ORIGINAL PAPER



Evaluation of follow-up colposcopy procedures after abnormal cervical screening result across a statewide study in Mississippi

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Received: 9 February 2024 / Accepted: 7 August 2024 / Published online: 17 August 2024 This is a U.S. Government work and not under copyright protection in the US; foreign copyright protection may apply 2024

Abstract

Purpose Cervical screening is used to detect and treat precancers to prevent invasive cancers. However, successful prevention also requires adequate follow-up and treatment of individuals with abnormal screening results. The aim was to investigate demographics, clinical characteristics, and follow-up status for individuals needing colposcopy after an abnormal screening result.

Methods The STRIDES (Studying Risk to Improve DisparitiES) cohort comprises individuals undergoing cervical cancer screening and management at a Mississippi Health Department or University of Mississippi clinic. Follow-up status, demographics, and clinical data were assessed from electronic health records and, if necessary, patient navigation on individuals identified as needing a colposcopy after an abnormal screening.

Results Of the 1,458 individuals requiring colposcopy, 43.0% had the procedure within 4 months, 16.4% had a delayed procedure, and 39.5% had no documented colposcopy follow-up, with significant predictors of follow-up identified as age and cytology diagnosis. Individuals 30 + were more likely to have follow up with a colposcopy compared to individuals < 30 years (49% and 38.7%, respectively; p < .001). Individuals with cytology diagnoses of LSIL (52.9%), ASC-H (51.4%), and HSIL (62.3%) had higher percentages of adherence to follow-up guidelines (p < .001). In total, we found that 78% of individuals had some type of follow-up, including a repeat screening visit.

Conclusion Despite high cervical cancer screening rates among Mississippians, a substantial proportion did not have adequate next-step intervention. However, it is encouraging that highest risk individuals were more likely to have a colposcopy. Regardless, continuing to understand the underlying causes for incomplete follow-up is crucial for timely secondary targeted interventions to reduce cervical cancer burden, promote awareness, and improve health outcomes.

Keywords Cervical cancer · Colposcopy · ASC-US · LSIL · ASC-H · HSIL

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Introduction

Cervical cancer (CC), caused by persistent infection with carcinogenic human papillomavirus (HPV), is one of the few preventable cancers. Dramatic reductions in incidence and mortality rates have been seen in the United States (U.S.) over the last several decades due to the utilization of the Papanicolaou (Pap) smear screening. With the more recent advent of the HPVvaccine and the increased use of HPVbased testing, either alone or with Pap cytology (co-testing) as primary screening approaches, even more reductions in CC incidence and mortality are expected [1]. However, despite availability of primary and secondary prevention approaches, over 13,000 new CC cases and more than 4,000 CC deaths are expected in the U.S. in 2024 alone [2]. More than half of new CC diagnoses in the U.S. occur in individuals who are never screened or who are under-screened [3]. Beyond screening, secondary prevention of CC requires effective and timely follow-up with diagnostic colposcopy and targeted biopsies of screen-positive individuals and treatment of cervical precancers if detected. Without these steps, CC prevention fails, and screened individuals remain at elevated risk for CC [4].

Within the U.S., there are known geographic disparities with respect to cervical cancer incidence and mortality. Mississippi ranks among the top five states in the nation for CC incidence and mortality, despite having among the highest CC screening rates [5, 6]. This scenario suggests that high rates of cervical cancer in Mississippi may, in part, be due to other factors, potentially including lack of diagnostic followup and treatment among screened individuals. We sought to evaluate patient demographics, clinical characteristics, and adherence to follow-up colposcopy care for individuals with an abnormal CC screening results within a large, statewide cohort of patients undergoing screening in Mississippi.

Methods

Study population

The National Cancer Institute (NCI), Mississippi State Department of Health (MSDH), and the School of Nursing at the University of Mississippi Medical Center (UMMC) developed the STRIDES—Studying Risk to Improve DisparitiES study in 2019. STRIDES represents a statewide cohort of individuals undergoing cervical cancer screening and management at UMMC and MSDH. A detailed description of the STRIDES study design is available elsewhere (5).

Procedures

For this study, nested within the STRIDES cohort, we identified individuals with the following criteria: screened from January 2018 through August 2021 at an MSDH clinic and had an abnormal screening result with an indication for colposcopy based on the 2012 ASCCP management guidelines. Although the ASCCP published new risk-based guidelines in April 2020, most patients (80%) were screened prior to this period, and guideline adoption may take months to years depending on the setting [7]. Once the sample was obtained, we performed chart reviews of electronic health records recurrently (EHR; i.e., the MSDH histology file) through September 2022 to search for documentation indicating receipt of any cervical cancer follow-up care by an MSDH or outside provider. Receipt of follow-up care was defined according to the presence of histology resulting from a cervical biopsy in the EHR and extracted for our records. After chart reviews any individual found to be lacking follow-up documentation in the EHR were then provided to a team of patient navigators, who worked with MSDH clinic staff to contact individuals and schedule follow-up appointments.

