

Impact of Total Body Weight on Efficacy of Ceftriaxone in Obese Patients

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BACKGROUND

- Obese patients are at an increased risk of infection and poor therapeutic outcomes compared to non-obese patients
- Pharmacokinetic parameters of β -lactam antibiotics are impacted by excessive adipose tissue and increased glomerular filtration rates
- Minimal data exist that assesses the relationship between ceftriaxone efficacy and obesity
- Purpose: To evaluate the clinical outcomes of ceftriaxone when used as definitive monotherapy for obese patients versus non-obese patients

METHODS

Study Design

- Single center, retrospective cohort; approved by the University of Mississippi Medical Center Institutional Review Board

Outcome Measures and Definitions

- **Composite Primary – Clinical Treatment Failure:**
 - Presence of any of the following:
 - Change in definitive therapy between 72 hours and 14 days post-initiation due to clinical worsening
 - Persistent leukocytosis (WBC > 10x10⁹/L) between 72 hours and 14 days after treatment initiation
 - Fever (single temperature >100.9 F) between 72 hours and 14 days post-treatment initiation
 - Re-infection within 30 days leading to readmission to the hospital
- **Secondary:**
 - 30-day inpatient all-cause mortality
 - 30-day hospital readmission

Subjects

- The study population included patients admitted to the University of Mississippi Medical Center from July 2015 to July 2017

Inclusion Criteria	Exclusion Criteria
<ul style="list-style-type: none"> • Age \geq 18 years old • Treated with ceftriaxone monotherapy for > 72 hours 	<ul style="list-style-type: none"> • Source control not achieved within 72 hours • Polymicrobial infections requiring > 1 antibiotic for definitive therapy

Analysis

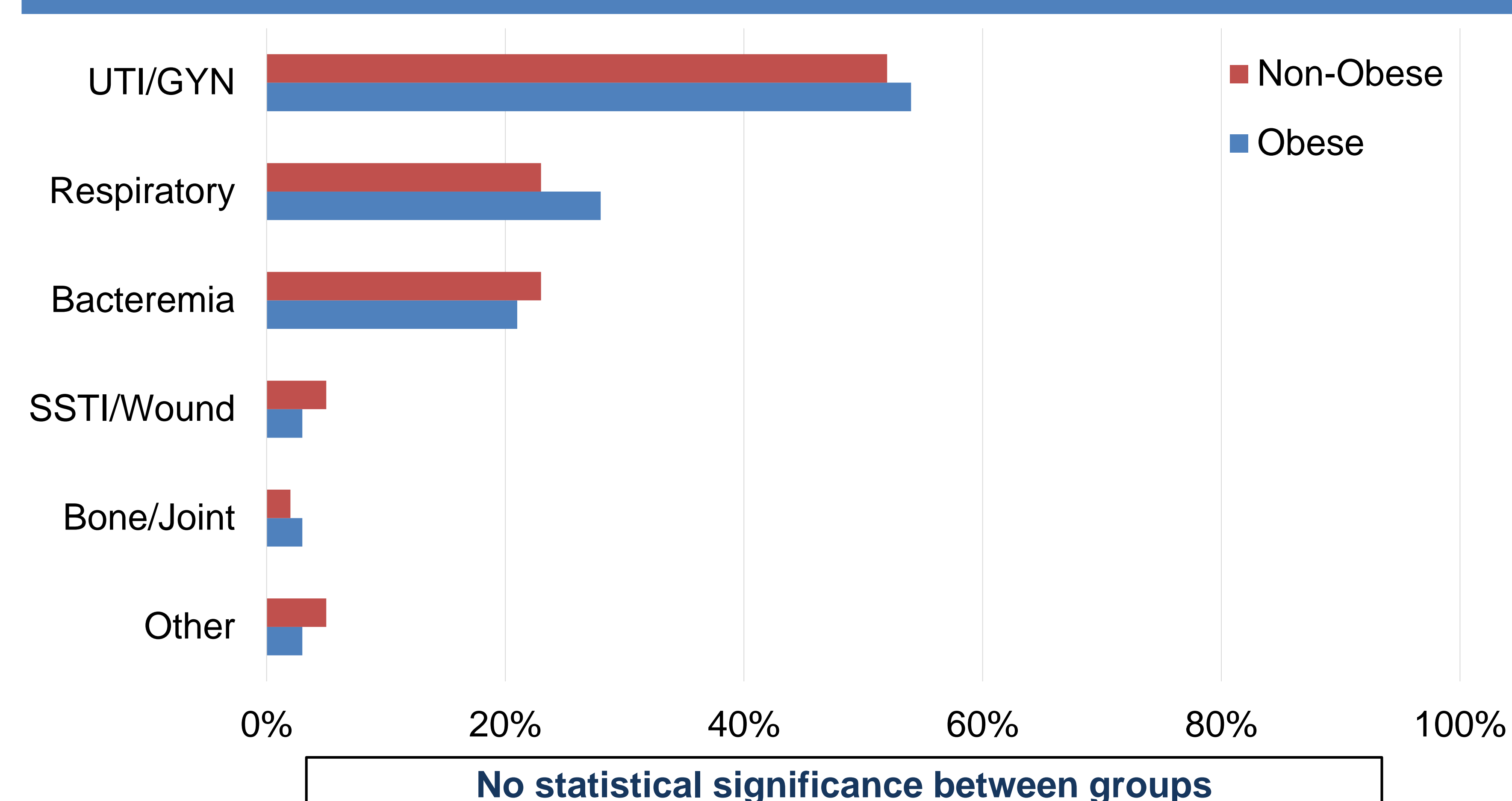
Results were reported as proportions or medians [interquartile range (IQR)]. Comparisons between the obese group and non-obese group were analyzed using Student's t-test or Mann-Whitney U test for continuous variables and X² test or Fisher's Exact test for categorical variables. A p-value of < 0.05 was considered statistically significant. Statistical analysis was performed using SPSS software version 24.0 (IBM).

