## Rates and Risk Factors for Multiple Periprosthetic Joint Infections in Patients with Multiple Arthroplasties

Kingsley A Oladeji, Jonathan J Lee<sup>1</sup>, Shay Ivan Warren, Brian Francis Sweeney, Tatenda L Chakoma, Prerna Arora<sup>2</sup>, Andrea Finlay, Derek F Amanatullah<sup>2</sup>

<sup>1</sup>Stanford University Department of Orthopedic Surge, <sup>2</sup>Stanford University

INTRODUCTION: The rate for periprosthetic joint infection (PJI) exceeds 1% of all primary arthroplasties. Over 30% of patients with a primary total hip or knee arthroplasty end up with more than one arthroplasty. Given that patients with multiple arthroplasties incur the risk for PJI in each operated joint, we hypothesized that the rate in subsequent arthroplasty is high. Our objective was to determine the rates and risk factors for synchronous and metachronous PJI in subsequent prostheses in this population.

METHODS: Patients with multiple prostheses with more than one PJI were categorized as synchronous (i.e., presenting at the same time as the initial infection) or metachronous (i.e., presenting at a different time as the initial infection). Differences were assessed using a chi-square test for categorical variables and an t-test for continuous variables. Odds ratios (OR) and confidence intervals (CI) were calculated using logistic regression. P < 0.05 was considered statistically significant.

RESULTS: A total of 337 patients with a PJI in at least one joint were reviewed from 2003 to 2021. Demographics and risk factors were collected at a mean follow up of  $2.9 \pm 2.9$  years (range: 0.0 - 17.2 years). Thirty-one (9%) patients developed a PJI in more than one joint. Patients with multiple PJI were significantly more likely to have a positive blood culture (48% vs. 16%, p < 0.001) and longer follow-up times than patients with single PJI (4.8 years vs. 2.7 years, p < 0.001). The odds of developing a second PJI in the contralateral limb (n = 27, 87%) were elevated by almost 7-fold (OR: 6.8, CI: 2.1-21.5, p = 0.001) when compared to a second PJI in the ipsilateral limb (n = 4, 13%, Figure 1). There was no difference for the odds of a synchronous (n = 13, 4%) versus metachronous (n = 18, 5%) presentation (OR: 0.72, CI: 0.30-1.72, p = 0.463, Table 1). Tobacco users were 75% more likely to have a metachronous presentation of their second PJI compared to patients without tobacco use (OR: 1.75, CI: 1.1-2.9, p = 0.041, Table 2).

DISCUSSION AND CONCLUSION: Almost 10% of patients with an initial PJI will develop a second PJI in a subsequent arthroplasty with the contralateral limb being at particular risk. Caution should be taken in patients with a history of tobacco use and prior PJI as the relative risk of a subsequent PJI is high. Given that many metachronous PJI patients presented years later with the same bacterial culture, the results point toward altered host immunity after PJI as a potential mechanism. Optimizing the management of patients with a prior PJI is essential to reducing morbidity and additional PJI-related healthcare spending in this high-risk group.