Study variables

MSDH screening procedures included cytology with HPV triage of atypical squamous cells of undermined significance (ASC-US) for patients aged < 30 and co-testing for patients aged 30 + . Cytological diagnoses were obtained from the EHR and classified according to the Bethesda System as NILM, negative for intraepithelial lesion or malignancy; ASC-US; LSIL, low-grade intraepithelial lesion; ASC-H, atypical squamous cells cannot exclude high grade; or HSIL, high-grade intraepithelial lesion. HPV testing was performed using cobas4800 on the Thin-Prep sample collected during the Pap smear screening with results reported in the EHR. HPV genotypes were reported as HPV type 16, HPV type 18, and pooled HPV other high-risk types (HR12): 31, 33, 35, 39, 45, 51, 52, 56, 58, 59, 66, and 68.

At the time of the CC screening visit, we collected sociodemographic characteristics from the EHR. Age was categorized as < 30 years and \geq 30 years. We categorized race as White or Caucasian ("White"), Black or African American ("Black"), American Indian or Alaska Native, Asian, Native Hawaiian or Other Pacific Islander, Multiracial, Other Race, and Patient refused or Unknown. Due to low sample size, American Indian or Alaska Native (n=4), Native Hawaiian or Other Pacific Islander (n=1), Multiracial (n = 6), and Other race (n = 69) were combined into one category ("Other"). We categorized ethnicity as Non-Hispanic, Hispanic, Asian, and Native American. For analyses, race and ethnicity were combined and categorized as non-Hispanic White, non-Hispanic Black, Hispanic, and non-Hispanic other, and unknown. Smoking was recorded as never, former, current, and unknown/ missing. Body mass index (BMI; kg/m²) was categorized according to standard definitions as < 25 (underweight/ healthy weight), 25 to < 30 (overweight), 30 to < 35 (class I obesity), and >/= 35 (class II-III obesity).

The primary outcome was adherence to follow-up recommendations for a colposcopy based on the 2012 ASCCP guidelines used at the time of the screening event. Follow-up status was determined based on the evaluation of any followup documentation to determine if individuals returned for their follow-up colposcopy after an abnormal cervical cancer screening result. Individuals were placed into three groups: 1. Adherent to Follow-Up with Colposcopy (on time) = individuals who returned to their provider and had a colposcopy within the recommended 4 month timeframe; 2. Adherent to Follow-up with Colposcopy (delayed) = Individuals who returned to their provider and had a colposcopy outside of the recommended 4 month timeframe; 3. Non-Adherent to Follow-Up Recommendations = Individuals who did not return for the recommended follow-up colposcopy by September 2022. The choice of 4 months was used to indicate delayed follow-up and based on ASCCP management guidelines [7, 8]. The non-adherent group also includes individuals who may have returned for follow-up but received a cotest instead of a colposcopy.

Data analysis

We assessed patient demographics, clinical characteristics, and adherence to follow-up recommendations using descriptive statistics and reported means and standard deviations for continuous variables, as well as frequencies and percentages for categorical variables. We compared characteristics of individuals who were adherent (on time), adherent (delayed), and non-adherent to follow-up care using chi-square analysis. We estimated the odd ratios (ORs) and 95% confidence intervals (CIs) for associations of individual and clinical characteristics with follow-up using multivariable logistic regression analyses to predict follow-up vs. no follow-up within the sample. We used Kaplan-Meier methods to assess receipt of colposcopy over time among individuals with abnormal screening results and compared the survival follow-up curves by both on age and cytology using log-rank tests. Statistical analyses were performed using IBM SPSS Statistics version 28 (IBM Inc., Armonk, NY, USA). To account for potential changes in practice that may have occurred during the COVID-19 pandemic, we performed a sensitivity analysis evaluating characteristics associated with follow-up, stratified by time (pre-pandemic: February 2018-February 2020 and post-pandemic: March 2020-August 2021).

Results

Between 2018 and 2021 the STRIDES study had a population size of 32,735 individuals undergoing screening within Mississippi's public health system, with 20,792 individuals being seen at a MSDH clinic. We identified 1,458 MSDH individuals that had an abnormal result indicating need for a colposcopy based on the 2012 ASCCP guidelines. Characteristics of these individuals by follow-up status are shown in Table 1 with a full breakdown of cytology and HPV results shown in Supplemental Table S1 and Table S2. A total of 627 individuals (43.0%) with an abnormal screening result had a follow-up colposcopy procedure documented within 4 months (on time). There were 239 (16.4%) individuals that had follow-up documentation of a colposcopy procedure, but outside of the recommended four-month period with a delayed mean of 15.6 months (SD = 12.91) and a maximum of 53 months seen. Of the 592 individuals (40.6%) identified as not adherent to the follow-up guidelines, 264 (18.1%)had documentation of a return visit where they received an additional co-test instead of the recommended colposcopy. Characteristics of individuals who returned for a repeat cotest compared to those who did have follow-up care documented are shown in Supplemental Table S3.

