

Clinical epidemiology and outcomes of invasive pulmonary aspergillosis as a complication of respiratory viral infection in hospitalized patients.

Lily He, PharmD, MS (candidate); Bertha De Los Santos, PharmD, MS (candidate); Brian J. Barnes, PharmD, MS; Nicholas S. Britt, PharmD, MS, BCPS, BCIDP
University of Kansas School of Pharmacy (KUMC Health Informatics , KU Frontiers Clinical and Translational Science Institute)

INTRODUCTION

- Invasive pulmonary aspergillosis (IPA) infections are difficult to diagnose, treat, and are associated with a high risk of death
- IPA is previously associated with immunosuppressive treatments in those undergoing stem cell or solid organ transplantation
- IPA is recognized as a complication of severe influenza infection, although the association between IPA and other respiratory viral infections (RVI) is unclear

OBJECTIVES

- Investigate the risk of developing IPA following RVI caused by a variety of respiratory viral pathogens
- Identify associated risk factors
- Evaluate the impact of RVI-associated IPA on patient outcomes and mortality

METHODS

- In-progress study using a retrospective cohort design
- Evaluated inpatient admissions between September 30, 2018, and October 5, 2019
- Inclusion criteria:**
 - Patients 18 years old and above
 - Admission to the University of Kansas Hospital
 - Multiplex PCR positive for respiratory virus from a nasopharyngeal swab, nasal wash, or bronchoalveolar lavage specimen
- Exclusion criteria:**
 - Proven or probable IPA prior to RVI
 - Admission duration <24 hours
- Statistical analysis and time-to-death was evaluated using Kaplan-Meier curves, hazard ratios, and log-rank testing

