How to Implement Adherence-Promoting Programs in Clinical Practice? A Discrete Choice Experiment on Physicians' Preferences

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Sabrina Müller 10
Tjalf Ziemssen 10
Curt Diehm
Tobias Duncker
Philipp Hoffmanns
Inga-Marion Thate-Waschke
Markus Schürks 10
Thomas Wilke 10

¹Ingress-Health, Wismar 23966, Germany; ²Zentrum für klinische Neurowissenschaften, Klinik und Poliklinik für Neurologie, Universitätsklinikum Carl Gustav Carus an der Technischen Universität Dresden, Dresden 01307, Germany; ³Private Practice, Ettlingen 76275, Germany; ⁴Institut für Augenheilkunde Halle, Halle 06114, Germany; ⁵Private Practice, Rheinstetten 76287, Germany; ⁶Bayer Vital GmbH, Leverkusen 51368, Germany; ⁷Institut für Pharmakoökonomie und Arzneimittelogistik, Wismar 23966, Germany **Introduction:** The aim of this study was to examine physicians' preferences regarding adherence-promoting programs (APPs), and to investigate which APP characteristics influence the willingness of physicians to implement these in daily practice.

Materials and Methods: A discrete choice experiment was conducted among general practitioners, cardiologists, neurologists and ophthalmologists in Germany. The design considered five attributes with two or three attribute levels each: validation status of the APP; possibility for physicians to receive a certificate; type of intervention; time commitment per patient and quarter of the year to carry out the APP; reimbursement for APP participation, per included patient and quarter of the year.

A multinomial logit model was run to estimate physicians' utility for each attribute and to evaluate the influence of different levels on the probability of choosing a specific APP. The relative importance of the attributes was compared between different predefined subgroups.

Results: In total, 222 physicians were included in the analysis. The most important characteristics of APPs were time commitment to carry out the program (34.8% importance), reimbursement (33.3%), and validation status of the program (23.7%). The remaining attributes (type of intervention: 3.6%; possibility to receive a certificate: 4.7%) were proven to be less important for a physician's decision to participate in an APP. Physicians on average preferred APP alternatives characterized by little time commitment (β =1.456, p<0.001), high reimbursement for work (β =1.392, p<0.001), "positive validation status" (β =0.990, p<0.001), the "possibility to get a certificate" (β =0.197, p<0.001), and the provision of "tools for both physicians and patients" (β =0.150, p<0.001).

Conclusion: For the majority of the physicians participating in this survey, the willingness to implement an APP is determined by the associated time commitment and reimbursement. Considering physicians' preferences regarding different APP features in the promoting process of these programs may enhance physicians' participation and engagement.

Keywords: adherence promoting programs, adherence interventions, compliance programs, discrete choice experiment

Introduction

Patients' non-adherence to medication for chronic diseases is a common phenomenon^{1–3} and may have substantial negative clinical and health-economic impact.^{4,5} Adherence to medication is a complex and multidimensional behavior.⁶ Therefore, direct engagement with patients is crucial to address any "adherence barriers."^{3,7} Nevertheless, in real-world

Correspondence: Sabrina Müller Ingress-Health, Alter Holzhafen 19, Wismar 23966, Germany Email sabrina.mueller@ingress-health.com treatment settings, systematic interventions to measure and improve patients' adherence are rarely used.^{8,9} For example, the Global Asthma Physician Survey (GAPS) estimated that written action plans (37%) and technology (15%) were rarely employed by physicians to aid patients' asthma management. Moreover, only a minority of physicians (10%) used validated patient-reported outcomes questionnaires to monitor asthma control.¹⁰

Even if adherence-promoting programs (APPs) are implemented, the evidence with regard to their success rates is still ambiguous. ¹¹ While some interventions were proven to increase patients' adherence ^{12,13} few were able to also improve patients' clinical outcomes. ¹⁴

For an APP implementation, different options exist. Health care professionals such as physicians and pharmacists might be engaged in adherence-promoting interactions with patients. However, physicians' roles in these programs are crucial, as they are usually the central contact persons for patients in the prescription of a medication, in educating patients on why and how to take medication, and for re-engagement with patients and re-prescription of medication. Therefore, it may be hypothesized that limited success of many APPs is at least partly due to a lack of inclusion of and support by treating physicians. Hence, successful implementation of an APP requires physicians' willingness to fulfill their pivotal roles. ¹³ So far, factors associated with the willingness to engage in APPs have not been studied.