The follow-up outcome groups had significant differences noted between adherence to follow-up recommendations by age, race/ethnicity, and cytology diagnosis shown in Table 1. Regarding age, 49.0% of individuals ≥ 30 years of age compared to 38.7% of individuals < 30 years of age were adherent to follow-up guidelines (p < 0.001). Regarding race and ethnicity (p=0.003), individuals who were Hispanic had the highest percentage (63.3%) of having follow-up care based on recommendations. Individuals listed as other had the highest percentage (48.2%) of not having documented follow-up care. When looking at cytology diagnosis (p < 0.001), individuals with a low-risk cytology diagnosis such as ASC-US (53.6%) and LSIL (31.9%) were less likely to receive follow-up compared to individuals with a more severe cytology diagnosis, such as ASC-H (25.0%) and HSIL (21.6%). Individuals with an HPV-positive NILM diagnosis (44.2%) also had a higher percentage of being less likely to receive follow-up. There was no difference in follow-up outcomes by BMI category (p = 0.105) or smoking status (p = 0.216). As expected, overall follow-up was lower during the height of the COVID-19 pandemic; however, patterns by age, race/ethnicity, and cytology were

 Table 1
 Frequencies and Chi-Square Results for Follow-up Outcomes Among Individuals Following Abnormal Cervical Cancer Screening

	Adherent: Follow-Up with Colposcopy (on time) $(n = 627)$		Adherent: Follow-Up with Colposcopy (delayed) $(n = 239)$		Non-Adherent: No Follow-	p value		
	n	row%	n	row%	n	row%		
Age							<.001	
< 30 years old	327	38.7	126	14.9	394	46.5		
\geq 30 years old	300	49.0	113	18.5	199	32.5		
Race							.003	
Non-Hispanic White	156	43.7	57	16.0	144	40.3		
Non-Hispanic Black	339	41.4	143	17.5	336	41.1		
All Hispanic	57	63.3	12	13.3	21	23.3		
Other	40	36.4	17	15.5	53	48.2		
BMI							.105	
<25	166	39.8	68	16.3	183	43.9		
25-<30	125	38.2	60	18.3	142	43.4		
30-<35	135	50.0	40	14.8	95	35.2		
35+	147	43.5	55	16.3	136	40.2		
Smoking							.216	
Never Smoker	353	43.4	125	15.4	336	41.3		
Former	84	49.1	29	17.0	58	33.9		
Current	190	40.3	85	18.0	197	41.7		
Cytology Diagnosis								
NILM ^a	57	34.5	35	21.2	73	44.2		
ASC-US	191	31.6	89	14.7	324	53.6		
LSIL	212	52.9	61	15.2	128	31.9		
ASC-H	37	51.4	17	23.6	18	25.0		
HSIL	124	62.3	32	16.1	43	21.6		

ASC-H atypical squamous cells cannot exclude high grade, ASC-US atypical squamous cells of undetermined significance, HSIL high-grade intraepithelial lesion, LSIL low-grade intraepithelial lesion, NILM negative for intraepithelial lesion or malignancy, Pap Papanicolaou test ^aNILM diagnoses were HPV 16, HPV 18, or HPV 12 HR other positive, indicating the need for further management with a colposcopy based on ASCCP 2012 guidelines

generally similar to those observed pre-pandemic (Supplemental Table 4).

We conducted multivariate logistic regression analyses to investigate factors associated with follow-up care, shown in Table 2. Overall, age (per one-year increase) was positively associated with an increased likelihood of following up with a colposcopy recommendation (OR = 1.07; 95% CI = 1.05, 1.09). When further stratified by age, younger individuals (<30 years) observed a 36% increase (OR = 1.36; 95% CI = 1.26, 1.47; p < 0.001) in the likelihood of following up with a colposcopy with every one-year increase in age. However, in individuals \geq 30 years old, the same pattern of increased likelihood of colposcopy follow-up was not significantly identified. (OR = 1.01; 95% CI = 0.98, 1.47; p = 0.556). Cytology diagnoses were also associated with increased likelihood of completing a colposcopy follow-up in the various regression models. When comparing individuals with a NILM cytology result to the full sample, those with LSIL, ASC-H, and HSIL were more likely to follow up with colposcopy, (OR = 2.67, 3.9, and 5.37, respectively; p < 0.001). Similar findings among cytological diagnoses were observed when stratified by age groups.

Supplemental Table S1 and Table S2 show the distribution of cytology screening results by HPV genotype. Among the 617 individuals \geq 30 years of age, 90.4% (n = 553) of the cytology results had an associated HPV test result, with 96.2% (n = 532) of those screening HPV positive. Among individuals < 30 with ASC-US cytology diagnosis, HPV testing was completed on 99.8% (n = 428) individuals. HPV Other HR12 was the most common result among **Table 2** Regression Results for Follow-up with a Colposcopy Based on Total Sample (n = 1458) and Age-Stratified for < 30 (n = 846) and ≥ 30 (n = 612)

