Supplementary Online Content

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This supplementary material has been provided by the authors to give readers additional information about their work.

eMethods. Study Participants and Intervention

The study was included as part of a large megastudy conducted in fall 2020 by Behavior Change for Good in partnership with Penn Medicine and Geisinger Health (see Milkman et al, 2021). Participants in the megastudy included all patients with new or routine (non-sick) primary care appointments at Penn Medicine between September 24, 2020 and March 31, 2021 and at Geisinger Health between September 28, 2020 and March 31, 2021 who met the following eligibility criteria: (1) they had a cell phone number recorded in their electronic health record, (2) they had not opted out of receiving SMS appointment reminders from their healthcare provider or asked not to be contacted for research purposes, (3) they did not have a documented allergy or adverse reaction to the flu vaccine and (4) they had not yet received an influenza vaccine in the 2020-2021 flu season according to their electronic health record.

A total of 74,811 patients were randomized to one of the 19 experimental conditions or a Usual Care Control condition that did not receive any text-based reminders to get a flu vaccine. Randomization was blockstratified with randomly-selected block sizes of 40, 80, and 120, and took place automatically based on patient lists generated via the electronic health record. All patients received standard appointment reminders (the usual care). Full details of the megastudy are reported in Milkman et al. (2021). The current paper reports results from the two messaging conditions designed by the authors as a well as a comparison with the Usual Care condition.

Patients in the Usual Care Control condition received only the standard appointment SMS reminders from their health system, indicating the date, time, and location of their appointments. Patients in the two messaging conditions received a sequence of three back-to-back SMS messages at 6 pm the day before their appointment, as shown in Figure 1. The first SMS contained general information, the second showed a photo of a flu shot vial that was labeled as "your flu dose" in the Reserved condition, and the third SMS used "reserved" vs. "available" language. The dependent variable was a binary indicator for receipt of a flu shot on the date of scheduled appointment

As shown in Figure 2, not all participants received their allocated messages. Phone carriers blocked some participants in the Available condition from receiving the third SMS that comprised the manipulated portion of the message. Among participants in the Available condition, 539 were sent messages between September 23 and October 11, 2020 worded as "Reply Y if you would like to receive a flu shot at your appt tomorrow, N if not." For 85 of these patients (16%) the third SMS was not delivered. Because the message failures were limited to the Available question, we altered the wording of the message in an attempt to avoid the carrier filters that blocked the message. On October 12 and 13, 2020, participants in the Available condition received the wording "If you would like to receive a flu shot at your appt tomorrow reply Y, or N if you don't" but message delivery failed for 12 out of the 95 participants (13%). Beginning October 14, 2020 participants in the Available condition received the wording "Flu shots will be available at your appt tomorrow. Reply Y if you would like to receive one, N if not," which resulted in very few additional message failures (11, or <1% out of 2686 participants). All told, 108 out of 3320 participants in the Available condition experienced message delivery failure compared to only 7 out of 3347 in the Reserved condition. Some additional participants had message delivery failures for other reasons. All participants who experienced message delivery failure were retained in the analysis to preserve the intent to treat analytic plan and to avoid selection bias that might make the two message conditions non comparable to the no message Usual Care Control condition. A robustness check analysis (not shown) repeated our main regression after excluding the 174 participants who experienced message delivery failures. This analysis showed a 0.8 percentage point difference (p=.42) in vaccination rates between the Reserved and Available conditions.

eResults. Preregistered and Exploratory Analyses

Participant characteristics

Characteristics of participants in each study condition are shown in Table S1. There were no statistically significant differences in key sociodemographic characteristics across the two study arms in the present study or in the Usual Care Control.

Main pre-registered analysis: Logistic model

Pre-registered analysis plans are available at <u>https://aspredicted.org/rf4hm.pdf</u> and <u>https://aspredicted.org/np99d.pdf</u>. <u>Table 2</u> and <u>eTable 2</u> show pre-registered linear probability model regressions and logistic regressions including all participants in the Reserved and Available conditions. Vaccination rates do not

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differ statistically between the two message conditions, but vaccination rates were higher among patients from Penn Medicine compared to Geisinger Health, higher among older patients, lower among Black patients compared to White patients, and higher for men than for women. In addition, vaccination rates were higher among patients who had received a flu shot the previous year and higher among patients who had appointments earlier in the season.

Additional pre-registered analyses: Appointment and previous vaccination

eTable 3shows additional pre-registered analyses. We analyzed separately patients who had appointments prior to Dec 31, 2020 from those who had appointments on or after Jan 1, 2021. The effect of the Reserved message relative to the Available message was directionally larger for patients with earlier appointments, but even among this group the 1.6 percentage point difference was not statistically significant (p=.24).