The main aim of this study was to understand physicians' preferences determining their willingness to carry out an APP.

Methods

Setting

A physician survey among quota-sampled general practitioners (GPs), cardiologists, neurologists and ophthalmologists was conducted in Germany. Based on a database of physicians throughout Germany 4,000 physicians were invited to participate in the study via mail and phone. Participating physicians filled in paper-and-pencil or online questionnaires or participated in phone interviews. Since we did not collect patient-related data and did not interact with patients, we did not consult an ethics committee. Physicians received a reimbursement for their participation in this study (90,- \in to 120,- \in per completed questionnaire).

Preference Measurement

Among the different methods used to investigate preferences in health care, discrete-choice experiments (DCEs) have become a widely accepted approach in health care.^{15–19} In a DCE, participants are presented with descriptions of two (or more) complete hypothetical options based on a combination of different attributes. Interviewees are then asked to select their preferred alternative from the presented choices.^{15,19} The underlying assumption of a DCE is that rational individuals will always choose the alternative with the higher level of expected utility. Willingness to engage in APPs may be investigated using a DCE study, as APPs are associated with obvious benefits from a physician's perspective (better adherence, additional income, etc.), but are also associated with potential cost, mainly in terms of time commitment.

Our first step in developing a DCE design suitable to assess physicians' preferences regarding different characteristics of APPs was to conduct a qualitative study. This included a targeted literature review regarding key attributes describing different APP options and four different focus groups discussions, each with 3-4 physicians. With these focus groups consisting of GPs, ophthalmologists, cardiologists and neurologists, we discussed the importance and relevance of attributes identified from the literature as well as the need of considering any additional attribute. Moreover, the questionnaire draft, which was generated after assessment of the results of the focus group discussions, was assessed by an advisory board of four clinical experts (TZ, CD, TD, PH). Finally, five attributes with two or three levels for each attribute were agreed upon and included in the questionnaire for the quantitative study:

- Validation status of the APP
 - Validated,
 - Not validated.
- Possibility for physicians to receive a certificate, confirming participation in the APP (which can be used for quality-oriented patient communication)
 - o Yes,
 - o No.
- Type of intervention
 - o Tools for patients (digital and non-digital reminder),
 - Digital tools for physicians (simplified generation of medication plans, appointment management, patients can document data that physician have access to),
 - Comprehensive package including both tools for patients and physicians, which will be based on behavioral and knowledge-related patient discussions.

- Time commitment per patient and quarter of the year, to implement the APP
 - o 10 mins for practice personnel,
 - o 30 mins for practice personnel,
 - o 30 mins for physician.
- Reimbursement for APP participation, per included patient and quarter of the year
 - 0 20 €,
 - 0 50 €,
 - 0 80 €.

Based on the full-factorial design, including the complete set of combinations of all attribute levels and choice situations, all possible main and interaction effects could be estimated. However, the full-factorial design is not feasible in a survey due to the high number of choices and its complexity. Thus, a fractional-factorial design was considered (using IBM SPSS Statistic software), consisting of a subset of all theoretically possible choice sets with which all effects of interest still can be estimated. The fractional-factorial design of this study contained 16 different choice sets (each with two alternative treatment options A and B). An opt-out option was added to each of the generated choice sets allowing the physicians to state that they would neither support alternative A nor alternative B. An example of a choice set is given in Figure 1.