Variable	Model 1: Full Sample			Model 2: < 30-Year-Old			Model 3: ≥ 30-Year-Old		
	OR	95% CI	р	OR	95% CI	р	OR	95% CI	р
Age	1.07	[1.05, 1.09]	<.001	1.36	[1.26, 1.47]	<.001	1.01	[0.98, 1.04]	.556
Race									
Non-Hispanic White	Ref		.203	Ref		.665	Ref		.737
Non-Hispanic Black	1.17	[0.86, 1.57]	.319	1.13	[0.74, 1.71]	.579	1.14	[0.72, 1.80]	.587
All Hispanic	1.72	[0.93, 3.18]	.083	1.69	[0.68, 4.21]	.260	1.17	[0.49, 2.75]	.727
Other	0.88	[0.55, 1.43]	.611	0.96	[0.51, 1.81]	.901	0.75	[0.33, 1.71]	.490
BMI									
< 25	Ref		.312	Ref		.709	Ref		.792
25- < 30	0.87	[0.63, 1.21]	.410	0.79	[0.51, 1.24]	.314	0.99	[0.58, 1.71]	.995
30- < 35	1.19	[0.84, 1.68]	.338	1.02	[0.62, 1.66]	.952	1.27	[0.74, 2.18]	.390
35 +	1.14	[0.83, 1.58]	.417	1.01	[0.65, 1.57]	.953	1.03	[0.60, 1.77]	.907
Smoking									
Never Smoker	Ref		.227	Ref		.624	Ref		.064
Current	0.86	[0.65, 1.14]	.298	0.97	[0.66, 1.44]	.891	0.62	[0.40, 0.96]	.034
Former	1.22	[0.82, 1.81]	.320	1.28	[0.74, 2.19]	.378	1.07	[0.56, 2.05]	.847
Cytology Diagnosis									
NILM	Ref		<.001				Ref		.005
ASC-US	1.18	[0.77, 1.81]	.451	Ref		<.001	1.59	[0.98, 2.58]	0.60
LSIL	2.67	[1.72, 4.14]	<.001	1.34	[.893, 2.01]	.157	2.78	[1.60, 4.81]	<.001
ASC-H	3.90	[1.91, 7.97]	<.001	6.60	[2.79, 15.62]	<.001	2.06	[0.80, 5.31]	.134
HSIL	5.37	[3.13, 9.20]	<.001	8.95	[4.97, 16.12]	<.001	2.48	[1.32, 4.68]	.005
Atypical	1.12	[0.31, 4.10]	.865				0.98	[0.26, 3.79]	.982

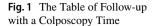
ASC-H atypical squamous cells cannot exclude high grade, ASC-US atypical squamous cells of undetermined significance, HSIL high-grade intraepithelial lesion, LSIL low-grade intraepithelial lesion, NILM negative for intraepithelial lesion or malignancy, OR Odds Ratio

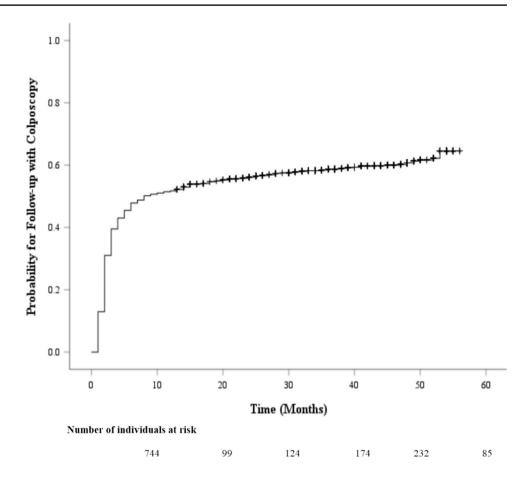
screened-positive HPV genotypes for the sample of individuals < 30 (n = 370, 86.2%) whereas HPV 16/18 was the most frequent subtype among HPV screened positive individuals \geq 30 (n = 281, 45.9%). Among individuals with HPV 16/18, both LSIL (OR = 4.00; 95% CI = 1.25, 12.79; p = 0.019) and ASC-H/HSIL (OR = 2.07, 95% CI = 0.93, 4.59, p = 0.074) cytology diagnoses results were more likely to go to colposcopy, whereas among Other HR12 only the cytology diagnosis of ASC-H/HSIL (OR = 4.16; 95% CI = 1.27, 13.67; p = 0.019) were more likely to receive a colposcopy for follow-up care (data not shown).

The Kaplan–Meier survival curve probability for followup care with a colposcopy after an abnormal screening result is displayed in Fig. 1. Most individuals who underwent a colposcopy did so within 12 months (0.5 probability by 8 months), then continued to increase at a slower rate up to 0.6 at year 5. Figure 2 provides a comparison of colposcopy probability between different age groups. The curve identifies individuals aged 30 years and older had a significantly higher probability for follow-up colposcopy at nearly 0.6 after one year and 0.7 at year 5 compared to individuals aged less than 30 years (p < 0.001). Figure 3 represents a comparison of follow-up colposcopy probability among cytological diagnoses with significant differences noted among the five cytology diagnoses (p < 0.001). Individuals with a higher risk cytology diagnosis of HSIL or ACSUS had a higher probability of following up with a colposcopy (0.7 and 0.6 after one year, respectively) compared to other diagnoses.

