Supplementary Appendix

Supplementary Methods

- Interventions were input by the Handshake antimicrobial stewardship team after rounds using the institutional Epic Antimicrobial Stewardship Standard Operation Procedure. Definitions of Antimicrobial Stewardship Intervention types:
 - Allergy clarification: Use when documenting/updating reactions to antimicrobial allergies
 - Broaden coverage/address absent therapy: Use when empiric or pathogenrecommended antimicrobial therapy is recommended to be broadened due to insufficient coverage.
 - De-escalation: Use when antimicrobial therapy has been de-escalated either empirically or based on culture results.
 - Diagnostic-related: Use when recommending a diagnostic test (ex. MRSA nasal swab).
 - Discontinue: Use when an antimicrobial is discontinued and not replaced with another.
 - Dose optimization: Use when an antimicrobial dose and/or frequency is changed.
 - Duration of therapy: Use when a duration of therapy is extended.
 - Drug-interaction/Contraindicated: Use when a patient's antimicrobial is switched to an alternative safer regimen due to drug interaction or contraindication.

- Monitoring-related: Use when recommending a non-therapeutic drug level
 lab (ex. Requesting additional susceptibilities from microbiology).
- Route change intravenous to oral: Use when changing the same antibiotic
 from intravenous to oral or per tube.
- Interventions were reviewed for acceptance by the ID pharmacist the next business day and any recommendations that were implemented within 24 hours were counted as accepted.
- Partial acceptance was accounted for by splitting recommendations into sub-recommendations and marking each one as accepted or rejected (i.e. the recommendation was to switch from ceftriaxone to amoxicillin-clavulanate and shorten the duration by 3 days, but only the duration was shortened. This would be recorded as two separate recommendations with the former being rejected and the latter being accepted).
- Antimicrobial Spectrum Index we calculated an antimicrobial spectrum index to describe potential shifts in broad to narrow spectrum agents. We utilized the methodology described by Dr. Gerber (reference 10) but modified and updated it to include newer antimicrobials as described in detail by Dr. Ilges (reference 11).
- dDOT/tDOT as described in detail by Dr. Moehring, et al. (reference 12) we calculated digestive days of therapy (dDOT) over total days of therapy (tDOT) as a marker of intravenous to oral route conversions. dDOT as defined by NHSN was the number of days of therapy administered via an oral, tube, or per rectum route. tDOT was defined as the dDOT plus intravenous DOT.

Supplementary Tables

Table 1. Existing Antimicrobial Stewardship Program Details

ASP Strategy	Affected Antimicrobials			
Prior Authorization	Amphotericin B, liposomal, Artesunate, Cabotegravir-			
	Rilpivirine, Cefiderocol, Ceftazidime-avibactam,			
	Ceftolozane-tazobactam, Chloramphenicol, Dalbavancin,			
	Eravacycline, Imipenem-relebactam, Meropenem-			
	vaborbactam, Tedizolid			
72 hour Prospective audit	· · ·			
and feedback	Foscarnet, Maribavir, Quinupristin/dalfopristin, inhaled			
	Ribavirin, Telavancin, Tigecycline			
Restricted by Use Criteria	Aztreonam, Baloxavir, Letermovir, Remdesivir, SUBA-			
	itraconazole			
Guidelines and order sets	C. difficile Treatment Guidelines			
	COVID-19 Treatment Guidelines			
	Intra-abdominal Infection Guidelines			
	Pneumonia Treatment Guidelines			
	BioFire Pneumonia Panel Treatment Guidelines			
	Sexually Transmitted Infection Guidelines			
	Skin and Soft Tissue Infection Guidelines			
	Surgical Antimicrobial Prophylaxis Guidelines			
	Urinary Tract Infection for Hospitalized Patients			
	Guidelines			
Renal dosing policy	No automatic pharmacist renal dosing policy. However we			
	utilize renal dose context on order entry to guide			
	prescribers to the initial correct renal dose. In addition we			
	have retrospective renal dosing clinical decision support			
	rules that alert pharmacists 365 days/year when the dose			
	ordered does not match the recommended dose.			
Automatic IV to PO policy	There was a pharmacist Medication Therapy			
	Management protocol for IV to PO antimicrobials in			
	existence until December 2021 but given the low			
	utilization of pharmacists using this automatic protocol			
	(average 3 per month) compared to the number of IV to			
	PO interventions being made and accepted outside of			
	the confines of the protocol (average 55 per month) it			
	was retired.			

Table 2. Feedback Survey Results

Demographics	Medicine Residents	Hospitalists N = 49
	N = 45	
Level of Training PGY-1	16 (36%)	N/A
PGY-2	12 (27%)	
PGY-3	16 (36%)	
Other	1 (1%)	
Clinical role Hospitalists	N/A	41 (84%)
Nurse Practitioner		8 (16%)
Years worked at BJH as Hospitalists	N/A	
< 1 year		6 (12%)
1-5 years		24 (49%)
5-10 years		8 (16%)
> 10 years		11 (23%)
Have you interacted with the Handshake ASP	Yes = 35 (78%)	Yes = 43
team in the past 5 months?		(88%)

Survey Questions	Medicine	Hospitalists
	Residents	N = 42*
	N = 35	
How often do you contact the ASP team?	N/A	
Never		7 (17%)
< Once per month		5 (12%)
Once per month		12 (29%)
A few times per month		13 (31%)
Once per week		4 (10%)
Several times per week		1 (2%)
Is the frequency of contact adequate?		
Would prefer less frequently	1 (3%)	1 (2%)
Just right	31 (88.5%)	37 (88%)
Would prefer more frequently	3 (8.5%)	4 (10%)
How would you prefer to receive communications from		
the ASP team?	22 (222()	22 (222()
In-person rounds	23 (66%)	22 (52%)
Epic chat message	12 (34%)	18 (43%)
Phone call	0	1 (2%)
Prefer zero communication	0	1 (2%)
How often do you find the recommendations from ASP		
to be helpful in patient care?		
Never	0	0
Sometimes	1 (3%)	3 (7%)