Data Collection and Statistical Analysis

Demographic characteristics and additional information about the implementation of APPs were documented by participating physicians or collected by trained interviewers on paper-and-pencil or online questionnaires.

Before data analysis, the consistency of given answers within the DCE choice situations was checked. Therefore, the so-called "straight-liners" (respondents who decided always for the same option, eg, in all 16 choice situations option A was selected) were identified and excluded from the analysis set.

Physicians' characteristics were analyzed using descriptive statistics, including absolute and relative frequencies for categorical variables and summary statistics (eg, mean, standard deviation) for continuous variables. With regard to the DCE data, the influence of different attribute levels on the probability of a physician's choice of a specific APP alternative as well as utility levels for each attribute level were estimated, based on a multinomial logit model (MNL). The

relative importance of each attribute for the overall decision for/against an APP alternative was calculated based on the absolute distance between the minimum and maximum value/coefficients of the level of each attribute. The ratio of the coefficient distance of each attribute to the sum of the absolute coefficient distances of all attributes were interpreted as the relative importance.²²

Relative importance of the different attributes was compared between specific subgroups of interviewed physician. Mainly, it was investigated whether preferences differ by age, gender and specialty of the physician as well as by her/his experience and the type of practice. In addition, the interaction effects between individual characteristics and attributes were estimated. The specific information to define respective subgroups was collected at the end of the survey. Analyses were done with Stata/MP 14, IBM SPSS Statistic software (version 20) and Microsoft Excel 2016.

Results

Study Population

The results of 222 physicians were included in the DCE analysis, after exclusion of the outcomes of 4 physicians (3 GPs and 1 neurologist) due to inconsistent answers. Of these, 47 were GPs, 47 cardiologists, 83 neurologists, and 45 ophthalmologists. The mean age was 49.6 (SD 8.8) years, and 36.5% were female (Table 1). On average, physicians treated 1691.8 patients per quarter, and they had a mean job experience of 13.1 years (SD 9.1).

Physicians' APP Preferences

An initial descriptive analysis showed that in 31.9% of all decision situations (in total 3,552 different choice situations – 16 choice tasks per physician), physicians selected the "opt-out option" and, thus, decided not to participate in none of the provided APP alternative. 77.5% of all the interviewed physicians decided at least once for an opt-out alternative in one of the 16 choice cards, whereas 51.8% chose the opt-out alternative in at least 5 choice situations. Ophthalmologists were more often choosing for an opt-out alternative whereas cardiologists showed the lowest rate of choices of opt-out alternative (53.3% versus 40.4% with at least 5 opt-out).

Table 2 shows the results of the multivariable logistic regression that included as a dependent variable whether the physician chose the opt-out option or one of the opt-in options (either A or B). The attribute levels were encoded as dummy variables: "best option" included versus "best

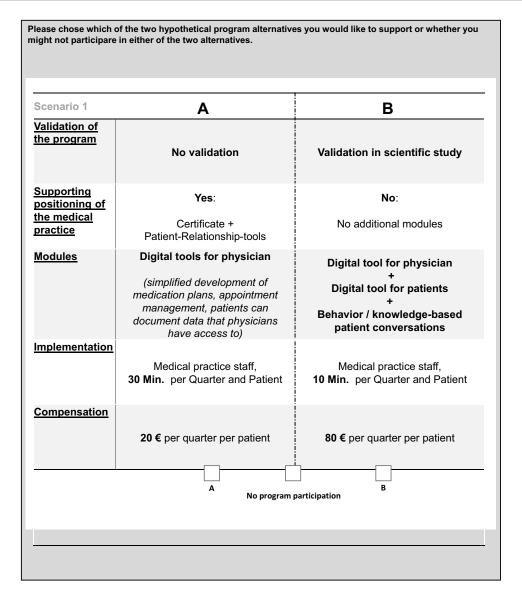


Figure I An example of the applied choice sets. The DCE card shows two different hypothetical APP alternatives derived from a combination of the defined attributes and the respective levels. Physicians needed to decide between option A, B or an opt-out option.