Discussion

Colposcopy is a critical secondary preventative diagnostic procedure used to evaluate cervical abnormalities detected during routine screening, and the adherence to guidelines ensures standardized and evidence-based management recommendations [7, 8]. In our study, we evaluated patient demographics and clinical characteristics in relation to follow-up status among individuals needing a colposcopy based on ASCCP guidelines after an abnormal cervical cancer screening result. We identified 1,458 individuals screened at a MSDH clinic, between 2018 and 2021, who needed colposcopy after their abnormal screening result. A majority (43.0%) of individuals with an abnormal screening result had a follow-up colposcopy procedure within 4 months (on time), with 59% having any follow-up colposcopy



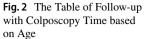


(regardless of time) and 78% with some type of follow-up including a repeat screening visit. While this is reassuring, approximately 40% of individuals with abnormal screening results are not receiving recommended diagnostic follow-up with colposcopy in our population. Thus, challenges persist to ensure that patients with abnormal cervical cancer screening results receive necessary follow-up care. Adherence to guidelines also significantly differed between age, race/ethnicity, and cytology diagnosis. The likelihood of follow-up colposcopy adherence increased with age and with increased cytology diagnosis severity (i.e., LSIL, ASC-H, and HSIL).

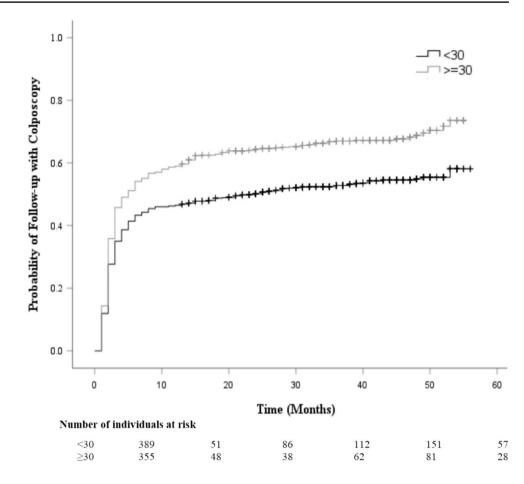
Further categorization of timely adherence to the recommended colposcopy was categorized into three groups and a significant concern identified was that 39.5% of the individuals were not adherent to the guidelines for follow-up care. Further, 328 (55.4% of the 39.5% not adherent) individuals had no documented follow-up of any kind. Thus, challenges exist to ensure that patients with abnormal cervical cancer screening results persist in receiving necessary follow-up care. A previous study, evaluating adherence to colposcopy follow-up found that the majority of individuals fell within the adherent (on-time) category, but 42.3% were not adherent, which is consistent with our findings [9]. Together, these findings identify the need to continue to explore challenges surrounding timely and appropriate cervical cancer follow-up care, particularly among Mississippi's vulnerable populations.

Martinez-Gutierrez et al. (2023) recently published a systematic review of 26 studies including 265,041 individuals from high-income countries who required follow-up after a cervical cancer screening. Over 40 factors were used to define inadequate follow-up, with younger, less educated, and lower socioeconomic status (SES) being associated with inadequate follow-up [10]. While socioeconomic variables were not included within our study analysis, this study is best interpreted within the context of the patient population seeking care at MSDH clinics. MSDH is the primary source of healthcare for Mississippians that are underinsured/ uninsured and that fall within the lower ranges of SES [11]. However, the problem of inadequate follow-up for patients at risk for cervical cancer is not limited to socioeconomic status.

Aligned with previous findings, age was also found to be a significant factor in relation to follow-up colposcopy care [10]. In our study, 46.5% of individuals aged < 30 years old were non-adherent with follow-up colposcopy care but an increased likelihood of colposcopy with every one year of increased age up to 30 was noted. In Sharp et al. (2012), who published a prospective cohort study including 2,213 individuals needing colposcopy, age was also significantly associated with not returning to the clinic for follow-up







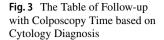
colposcopy care. In this report, and like our findings, individuals < 30 years old were found at the highest risk of non-adherence with risk of non-adherence reducing by one-third in patients aged 30–39 years and by two-thirds in those aged 40–59 [12].

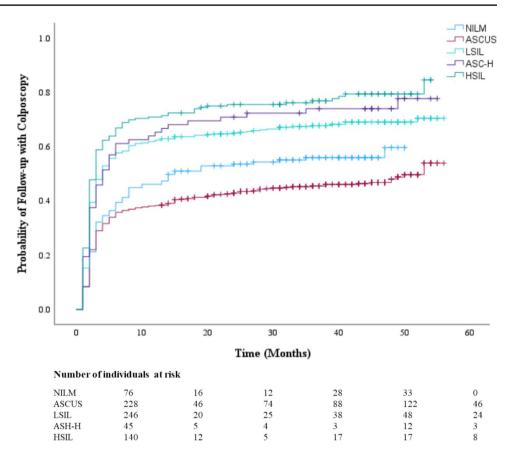
Our study found that increased likelihood of colposcopy follow-up care increased with cytology severity. Literature has found that colposcopy follow-up care tends to correlate well with high-grade abnormalities but is less efficient for women with lower-level cytological abnormalities [13]. Other studies have shown that individuals with lower cytological diagnosis (ASCUS or NILM but HPV positive) are less likely to receive a colposcopy as their follow up care as some providers and/or patients may prefer conservative management [14]. Perkins et al. (2021) found that adherence to follow-up colposcopy guidelines, within 6 months, had the following rank order: high-grade cytology (defined as HSIL & ASCH) > low-grade cytology (LSIL, ASCUS) > HPV-positive NILM cytology [15]. This same pattern was identified within our study with 62.3% of HSIL diagnosis and 51.4% of ASC-H diagnosis being adherent (on time). The fact that over 75% of patients with an HSIL diagnosis received any follow-up colposcopy during the study period is reassuring, given that this screening result is associated with an elevated risk of cervical precancer/cancer [16].