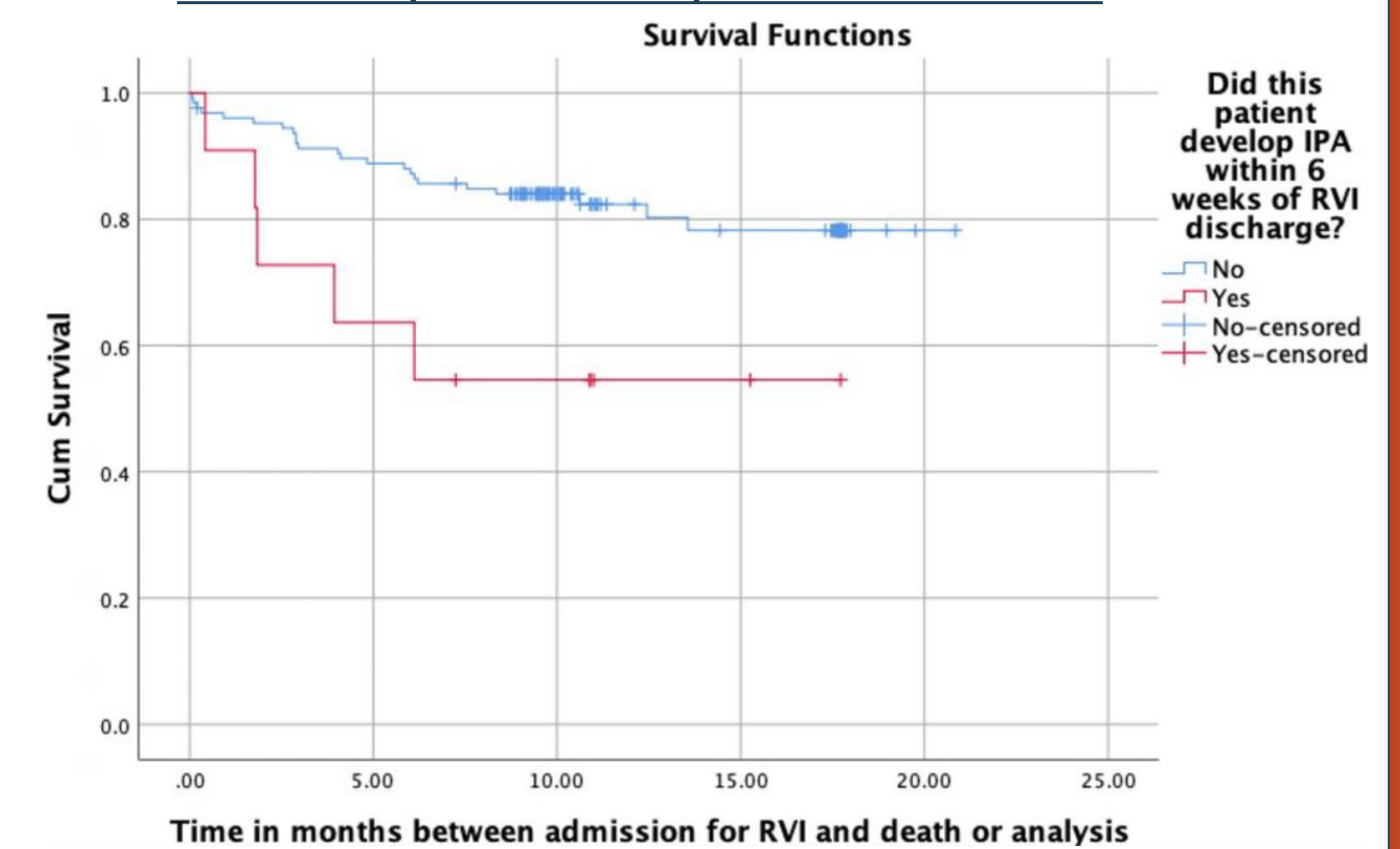
TABLE 1: BASELINE CHARACTERISTICS

Characteristic n (%) unless specified otherwise	IPA Group (n=11)	Non-IPA Group (n=126)	P Value
Age in years, mean (SD)	64 (± 11.6)	57 (± 18.1)	0.235
Female	3 (27.3)	65 (52.4)	0.128
Tobacco Use			
Current	1 (9.1)	23 (18.3)	0.905
Former	5 (45.5)	46 (36.5)	
Never	5 (45.5)	51 (40.5)	
Unknown	0 (0.0)	6 (4.8)	
Chronic Pulmonary Disease	5 (45.5)	54 (42.9)	> 0.999
Diabetes	1 (9.1)	37 (29.4)	0.29
Renal Disease (moderate-severe)	2 (18.2)	28 (22.2)	> 0.999
Liver Disease (moderate-severe)	0 (0.0)	10 (7.9)	> 0.999
Non-Metastatic Solid Tumor	0 (0.0)	10 (7.9)	> 0.999
Leukemia	6 (54.5)	10 (7.9)	< 0.001
Lymphoma/Multiple Myeloma	1 (9.1)	20 (15.9)	> 0.999
Metastatic Solid Tumor	0 (0.0)	4 (3.2)	> 0.999
AIDS	0 (0.0)	3 (2.4)	> 0.999
Charleston Index Score, median (IQR)	5 (3-5)	5 (3-6)	> 0.866
Prior Immunosuppressant Use	5 (45.5)	30 (23.8)	0.15
Prior Corticosteroid Use	10 (90.9)	34 (27.2)	< 0.001
Prophylactic Antibiotic Use	10 (90.9)	83 (66.4)	0.173
Prophylactic Antifungal Use	7 (63.6)	16 (12.7)	< 0.001
ID Consult	7 (63.6)	25 (20.3)	0.004
SOFA Score, median (IQR)	2 (0-3)	1 (0-3)	0.670
Total APACHE II Score, median (IQR)	16 (11-25)	12 (7-17)	0.154
ICU Admission	5 (45.5)	37 (30.1)	0.320

OUTCOME MEASUREMENTS

Outcome n (%)	IPA Group (n=11)	Non-IPA Group (n=126)	P Value
Inpatient Mortality	1 (9.1)	4 (3.2)	0.346
6-Week Mortality	1 (9.1)	6 (4.8)	0.451
12-Week Mortality	3 (27.3)	12 (9.5)	0.103

LOG-RANK (MANTEL-COX) MORTALITY CURVE



CONCLUSIONS

- IPA more commonly occurred in patients with non-influenza RVIs, suggesting that this complication is not limited to influenza
- There is an increased need for mechanical ventilation and increased mortality among those experiencing IPA following RVI
- Clinicians should be aware of the potential complications of severe RVI
- Future research into appropriate prophylaxis and treatment of post-RVI IPA is needed

ACKNOWLEDGMENT

We would like to recognize and thank Aletha Loeb, PharmD (candidate), Mallory Clay, PharmD (candidate), Haley Gandy, PharmD (candidate), and Xuan Gu, PharmD (candidate) for taking part in collecting and analyzing the data found for this study.

PREVIOUS RVI IN IPA GROUP

Of the 11 eligible IPA patients, 5 of them had a recent history of human rhinovirus or enterovirus and only 2 had a history of influenza (including H1N1 and Influenza A)