Notes: The figure shows one choice set as presented to participating physicians. All in all, physicians were asked to decide on 16 different choice sets.

option" not included in the choice (e.g., either option A or B include 80€ of compensation vs neither A nor B include 80€ of compensation). Validation status of APP and the possibility to receive a certificate were dropped by the model since the attributes are presented with just two levels.

Table 2 presents the results from the logistic regression analysis investigating the effect of the variation in attribute levels and physicians' as well as their practices' characteristics on the probability to choose the opt-out option. It can be seen that if "the best alternative" (comprehensive package as type of intervention, 10 mins of time commitment from the clinic staff and 80€ compensation) was included

as an option in one of the alternatives, the probability to choose the opt-out option was significantly lower (odds ratios [OR]: 0.789, 0.396 and 0.512, respectively). In addition, neurologists have a significantly lower probability to choose the opt-out option, as well as physicians working in a single practice rather than other types of medical practice (OR: 0.787 and 1.367, respectively). Physicians with more than 20 years of practice experience have a significantly lower probability to choose the opt-out option than physicians with ≤ 10 years of practice experience (OR: 1.367).

Figure 2 shows the results of the MNL regression analyses. Overall, three attributes were proven to be

Table I Characteristics of Interviewed Physicians

	Overall Sample	GPs	Cardiologists	Neurologists	Ophthalmologists
N (%)	222 (100.0%)	47 (21.2%)	47 (21.2%)	83 (37.4%)	45 (20.3%)
Age in years - mean (Valid N SD)	49.6 (164 8.8)	50.9 (36 8.9)	48.9 (34 8.8)	51.1 (66 7.9)	44.9 (28 9.4)
Female gender – N (%)	81 (36.5)	25 (53.2)	7 (14.9)	32 (38.6)	17 (37.7)
Type of medical practice					
Single practice – N (%)	74 (37.6)	17 (36.2)	12 (25,5)	31 (37.3.)	15 (33.3)
Joint/community practice - N (%)	78 (39.6)	17 (36.2)	19 (40,4)	27 (32.5)	14 (31.1)
Practice cooperation - N (%)	10 (5.1)	2 (4.3)	I (2.I)	5 (6.0)	2 (4.4)
Medical care center- N (%)	28 (14.2)	2 (4.3)	8 (17.0)	9 (10.8)	9 (20.0)
Others- N (%)	7 (3.6)	I (2.I)	2 (4.3)	4 (4.8)	0 (0.0)
Patients per quarter – mean (Valid N SD)	1,691.8 (159	1,579.1 (35	1,667.7 (31	1,263.3 (67	2,976.5 (26
	1526.8)	1177.9)	1537.9)	1074.1)	2175.9)
Total number of employees – mean (Valid N SD)	9.4 (154 10.5)	6.3 (35 5.0)	10.3 (32 8.3)	7.0 (60 6.3)	17.3 (27 18.9)
Years since start of practice – mean (valid N SD)	13.1 (169 9.1)	15.2 (38 10.2)	10.4 (34 8.2)	13.7 (67 8.9)	12.0 (30 8.8)
Number of physicians – mean (Valid N \mid SD)	2.6 (158 2.3)	2.1 (35 1.7)	2.6 (34 2.1)	2.4 (62 2.4)	3.5 (27 2.9)
Location of the practice					
City Center – N (%)	124 (63.9)	18 (38.3)	30 (63,8)	48 (57.8)	29 (64.4)
Countryside – N (%)	23 (11.9)	8 (17.0)	4 (8.5)	5 (6.0)	6 (13.3)
Suburban – N (%)	33 (17.0)	11 (23.4)	5 (10.6)	14 (16.9)	3 (6.7)
Medical Center – N (%)	14 (7.2)	4 (8.5)	I (2.I)	7 (8.4)	2 (4.4)

important from a physicians' perspective: time commitment to carry out the program (34.8% importance), reimbursement (33.3%), and validation status of the program (23.7%). The remaining attributes (type of intervention: 3.6%; possibility to receive a certificate: 4.7%) were much less important for a physician's decision to participate in a specific APP.