Overall, we found that approximately 45% of individuals lacking colposcopy follow-up had a repeat screening procedure. Among those, approximately 13% had a high-grade cytology result (ASC-H or HSIL). This shows a high rate of continued engagement with the healthcare system and an opportunity to schedule and plan for the necessary diagnostic and treatment services, ensuring those who need care can receive it promptly. Further, since the majority of individuals referred to colposcopy do not have cervical precancer, repeat testing can provide some level of reassurance against risk of precancer and cancer even in those who do not receive the recommended colposcopy.

Strength/limitations

Mississippi has a racially and ethnically diverse population, and one notable aspect of our study is its inclusion of a substantial percentage of African American individuals and those residing in rural areas. These groups have been underrepresented in cervical cancer research despite having a disproportionate burden of the disease and mortality [5]. This diversity is essential to display





as tailored screening strategies may result from a clearer understanding of any differences among groups [7, 8, 17]. Additionally, all individuals who went to a MSDH clinic for cervical cancer screening and follow-up during the data collection period were included in the biorepository. Other than individuals sent to outside providers, all specimens were routed through a single cytopathology laboratory, allowing for consistent and complete ascertainment of data from patients who underwent care at MSDH. Further, we performed multi-level chart reviews to collect any information from patients who sought follow-up care by an outside provider.

Even with the valuable insights obtained from our study, a few limitations should be acknowledged. First, this data was obtained from EHR. Despite best efforts, missing or incomplete information is possible. Further, while it would have been of interest to evaluate follow-up according to HPV vaccination status, this information is currently not available in the STRIDES database. HPV vaccination coverage is disproportionately lower in MS compared with the U.S. overall (49.5% in MS versus 71.5% in the U.S. overall receiving at least one dose in 2019); therefore, screening, and appropriate management of abnormal screening results remains critically important for cervical prevention efforts in the state [5]. Additionally, the study was conducted within the context of the Mississippi public health care population, therefore the findings of this study may not be directly generalizable to other regions/populations within and outside of the U.S.

Also, our study includes individuals screened from January 2018 to August 2021, with follow-up extending to September 2022, which includes the COVID pandemic period. MSDH clinic efforts were shifted to support the COVID-19 response throughout the state, making it more difficult to adequately follow-up on abnormal cytology and/or HPV results. However, 80% of the population was screened and managed before this time period and in stratified analyses, we observed only a slight decrement in follow-up during the height of the pandemic, with higher rates of adherence among those with high-grade cytology, consistent with prepandemic patterns.

Future implications

Mississippi has one of the highest screening rates, but the highest cervical cancer incidence in the nation. Our study reveals that screening status alone is not a sufficient predictor of successful cervical cancer prevention. There may be additional external factors influencing follow-up, and understanding the factors influencing this is critical to improve cervical cancer prevention. There are multiple social and structural barriers identified that disproportionately impact underserved populations, like the population seen in Mississippi's public health system [18, 19]. Further research is needed to explore these barriers and facilitators that impact individual follow-up to cervical cancer care in Mississippi to address identified disparities and inequalities [20, 21]. Additionally, understanding of the social and structural factors impacting adherence to follow-up recommendations can inform the development of future targeted interventions and patient-centered strategies. Availability and access to colposcopy clinics likely plays a major role: Screening is offered at many health clinics throughout the state, but fewer clinics have colposcopy equipment. Engaging patient navigators and leveraging community resources might be crucial in facilitating and supporting individuals in their follow-up care [22–25]. Further, educational campaigns and outreach initiatives may help raise awareness about the importance of timely colposcopy and help eliminate misconceptions that might prevent individuals from seeking appropriate care [26, 27]. In relation to the 18.48% (of the total n = 1,458) of individuals who returned for a repeat co-test instead of the recommended colposcopy, further research is necessary to understand patient's perspective.

In conclusion, our study among patients attending MSDH clinics with an abnormal cervical screening result found that 59% received the recommended follow-up colposcopy, while an additional 19% undergoing repeat screening instead of colposcopy. Overall, this leaves 22% of individuals with abnormal screening results lacking any type of follow-up care, and 41% without recommended colposcopy. Adherence to management guidelines is essential for accurate diagnosis and appropriate management of cervical abnormalities, which will ultimately contribute to the reduction of cervical cancer incidence and mortality. Identifying and addressing barriers to timely follow-up care will be crucial for reducing the burden of cervical cancer and improving health outcomes in underserved populations, like Mississippi. Our findings call for collaborative efforts among healthcare providers, policymakers, and communities to implement targeted interventions to bridge the gaps in cervical cancer care and strive for better health equity and improved patient outcomes.