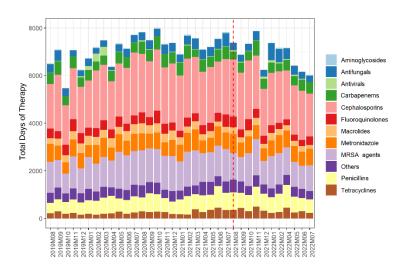
About half the time	1 (3%)	3 (7%)
Most of the time	14 (40%)	16 (38%)
Always	19 (54%)	20 (48%)
How often do you implement the ASP		
recommendations?		
Never	0	0
Sometimes	1 (3%)	3 (7%)
About half the time	3 (9%)	2 (5%)
Most of the time	19 (54%)	21 (50%)
Always	11 (30%)	16 (28%)
No response	1 (3%)	0
How easy is it for you to get in touch with members of		
the ASP team?		
Difficult	6 (17%)	1 (2%)
Neither easy nor difficult	10 (28.5%)	8 (19%)
Somewhat easy	10 (28.5%)	11 (26%)
Extremely easy	7 (20%)	22 (53%)
No response	2 (6%)	0
Would you be interested in rounding with the ASP		N/A
team as an elective?	1 (3%)	
Definitely not	9 (25.5%)	
Probably not	13 (37%)	
Probably yes	10 (28.5%)	
Definitely yes	2 (6%)	
No response		
Would you be interested in any of the following	N/A	
activities or interventions?		
Individual antibiotic use reports		15 (30%)
Peer comparison reports for antibiotic use		22 (44%)
Research collaboration with ASP		6 (12%)
Hospitalists stewardship champion		7 (14%)

^{*}While 43 survey participants responded they had interacted with the ASP team; only 42 participants completed the second part of the survey

Supplementary Figures

Figure 1. Total antibiotic usage by month and by antibiotic class excluding remdesivir, absolute (A) and percentage (B); intervention start shown as dashed red line

(A)



(B)

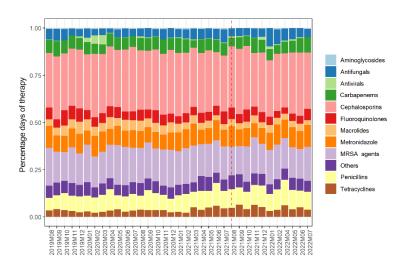


Figure 2. C. dificile infection rate by month; Intevention start red dashed line

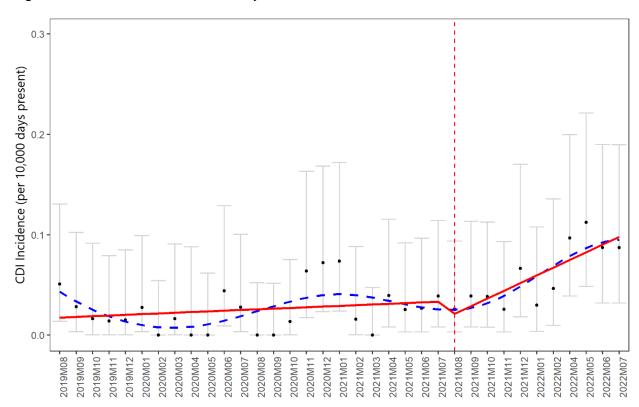
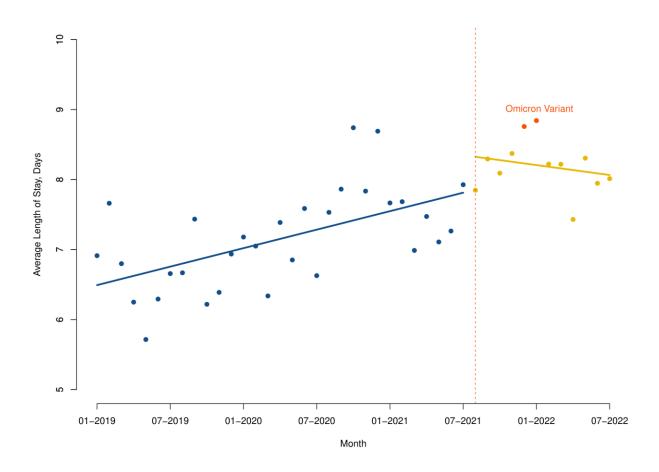


Figure 3. Length of stay by month; intervention start dashed red line; Omicron surge months labeled



Supplementary Discussion

Interrupted time series analysis of our length of stay data reveals that the slope did decrease during the post-implementation period. There wasn't a significant immediate effect at intervention (p = 0.162) or with a 3 month lag (p = 0.094). Additionally, there is no statistically significant difference in slope (p = 0.146) immediately post-intervention. However, the difference in slope is statistically significant (p = 0.039) when lagged by 3 months – this may be supportive of the theory that the intervention may have had a dampening effect in the overall secular increase in length of stay that has been seen nationally since the beginning of the pandemic.

Our *C. difficile* infection rates increased and this was reflected in the ITS, but there were numerous confounders here including a fairly major change in our *C. difficile* testing protocol as discussed in the main text. Additionally, the trend in the slope was highly influenced by two above average post-intervention months - more data would be required for concrete conclusions to be drawn.