15			PJI Type Synchronous PJI Metachronous PJI
0 -	Contralateral	Ipsilateral	

Risk factors	Unadjusted			Adjusted				Sync
	OR	95% CI	p-value	AOR	95% CI	p-value	Risk factors	N (%
lader survey				-	-		Index surgery	
Primary	Ref						Primary	6 (46
Beniden	0.07	0.011.03	0.162	0.00	0.021.07	A (20	Revision	7 (53
Revision	0.97	0.91-1.03	0.349	0.98	0.92-1.03	0.000	Total joint replaced	10.0
Total joint replaced							Khee	10(7
Knee	Ref						Hip Newber of inists with BU	3 (23
Hip	0.95	0.89-1.01	0.075	0.95	0.89-1.01	0.087	Number of Joints wan FM	0.000
Location of initial PJI							2+	13.0
Left knee	Ref						Location of initial PII	1.00
Right knee	0.98	0.90-1.06	0.581	0.97	0.89-1.05	0.481	Left knee	7 (53
Left hip	0.94	0.86-1.03	0.200	1.06	0.85-1.32	0.603	Right knee	2 (15
Right his	0.94	0.82-1.03	0.184	1.62	0.86(1.33	0.561	Left hip	3 (23
Sex	0.01			1,67			Right hip	1 (7.5
Pool.	3.6						Sex	-
remaie	Rei						Female	6 (46
Male	0.96	0.86-1.06	0.423	0.98	0.99-1.67	0.460	Male	7 (53.
Race							Race	
Non-White	0.99	0.92-1.07	0.775	0.97	0.89-1.06	0.515	Asian	0 (0.0
White	Ref						Black	0 (0.0
Dinicity							Native American	0 (0.0
Hispanie/Latino or Unknown	1.01	0.92-1.12	0.809	1.02	0.91-1.15	0.719	Other	3 (23
Non-Hispanic/ Non-Latino	Ref						Pacific Islander	0 (0.0
Ans at time of first PII	1.00	1.00-1.00	0.423	1.60	1.00-1.02	0.292	White	10.0
DMI	1.00	1.00.1.01	0.767	1.00	140.1.03	0.950	Ethnicity	10()
0.1	0.00	0.00-1.01	0.107	1.00	1.40-1.00	0.900	HistoricTatino	2 (15
Charleston Comorbiality Index	0.99	0.97-1.01	0.401	0.99	0.97-1.01	0.236	Non-Hispanic/ Non-	11 (8
ASA							Unknown	0 (0.0
182	Ref						Age at time of first PJI	68.18
384	1.05	0.99-1.12	0.108	1.05	0.98-1.12	0.183	BMI	28.25
Smoker							Charleston Comorbidity	3.08
No	Ref						ASA	<u> </u>
Current or former	1.01	0.92-1.12	0.770	1.02	0.92-1.12	0.748	1&2	3 (25.
Diabetes	-						3&4	10 (7:
No	Ref						Smoker	
Var	1.02	0.661111	0.505	1.01	0.041.00	0.366	No	13 (14
	1.00	0.954.11	0.505	1.01	0.941.09	0.750	Current or former	0 (0.0
Kheumatoud arthritis							Dubetes	0.00
No	Rel						No	9 (09
Yes	1.02	0.93-1.10	0.717	1.01	0.93-1.10	0.745	Phermatoid authritic	4 (30
Bacteria-positive blood culture							No	12/9
No	Ref						Ves	11/73
		-		1				1.0.
Yes	1.20	1.11+1.29	< 0.001	1.20	1.11+1.59	< 0.001	Bacteria-positive blood	
Yes Follow-up time (years)	1.20	1.11-1.29	< 0.001	1.02	1.01-1.03	< 0.001	Bacteria-positive blood	5 (38)

	Synchronous PJI	Metachronous PJI	
factors	N (%) or M (SD)	N (%) or M (SD)	Fisher or t-test p-value
X SUDNEY			1.00
rimary	6 (46.2%)	8 (44.4%)	-
rvision	7 (53.8%)	10 (55.6%)	-
l joint replaced			0.452
nee	10 (76.9%)	11 (61.1%)	
ip	3 (23.1%)	7 (38.9%)	-
ber of joints with PJI			
	0 (0.0%)	0 (0.0%)	-
+	13 (100.%)	18 (100.0%)	-
ation of initial PJI		,	0.151
eft knee	7 (53,856)	4 (22.2%)	
jeht knee	2 (15.4%)	7 (38,9%)	
eft hin	3 (23.1%)	2 (11.150)	
icht hin	1 (7.2%)	5 (27.8%)	
······		- (	0.794
emale	6 (46 2%)	12 (66 7%)	
Tale .	7(53,850)	6(33.3%)	
	. fearerey	· (*******	0.676
-	0.00000	0.00.0%)	*****
lack	0 (0.0%)	0 (0.0%)	
ation American	0(0.0%)	0(0.0%)	
that	3/21160	3716780	
noifis Islandar	0 (0.0%)	0(0000)	-
nknown	0 (0.0%)	0 (0.0%)	
/hite	10(76,9%)	15 (83 355)	
urity.	10(10576)	10 (001074)	1.00
imparied ating	2 (15.450)	2 (11 150)	1100
on-Dimanic/Non-	11 (84 (85)	16 (88 9%)	
nknown	0.00%)	0.00.0%)	_
at time of first Pll	68.18 (8.96)	65 (90 / 9 (95)	0.150
	28.29 (7.20)	33.07 (9.58)	0.145
rieston Comerhidity	3.08 (1.50)	2 22 (1 53)	0.525
			1.00
67	3/25/660	3/158%)	
84	10(75(85))	13 (23 255)	
iker.	to (crossil)	10 (10 / 0)	0.120
0	13 (10) (50)	14 (27.8%)	*****
o urrent or former	0.00%	4 (22 250)	
where	0 (0.074)	- (ama/4)	0.689
0.000	9 (69 750)	14 (77 8%)	1.007
	4(30.8%)	4/22/260	
es antipitie	+ (2000 AU)	- (aa.a/9)	010
amanova andifilis	12 (92 28))	11 (22.26)	v
	1.02.280	6/17 (14-479)	
we tasks monitive blood	10.070	2 (ar /8/8)	0.98
nam-possave bicos	6 (19 61()	11 (61 197)	9.687
	9 (61 66/)	11 (01.179)	
to	3 (01.270)	r (26,7%)	0.012
ow-up time (years)	3710 (3.37)	3.74 (4.30)	144007