The <u>Table 1</u> percentages suggest that the Reserved message was effective only for patients who did not receive a flu shot last year. The comparison between the Reserved vs. Available conditions for patients who did not receive a flu shot in the previous year showed a marginal effect (p=.099), whereas among patients who did vaccinate the previous year there was no effect (p=.954). A pre-registered analysis shown in eTable 4, however, did not detect a significant interaction between message condition (Reserved vs. Available) and previous flu shot status (p=.355 for the interaction).

Exploratory (post-hoc) analyses: Usual Care Control comparison and race/ethnicity

In exploratory analyses (eTable 5) we compared vaccination uptake in each of the message conditions to the Usual Care Control condition. Compared to the Usual Care Control, the Reserved message increased vaccination by 2.8 percentage points (p=.004), while the Available message increased vaccination by only 1.8 percentage points (p=.069).

An exploratory analysis shown in eTable 6 examined moderation by race. When comparing the Reserved and Available conditions, the message effect for Black patients was not significantly smaller than for White patients (p=.238 for the interaction). However, when comparing the Reserved condition to the Usual Care Control, the message effect was significantly smaller for Black than for White patients (p=.012 for the interaction). Indeed, the combined effect of the Reserved vs. Usual Care and the Available vs. Usual Care interactions was significantly greater than zero, indicating that the benefit of seeing any message relative to no message was larger for White patients than for Black patients. These results suggest that message effectiveness may vary across patient populations.

eFigure. Mediation Analysis



Indirect B= +.032, (95%CI: .018, .045) Direct B= -0.18, (95%CI: -.036, .000) Total B= +.014 (95%CI: -.009, .036)

Variable	Condition		
	Reserved (n=3375)	Available (n=3351)	Usual Care (n=3432)
Site			
Penn site	53.51%	53.06%	53.50%
Geisinger site	46.49%	46.94%	46.50%
Age (mean)	50.76	50.60	50.48
Race/Ethnicity			
Asian	2.16%	2.33%	2.30%
Black	19.91%	20.47%	20.40%
Hispanic	4.71%	4.86%	5.01%
Other	2.99%	3.73%	3.64%
White	70.22%	68.61%	68.65%
Gender			
Male	44.21%	44.29%	45.19%
Female	55.79%	55.71%	54.81%
Provider type			
Physician	76.36%	75.02%	76.17%
Physician assistant	9.36%	10.21%	9.76%
Resident	4.89%	5.94%	5.07%
Other	0.00%	0.06%	0.00%
Nurse practitioner	9.39%	8.77%	9.00%
Received influenza			
vaccine last year			
Yes	40.89%	40.56%	40.03%
No	59.11%	59.44%	59.97%

eTable 1. Characteristics of Participants in Each Study Condition

Note: Percentages shown are the fraction of participants in a given condition who matched each characteristic)

Variable	Model 1	Model 2
Reserved condition	1.064 (.055)	1.068 (.065)
Penn site		1.804 (.127)***
Age (centered)		1.010 (.002)***
Asian race/ethnicity		1.006 (.203)
Black race/ethnicity		0.699 (.061)***
Hispanic race/ethnicity		0.780 (.119)
Other race/ethnicity		1.068 (.190)
White race/ethnicity		REF
Male gender		1.110 (.069)*
Physician provider		0.992 (.109)
Physician assistant provider		0.996 (.149)
Resident provider		0.710 (.127)#
Nurse practitioner provider		REF
Flu shot last year		4.257 (.266)***
Time since study start (centered)		0.999 (.002)
Time squared (centered)		1.000 (.000)***
Constant	0.497 (.018)***	0.148 (.019)***
Ν	6726	6726
Pseudo R ²	.0002	.2500

eTable 2. Odds Ratios (SEs) From Preregistered Logistic Regression Results Comparing Reserved and Available Conditions

eTable 3. Coefficients (SEs) From Prere	egistered Linear	Probability Model
Regression for Patients	s With Appointm	ients Before vs /	After December 31