Supplementary Information The online version contains supplementary material available at https://doi.org/10.1007/s10552-024-01905-0.

Acknowledgements The authors would like to thank previous and current collaborators on the STRIDES work including: Bethany Sabins, NP, Laura Tucker, MSN, RN, and Thomas Dobbs, MD, Lei Zhang, PhD, Laree Hiser, PhD, and Ken Owens. Also, a special thank you to each health department nurse for their collaborative clinical site guidance.

Author contributions All authors contributed to the study conception and design. Material preparation, data collection and analysis were performed by Sydney Reaves, Katherine C. Hall, Mary W. Stewart, and Megan A. Clarke. The first draft of the manuscript was written by Sydney Reaves and all authors commented on previous versions of the manuscript. All authors read and approved the final manuscript.

Funding This project was funded by the Intramural Research Program of the National Cancer Institute including funding from the Cancer Cures Moonshot Initiative. Dr. Carolann Risley is supported by the National Institute of General Medical Sciences of the National Institutes of Health under Award Number U54GM115428. The content is solely the responsibility of the authors and does not necessarily represent the official views of the National Institutes of Health.

Data availability The datasets generated and analyzed during this study are not publicly available due to restriction to only investigators included on the study protocol decided by the Institutional Review Board of the University of Mississippi Medical Center. Summary data can be made available from the corresponding author upon request

Declarations

Competing interests The authors declare no competing interests.

Ethical approval The Institutional Review Boards at the University of Mississippi Medical Center and Mississippi State Department of Health approved this protocol (Number 2016–0189).

Consent to participate A HIPAA waiver of authorization and HIPAA waiver of informed consent were granted.

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References

- Yang DX, Soulos PR, Davis B, Gross CP, Yu JB (2018) Impact of widespread cervical cancer screening. Am J Clin Oncol 41(3):289–294. https://doi.org/10.1097/coc.00000000000264
- American Cancer Society. (2024). Cancer Facts & Figures 2024. https://www.cancer.org/content/dam/cancer-org/research/cancerfacts-and-statistics/annual-cancer-facts-and-figures/2024/2024cancer-facts-and-figures-acs.pdf
- Centers for Disease Control and Prevention. (n.d.). Cervical Cancer is Preventable. Centers for Disease Control and Prevention.
- Liang LA, Zeissig SR, Schauberger G, Merzweiler S, Radde K, Fischbeck S, Ikenberg H, Blettner M, Klug SJ (2022) Colposcopy non-attendance following an abnormal cervical cancer screening result: a prospective population-based cohort study. BMC Women's Health. https://doi.org/10.1186/s12905-022-01851-6
- Risley C, Stewart MW, Geisinger KR, Hiser LM, Morgan JC, Owens KJ, Ayyalasomayajula K, Rives RM, Jannela A, Grunes DE, Zhang L, Schiffman M, Wentzensen N, Clarke MA (2021) Strides—studying risk to improve disparities in cervical cancer in Mississippi—design and baseline results of a statewide cohort

study. Prev Med 153:106740. https://doi.org/10.1016/j.ypmed. 2021.106740

- Explore cervical cancer screening in Mississippi: AHR. America's Health Rankings. (n.d.). https://www.americashealthrankings.org/ explore/measures/cervical_cancer_screen_women/MS
- Management guidelines. ASCCP. (n.d.). https://www.asccp.org/ management-guidelines
- Wentzensen N, Massad LS, Mayeaux EJ Jr, Khan MJ, Waxman AG, Einstein MH, Conageski C, Schiffman MH, Gold MA, Apgar BS, Chelmow D, Choma KK, Darragh TM, Gage JC, Garcia FAR, Guido RS, Jeronimo JA, Liu A, Mathews CA, Mitchell MM, Moscicki AB, Novetsky AP, Papasozomenos T, Perkins RB, Silver MI, Smith KM, Stier EA, Tedeschi CA, Werner CL, Huh WK (2017) Evidence-based consensus recommendations for colposcopy practice for cervical cancer prevention in the United States. J Low Genit Tract Dis 21(4):216–222. https://doi.org/10.1097/ LGT.0000000000000322. (PMID: 28953109)
- Boitano T, Ketch P, Maier J, Nguyen C, Huh W, Straughn J, Scarinci I (2022) Increased disparities associated with black women and abnormal cervical cancer screening follow-up. Gynecol Oncol Rep. https://doi.org/10.1016/j.gore.2022.101041
- Martinez-Gutierrez J, Chima S, Boyd L, Sherwani A, Drosdowsky A, Karnchanachari N, Luong V, Reece JC, Emery J (2023) Failure to follow up abnormal test results associated with cervical cancer in primary and ambulatory care: a systematic review. BMC Cancer. https://doi.org/10.1186/s12885-023-11082-z
- Mississippi Primary Care Needs Assessment 2021 Mississippi state ... (n.d.). https://msdh.ms.gov/msdhsite/index.cfm/ 44,7357,112,63,pdf/Primary%20Care%20Needs%20Assessme nt.pdf
- Sharp L, Cotton S, Thornton A, Gray N, Cruickshank M, Whynes D, Duncan I, Hammond R, Smart L, Little J (2012) Who defaults from colposcopy? A multi-centre, population-based, prospective cohort study of predictors of non-attendance for follow-up among women with low-grade abnormal cervical cytology. Eur J Obstet Gynecol Reprod Biol 165(2):318–325. https://doi.org/10.1016/j. ejogrb.2012.08.001
- Hariprasad R, Mittal S, Basu P (2022) Role of colposcopy in the management of women with abnormal cytology. Cytojournal 14(19):40. https://doi.org/10.25259/CMAS_03_15_2021
- Clinical guidance: Recommended best practices for delivery of ... (n.d.-a). https://www.cancercareontario.ca/sites/ccocancercare/ files/guidelines/full/Colposcopy_Clinical_Guidance_Document. pdf
- Perkins RB, Adcock R, Benard V, Cuzick J, Waxman A, Howe J, Melkonian S, Gonzales J, Wiggins C, Wheeler CM, Joste NE, Kinney W, Wheeler CM, Wiggins C, Robertson M, McDonald R, Waxman A, Jenison S, Castle PE, English K (2021) Clinical follow-up practices after cervical cancer screening by co-testing: a population-based study of adherence to U.S. guideline recommendations. Prev Med 153:106770. https://doi.org/10.1016/j.ypmed. 2021.106770
- Egemen D, Cheung LC, Chen X, Demarco M, Perkins RB, Kinney W, Poitras N, Befano B, Locke A, Guido RS, Wiser AL, Gage JC, Katki HA, Wentzensen N, Castle PE, Schiffman M, Lorey TS (2020) Risk estimates supporting the 2019 ASCCP risk-based management consensus guidelines. J Low Genit Tract Dis 24(2):132–143. https://doi.org/10.1097/LGT.00000000000529
- Miller SM, Tagai EK, Wen KY, Lee M, Hui SA, Kurtz D, Scarpato J, Hernandez E (2017) Predictors of adherence to follow-up