RESULTS

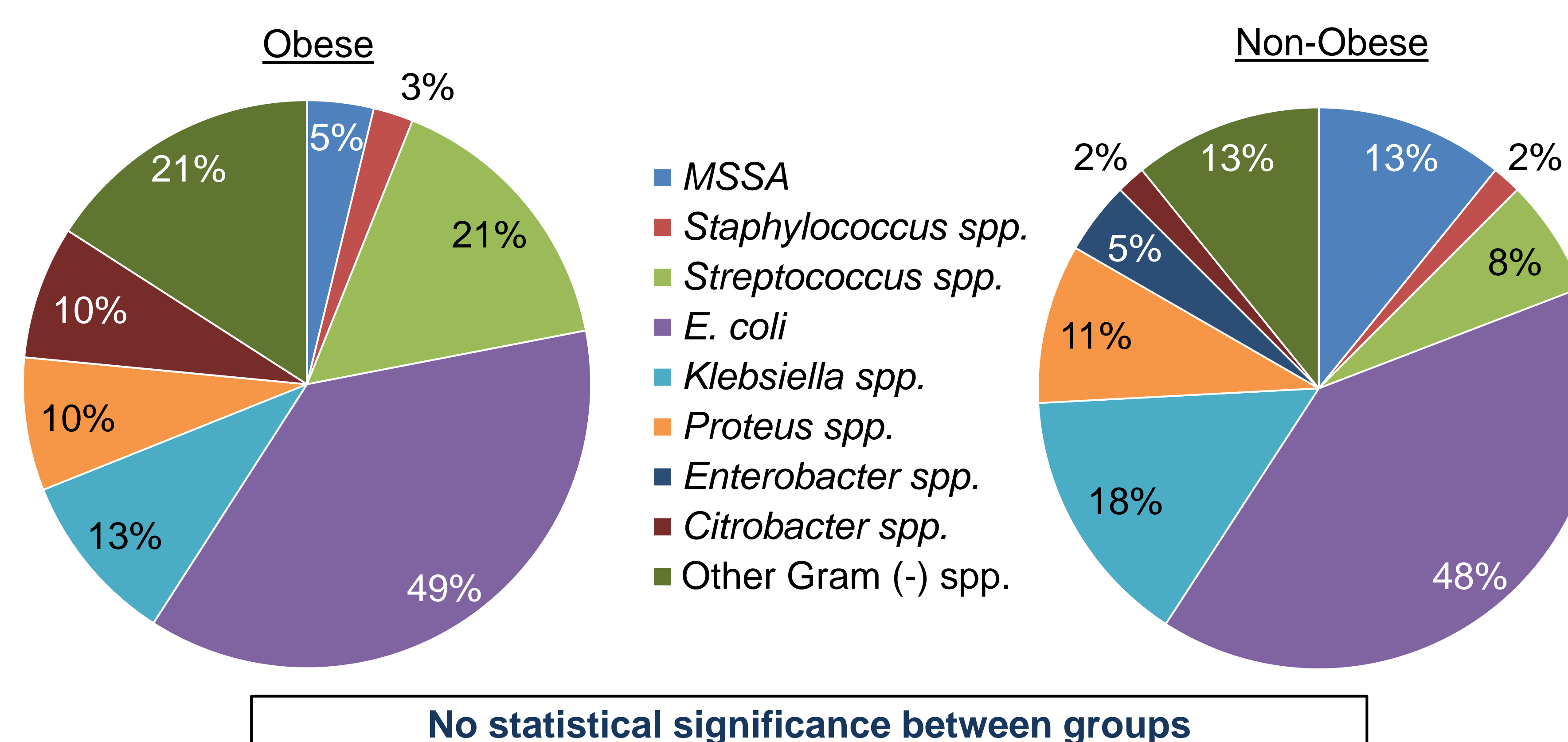
PATIENT DEMOGRAPHICS

Variable N(%) or Median [IQR]	Total (n=101)	Obese (n=39)	Non-Obese (n=62)	P-value
Age, years	62 [51-70]	62 [53-70]	62 [51-75]	0.761
Sex, male	56 (55.4)	24 (61.5)	32 (51.6)	0.329
Weight, kg	80 [64-98]	103 [95-120]	66 [59-77]	<0.001
Charlson Score	2 [1-4]	3 [1-5]	2 [1-4]	0.293