Variable	Model 1 Appointment	Model 2 Appointment
	Sep 23-Dec 31	Jan1 -Mar 31
Reserved condition	.016 (.014)	001 (.012)
Penn site	.159 (.016)***	007 (.014)
Age (centered)	.002 (.000)***	.000 (.0000)
Asian race/ethnicity	036 (.046)	.064 (.044)
Black race/ethnicity	105 (.020)***	.019 (.017)
Hispanic race/ethnicity	055 (.035)	008 (.027)
Other race/ethnicity	033 (.042)	.068 (.032)*
White race/ethnicity	REF	REF
Male gender	.013 (.014)	.023 (.012)#
Physician provider	.005 (.025)	011 (.022)
Physician assistant provider	021 (.034)	.019 (.028)
Resident provider	093 (.040)*	005 (.032)
Nurse practitioner provider	REF	REF
Flu shot last year	.348 (.015)***	.116 (.013)***
Time since study start	.002 (.001)#	007 (.003)*
Time squared	000 (.000)***	.000 (.000)
Constant	012 (.050)	.326 (.075)***
Ν	4114	2612
Adj R ²	.207	.071

eTable 4. Coefficients (SEs) From Preregistered Linear Probability Model <u>Regression Results Examining Moderation by</u> Previous Vaccination

Variable	Model 1
Reserved condition	.018 (.013)
Penn site	0.095 (.011)***
Age (centered)	.002 (.000)***
Asian race/ethnicity	008 (0.34)
Black race/ethnicity	059 (.014)***
Hispanic race/ethnicity	034 (.023)
Other race/ethnicity	.012 (.028)
White race/ethnicity	REF
Male gender	.018 (.010)#
Physician provider	.002 (.018)
Physician assistant provider	.003 (.024)
Resident provider	046 (.027)
Nurse practitioner provider	REF
Flu shot last year	.275 (.014)***
Time since study start	003 (.000)***
Time squared	000 (.000)
Flu shot last year x Reserve	019 (.020)
Constant	.177 (.020)***
N	6726
Adj R ²	.275

eTable 5. Coefficients (SEs) From Exploratory Linear Probability Model Regression Results Comparing Reserved, Available, and Usual Care Control Conditions

Variable	Model 1	Model 2
Reserved condition	.033 (.011)**	.028 (.010)**
Available condition	.019 (.011)#	.018 (.010)#
Penn site		.091 (.009)***
Age (centered)		.002 (.0002)***
Asian race/ethnicity		.013 (.027)
Black race/ethnicity		045 (.011)***
Hispanic race/ethnicity		026 (.019)
Other race/ethnicity		.006 (.023)
White race/ethnicity		REF
Male gender		.016 (.008)*
Physician provider		.017 (.014)
Physician assistant provider		.013 (.019)
Resident provider		002 (.022)*
Nurse practitioner provider		REF
Flu shot last year		.260 (.008)***
Time since study start		003 (.0003)***
Time squared		.000 (.000)
Constant	.313 (.008)***	.150 (.017)***
Ν	10,158	10,158
Adj R ²	.0006	.2748
F-test comparing Reserved to	F(1,10155)=1.48,	F(1,10142)=1.14,
Available	p=.22	p=.29

#p<.10, *p<.05, **p<.01 ***p<.001. Note: The Usual Care Control condition did not have the option to reply (as they did not receive a study message). Consequently the replied variable is not included in these analyses.

Variable	Model 1	Model 2
	2 conditions	3 conditions
Reserved condition	.016 (.011)	.040 (.011)***
Available condition		.024 (.011)*
Penn site	.095 (.011)***	.090 (.009)***
Age (centered)	.002 (.000)***	.002 (.0002)***
Asian race/ethnicity	008 (.034)	.014 (.027)
Black race/ethnicity	044 (.019)*	014 (.018)
Hispanic race/ethnicity	033 (.023)	025 (.019)
Other race/ethnicity	.013 (.028)	.007 (.023)
White race/ethnicity	REF	REF
Male gender	.018 (.010)#	.017 (.008)*
Physician provider	.002 (.018)	.017 (.014)
Physician assistant provider	.003 (.024)	.013 (.019)
Resident provider	047 (.027)#	002 (.022)
Nurse practitioner provider	REF	REF
Flu shot last year	.266 (.010)***	.260 (.008)***
Time since study start	003 (.000)***	003 (.0003)***
Time squared	000 (.000)	.000 (.000)
Black x Reserve	029 (.025)	061 (.024)*
Black x Available		032 (.024)
Constant	.177 (.020)***	.143 (.017)***
Ν	6726	10,158
Adj R ²	.275	.275
Test that coefficients for the two		F(1,10140) = 5.04,
interactions are jointly > 0		p=.025

eTable 6. Coefficients (SEs) From Exploratory Linear Probability Model Regression Results Examining Moderation by Black vs White Race