recommendations after an abnormal pap smear among underserved inner-city women. Patient Educ Couns 100:1353–1359

- Carrillo T, Montealegre JR, Bracamontes CG, Scheurer ME, Follen M, Mulla ZD (2021) Predictors of timely diagnostic follow-up after an abnormal pap test among Hispanic women seeking care in El Paso Texas. BMC Womens Health 21:11
- Kulkarni A, Glynn S, Gamble CR, Shen MJ, Cantillo E, Frey MK, Holcomb KM, Safford MM, Chapman-Davis E (2023) Understanding perceived barriers to colposcopy follow-up among underserved women at an Urban teaching hospital: a qualitative study. J Low Genit Tract Dis 27(1):87–92. https://doi.org/10.1097/LGT. 0000000000000700
- Abera GB, Abebe SM, Worku AG (2022) Impact of health education intervention on demand of women for cervical cancer screening: a cluster-randomized controlled trial. Trials. https://doi.org/ 10.1186/s13063-022-06765-0
- Wilding S, Wighton S, West R, Conner M, O'Connor DB (2023) A randomised controlled trial of volitional and motivational interventions to improve cervical cancer screening uptake. Soc Sci Med 322:115800. https://doi.org/10.1016/j.socscimed.2023. 115800
- 22. Battaglia TA, Bak SM, Heeren T, Chen CA, Kalish R, Tringale S, Taylor JO, Lottero B, Egan AP et al (2012) Boston patient navigation research program: the impact of navigation on time to diagnostic resolution after abnormal cancer screening. Cancer Epidemiol Biomark Prev 21:1645–1654
- 23. Bensink ME, Ramsey SD, Battaglia T, Fiscella K, Hurd TC, McKoy JM, Patierno SR, Raich PC, Seiber E et al (2014) Costs and outcomes evaluation of patient navigation after abnormal cancer screening: evidence from the patient navigation research program. Cancer 120:570–578
- Dhoj Shrestha A, Gyawali B, Shrestha A, Shrestha S, Neupane D, Ghimire S, Campbell C, Kallestrup P (2022) Effect of a female community health volunteer-delivered intervention to increase cervical cancer screening uptake in Nepal: a cluster randomized controlled trial. Prev Med Rep 29:101948. https://doi.org/10. 1016/j.pmedr.2022.101948
- Paskett ED, Dudley D, Young GS, Bernardo BM, Wells KJ, Calhoun EA, Fiscella K, Patierno SR, Warren-Mears V et al (2016) Impact of patient navigation interventions on timely diagnostic follow up for abnormal cervical screening. J Women's Health (Larchmt) 25:15–21
- Makadzange EE, Peeters A, Joore MA, Kimman ML (2022) The effectiveness of health education interventions on cervical cancer prevention in Africa: a systematic review. Prev Med 164:107219. https://doi.org/10.1016/j.ypmed.2022.107219
- Mboineki JF, Wang P, Dhakal K, Getu MA, Chen C (2022) The effect of peer-led navigation approach as a form of task shifting in promoting cervical cancer screening knowledge, intention, and practices among Urban women in Tanzania: a randomized controlled trial. Cancer Control. https://doi.org/10.1177/1073274822 1089480

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