Participating physicians preferred APP alternatives characterized by a minimal time commitment (β =1.456, p<0.001), but also by high reimbursement for work (β =1.392, p<0.001) as well as by a "positive validation status" (β =0.990, p<0.001), the "possibility to receive a certificate" (β =0.197, p<0.001) and the provision of "tools for both physicians and patients" (β =0.150, p<0.001) (Figure 2).

Subgroup Analysis

The subgroup analyses indicated heterogeneity among physicians with respect to preferences (Figure 3). In particular, physicians aged more than 50 years had a stronger relative preference for reimbursement (39.4%) compared to physicians aged ≤50 years, whose preferences were mainly determined by time commitment (35.4%). The interaction effect between the two age subgroups was statistically significant for the choice for having a certificate (p=0.011) and for choosing a higher compensation level (20€ vs 50€: p=0.044; 20€ vs 80€: p=0.029). Among male physicians, the decision for/

against an APPs was mainly determined by reimbursement (37.3%) and time commitment (34.8%), whereas for female physicians' reimbursement was less important (24.0%), while the validation status of the APPs (26.7%) and the type of intervention (8.9%) appeared more important. With regard to the differences in the interaction effect among genders, the compensation attribute showed statistically significant differences (20ε vs 50ε : p=0.065; 20ε vs 80ε : p=0.003).

Neurologists and cardiologists attributed roughly equal importance to time commitment and reimbursement. GPs' decisions were also mainly driven by these two attributes, with even a greater focus on reimbursement (39.3%). In contrast, time commitment was most important to ophthalmologists (44.3%) followed by the validation status of the APP (26.9%), whereas reimbursement had a lower relative importance (19.0%). The differences in attribute importance among type of physicians was statistically significant for cardiologist in comparison with GPs with regard to the time compensation (10 mins staff vs 30 mins physician: p=0.035) and for ophthalmologists compared to GPs regarding the monetary compensation (20 eV vs 50 eV : p=0.002; 20 eV vs 80 eV : p<0.001).

When stratifying physicians by work experience, relative importance of the reimbursement increased with the number of years in practice. In parallel, the relative importance of the validation status decreased. The estimated interaction effects showed that the importance attributed to the validation of the

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Table 2 Logistic Regression Exploring the Probability to Choose the Opt-Out Option

			Odds Ratio	95% Confidence Interval	p-value
Attribute Levels		N of Decisions			
Intervention tool	No option had the "comprehensive package tool"	888	Reference		
	At least one option had the "comprehensive package tool"	2,664	0.789	0.649-0.960	0.018
Time commitment	No option was "10 mins of the staff"	888	Reference		
	At least one option was "10 mins of the staff"	2,664	0.396	0.328-0.477	p<0.001
Compensation	No option was "80€"	1,776	Reference		
	At least one option was "80€"	1,776	0.512	0.431-0.609	p<0.001
Physician characteristics		N of physicians			
Age ≤50 years Age >50 years		140 82	Reference 0.938	0.751-1.172	p>0.100
Male gender Female gender		141 81	Reference	0.984–1.433	0.073
GP Cardiologist Neurologist Ophthalmologist		47 47 83 45	Reference 0.916 0.787 1.253	0.692-1.213 0.621-0.998 0.958-1.640	p>0.100 0.048 0.099
≤10 years of practice ≤20 years of practice >20 years of practice		84 46 38	Reference 1.164 0.560	0.915–1.481 0.421–0.745	p>0.100 p<0.001
Single practice Other type of medical practice		148 74	Reference	1.133–1.649	0.001

program statistically differed among the years of practice categories (\leq 10 years vs \leq 20 years: p=0.033; \leq 10 years vs \geq 20 years: p<0.001), as well as monetary compensation for physician with \geq 20 years practice compared to the ones with \leq 10 years' experience (20ϵ vs 50ϵ : p=0.026; 20ϵ vs 80ϵ : p=0.009).