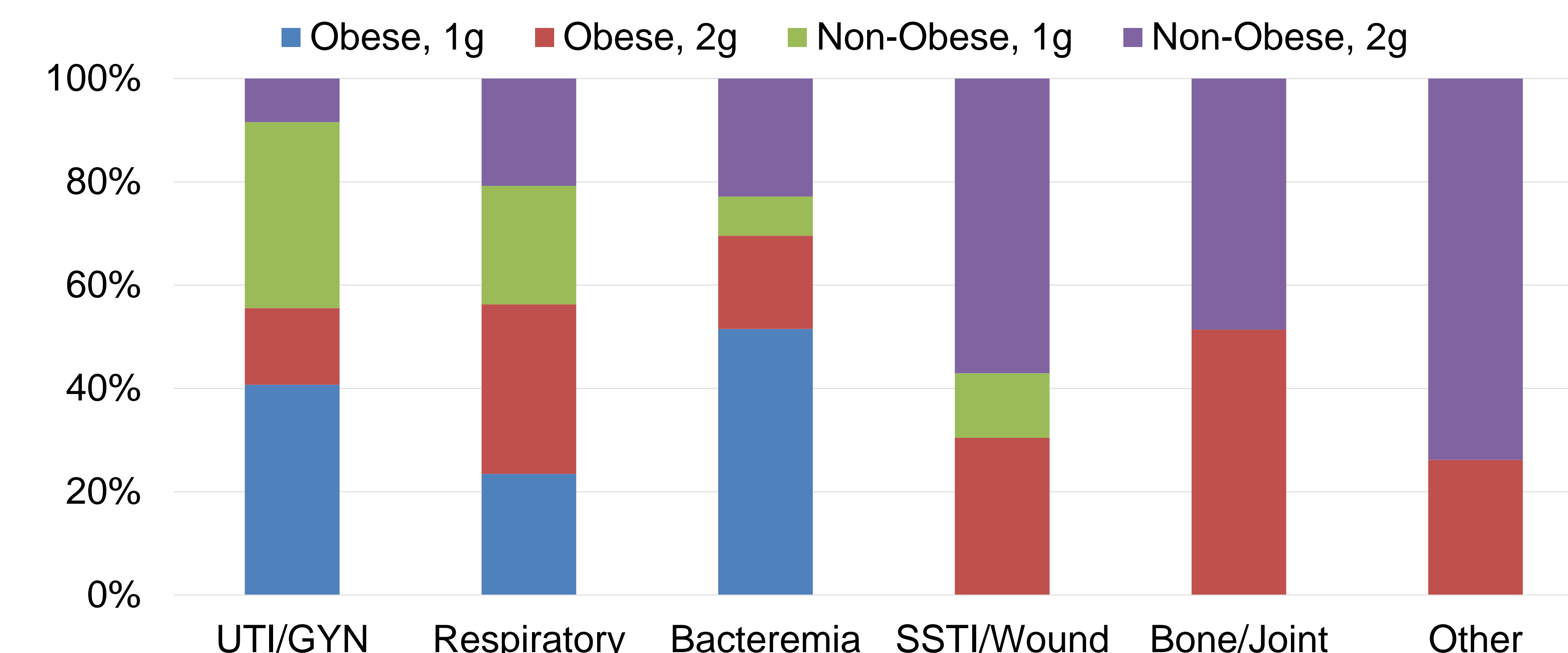
INFECTION SOURCE



MICROBIOLOGICAL RESULTS



CEFTRIAXONE DOSING



Obese patients more likely to receive 1g for UTI/GYN (p=0.014); non-obese patients more likely to receive 1g for UTI/GYN (p=0.010), 2g for bacteremia (p=0.028), and 1g for other infection sources (p=0.020).

COMPOSITE PRIMARY OUTCOME

Variable N (%)	Total (n=101)	Obese (n=39)	Non-Obese (n=62)	P-value
Overall Clinical Failure	49 (49)	24 (62)	25 (40)	0.038
Change in Therapy	26 (26)	14 (36)	12 (20)	0.064
Persistent Leukocytosis	39 (39)	21 (54)	18 (29)	0.013
Fever	12 (12)	6 (15)	6 (10)	0.529
30-day reinfection	11 (11)	3 (8)	8 (13)	0.523

SECONDARY OUTCOMES

Variable N (%)	Total (n=101)	Obese (n=39)	Non-Obese (n=62)	P-value
30-day readmission	18 (18)	6 (15)	12 (19)	0.791
30-day inpatient all-cause mortality	8 (8)	5 (13)	3 (5)	0.255

CONCLUSION

- Obese patients treated with ceftriaxone had higher rates of treatment failure compared with non-obese patients
- While not statistically significant, there was numerically higher inpatient mortality in obese patients compared with non-obese patients
- Further examination is needed to assess impact of ceftriaxone dose and organism MIC on clinical failure in obese patients