | −2.51 (2.28)                              | −3 (−5 to 0)                 | Individual         | No                               |
| “Pregnant women who take antibiotics are more likely to have babies with asthma” [11]. | −2.50 (2.33)                              | −3 (−5 to 0)                 | Others close to you | No                               |
| “Antibiotics can cause bad bacteria to overgrow in your gut. This can cause diarrhea and belly pain” [15] [12]. | −2.47 (2.17)                              | −3 (−5 to −1)                | Individual         | No                               |
| “In the future, antibiotic resistance will cause more deaths than cancer and diabetes combined” [6]. | −2.42 (2.23)                              | −3 (−5 to 0)                 | Society            | Yes                              |
| “Resistant bacteria can still be found in your gut four years after taking antibiotics” [16]. | −2.30 (2.25)                              | −3 (−4 to 0)                 | Individual         | Yes                              |
| “A single dose of antibiotics makes the bacteria in your body more resistant to treatment” [17]. | −2.12 (2.26)                              | −2 (−4 to 0)                 | Individual         | Yes                              |
| “Antibiotics kill your normal gut bacteria. This can cause bad bacteria to overgrow” [14]. | −1.99 (2.35)                              | −2 (−4 to 0)                 | Individual         | No                               |
| “Antibiotic resistant bacteria cause over two million illnesses and twenty-three thousand deaths in the U.S. each year” [9]. | −1.96 (2.45)                              | −2 (−4 to 0)                 | Society            | Yes                              |
| “Each antibiotic resistant infection costs up to $30,000 more to treat than other infections” [18] [19] [6]. | −1.93 (2.30)                              | −2 (−4 to 0)                 | Society            | Yes                              |
| “Taking antibiotics can cause you to get a yeast infection” [20].         | −1.90 (2.46)                              | −2 (−4 to 0)                 | Individual         | No                               |
| “Resistant bacteria can spread between people” [1] [9].                   | −1.85 (2.66)                              | −2 (−4 to 0)                 | Others close to you | Yes                              |
| “Antibiotics are the most common cause of Emergency Room visits for drug reactions in children” [9]. | −1.83 (2.33)                              | −2 (−4 to 0)                 | Others close to you | No                               |
| “One in five people who take an antibiotic in a hospital will have a side effect” [1]. | −1.72 (2.25)                              | −2 (−4 to 0)                 | Society            | No                               |
| “In some countries like India, people in hospitals have infections so resistant that antibiotics can't treat them” [21]. | −1.40 (2.60)                              | −1 (−4 to 0)                 | Society            | Yes                              |
| “Antibiotic resistance costs the U.S. between $20-35 billion each year” [18] [9]. | −1.35 (2.28)                              | −1 (−3 to 0)                 | Society            | Yes                              |

For the mean delta displayed, a “−5” meant the statement made the patient much less likely to request antibiotics, “0” meant the statement did not change their likelihood, and “5” would mean the statement made them much more likely to request antibiotics.
Table 3:
Mean Delta of Likelihood to Request Antibiotics Overall and by Category and Participant Characteristics

| Statement Category: | Individual | Close Contact | About Society | About Resistance | Not about Resistance | All Statements |
|---------------------|------------|---------------|---------------|------------------|---------------------|---------------|
| Total Sample (N=250)| −2.30      | −2.18         | −1.80         | −1.91            | −2.26               | −2.10         |
| Sex                 |            |               |               |                  |                     |               |
| Men (N=64)          | −1.75      | −1.49         | −1.35         | −1.43            | −1.77               | −1.62         |
| Women (N=184)       | −2.81      | −2.42         | −2.95         | −2.08            | −2.43               | −2.28         |
| Education           |            |               |               |                  |                     |               |
| HS Graduate or Less (N=54) | −1.90 | −1.63       | −1.15         | −1.33            | −1.72               | −1.60         |
| Any college or trade school up to bachelor's degree (N=140) | −2.30 | −2.33 | −1.86 | −1.95 | −2.33 | −2.16 |
| Master's degree or higher (N=56) | −2.65 | −2.34 | −2.25 | −2.38 | −2.50 | −2.45 |
| Race                |            |               |               |                  |                     |               |
| Black (N=152)       | −1.97      | −1.94         | −1.51         | −1.56            | −2.01               | −1.81         |
| White (N=69)        | −2.69      | −2.41         | −2.13         | −2.32            | −2.53               | −2.44         |
| Other (N=29)        | −3.09      | −2.89         | −2.49         | −2.81            | −2.88               | −2.84         |

A mean delta “−5” meant the statement made the patient much less likely to request antibiotics; “0” meant the statement did not change their likelihood, and “5” meant the statement made them much more likely to request antibiotics. (HS stands for High School).