Regarding the type of practice, single practices considered compensation more important (36.6%) than other types of practices (29.2%), but the validation status of the APPs less important (20.2% versus 25.1%). The importance of a validated program was statistically significant between the two subgroups (p=0.009), as well as the type of intervention tool (tool for physicians vs tool for patients: p=0.021).

Discussion

Engaging physicians in the conduct of APPs is a key element driving the success of these programs. However, their willingness to participate in APPs has not been investigated so far. Hence, the aim of our study was to explore physicians' preferences with regard to the implementation and of APPs.

In our study interviewed physicians expressed a preference for APPs that are characterized by high reimbursement, low time commitment, a positive validation status, the possibility to get a certificate, and intervention tools for both physicians and patients. Reimbursement, time commitment, and validation status were the most important characteristics accounting for 91.8% of the overall relative preference of physicians for/against an APP. Time commitment and compensation were also found to be the main drivers of physician's optout decision. In that respect, additional time commitment could potentially be made up for by additional compensation. The conditional logit used to compute the tradeoffs identified in our analysis showed that 24.8€ were

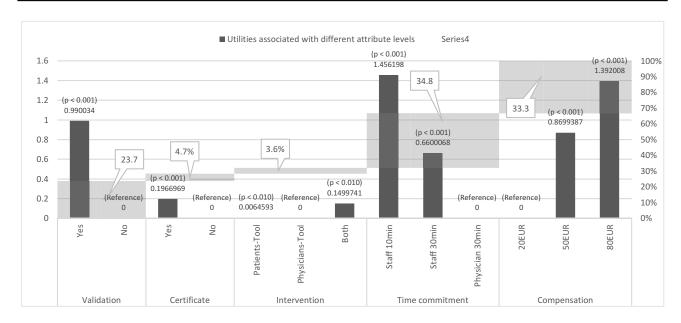


Figure 2 The results of the conditional logit estimation based on 222 physicians. For each attribute describing alternative APP options, its relative importance for overall (hypothetical) physicians' decisions for/against a specific APP is reported in % (light grey areas). Furthermore, utility per attribute level including significance of utility differences in comparison to the reference attribute level is shown (black bars).

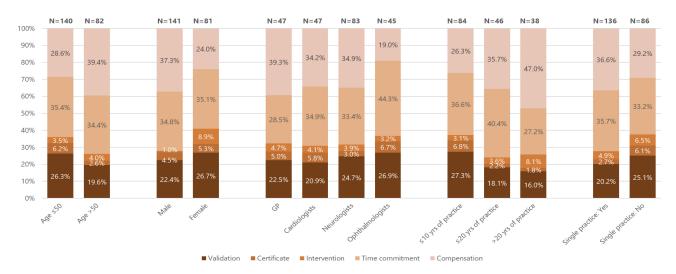


Figure 3 The relative importance of the different attributes across subgroups.

expected for a commitment of 30 mins of the stuff instead of 10 mins, and 69.2€ were expected for a 30 mins commitment by the physician instead of 10 mins of the staff.

However, preferences differed across subgroups. Reimbursement was more important for older versus younger physicians, men versus women, and physicians who had been in practice longer vs shorter. Neurologists, cardiologists and GPs showed similar results with regard to the high importance of time commitment and reimbursement, while for GPs reimbursement appeared more important than time commitment. In contrast, for

ophthalmologists, time compensation had the greatest relative importance, followed by the validation status of the APP. Reimbursement was of much lower importance. Preference heterogeneity with regard to time commitment and opportunity cost might be influenced by differences in wage (specialist have higher wages, thus, they would assume a higher compensation for the immediate loss of productivity).

Even if patients are usually in the focus with respect to medication intake behavior, physicians play a pivotal role when it comes to ensuring adherence. Patient—physician relation is largely determined by physicians' empathy and their ability to support patient' adherence to treatment. Moreover, physicians might be inquired about their patients' treatment adherence, hence, physicians are among the ones recognizing and addressing potential non-adherence. However, only few studies have investigated the role of physicians regarding adherence.²³ Furthermore, the available studies have analyzed physicians' perspectives and strategies to handle patient nonadherence in a more descriptive way. A multi-country study showed that physicians generally do not employ standardized tools to monitor asthma control or to manage its treatment with patients. 10 This may indicate that adequate engagement of physicians into APPs bears great potential for both physicians and patients. Another study analyzed physician participation in clinical research and reported a generally low willingness of physicians to participate in such studies. This was mainly related to inadequate financial and nonfinancial incentives and inadequate training that were not in line with physicians' interests.²⁴ Two studies investigated physicians' trade off with respect to financial and nonfinancial attributes of general patient supporting programs. With regard to cancer screening activities, GPs' trade-offs for screening incentives depend on the type of cancer: none of the attributes showed to be dominant over the others with respect to all the types of cancers (including the financial attribute).²⁵ The interaction between the payment and the issue of leaflets was never significant, explained by the fact that the leaflet was not an attractive incentive per se and the association with financial rewards did not change its intrinsic value. Another DCE study estimated the willingness to pay (WTP) by physicians for chronic disease management.²⁶ The highest WTP was associated with the duties of the nurse coordinators, underscoring the great importance of time commitment. The aspects determining physician participation in clinical research in this study and our results regarding factors important for physicians' participation in APPs partly overlap, specifically financial and nonfinancial interest. Hence, these appear to be general aspects to be considered in any form of patient involvement.²⁷

The strengths of our study include the study design, methodology, and conduct. First, we developed the questionnaire involving a literature review, focus groups meetings and obtaining expert advice. Second, DCE is an established method to investigate preferences circumventing socially desired responses. While a DCE design might be perceived as difficult to understand and complex, we believe that physicians are trained to process complex information. In addition, we provided detailed explanations and the choice situations were supported by structured choice sets. Third, sampling physicians from across

Germany and among different specialties ensured representativeness and allowed us to investigate potential differences by specialization.

However, some limitations need to be considered when interpreting our results. First, despite random and widescaled invitation of physicians, we cannot rule out the fact that those participating in our study already showed a higher preponderance for engagement in APPs than those who declined to participate. Second, our study cannot account for local differences (city vs countryside; differences among different German regions) regarding the organization of health care provision and established treatment patterns. For example, in some regions, the follow-up and the control of proper medication adherence might be done by specialists while in other regions specialists may be the initial prescribers and follow-up is taken care of by GPs. Finally, the low-response rate in this study as well as the high "opt-out" rate might indicate a generally low interest of physicians in engaging in and carrying out APPs. Moreover, the high number of choice tasks might have discouraged physicians to complete the questionnaire (mainly applicable for questionnaires completed via post). However, for a successful implementation of APPs physicians are crucial; hence, better understanding their views and preferences will inform the success of such programs.

Our study highlights the importance of considering physicians' preferences when designing, implementing and promoting APPs. Engaging physicians prior to implementing an APP and considering their preferences may enhance the success of the programs for their and the patients' benefit.

Ethics Approval

An ethical approval was not required due to Section 15 (1) of the Professional Code for Physician in Mecklenburg-Western Pomerania. Based on that an ethical approval is not required if only anonymized data (no personal data) regarding the opinion of physicians are collected.

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Disclosure

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