Supplemental Online Content

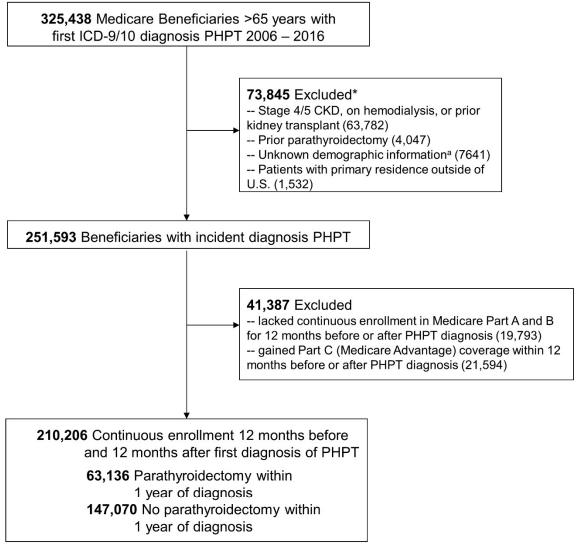
Seib CD, Meng T, Suh I, et al. Risk of fracture among older adults with primary hyperparathyroidism receiving parathyroidectomy vs nonoperative management. *JAMA Intern Med.* Published online November 29, 2021. doi:10.1001/jamainternmed.2021.6437

eFigure 1. Cohort enrollment criteria

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This supplemental material has been provided by the authors to give readers additional information about their work.

eFigure 1. Cohort Enrollment Criteria



^{*} The sum of subcategory totals exceeds the number of total exclusions because some patients met multiple exclusion criteria-

^a Patients were excluded if unknown age, sex, race/ethnicity, area deprivation index, or urban/rural inhabitance.

eTable 1. Adjusted hazard of any fracture and hip fracture based on inverse-probability weighted Fine-Gray competing risk regression accounting for the competing risk of death among Medicare beneficiaries with PHPT.

	ı	Any Fracture		Hip Fracture Fine-Gray Competing Risk Regression				
		ay Competin Regression	g Risk					
Characteristic	IPW Adjusted HR	95% CI		95% CI Adjusted HR		% CI		
Treatment								
Non-operative management	Ref			Ref				
Parathyroidectomy	0.84	0.82	0.85	0.83	0.80	0.85		
Sex								
Male	Ref			Ref				
Female	1.49	1.45 1.54		1.37	1.30	1.44		
Age group								
66 to 75	Ref			Ref				
76 to 85	1.55	1.52	1.58	2.16	2.08	2.24		
86 and above	1.85	1.80	1.91	3.01	2.86	3.16		
Race								
White	Ref			Ref				
Black	0.44	0.42	0.46	0.42	0.39	0.46		
Asian	0.73	0.66	0.81	0.77	0.64	0.92		
Hispanic	0.93	0.86	1.01	0.83	0.71	0.97		
American Native	0.98	0.82	1.18	0.78	0.54	1.12		
Unknown	0.63	0.56	0.70	0.54	0.44	0.68		
ADI Group								
Most advantaged	Ref			Ref				

Slightly advantaged	1.01	0.99	1.04	1.07	1.02	1.11
Slightly disadvantaged	1.01	0.98	1.04	1.10	1.05	1.15
Most disadvantaged	1.04	1.01	1.08	1.22	1.15	1.30
Geography						
Urban	Ref			Ref		
Rural	1.02	1.00	1.53	1.05	0.99	1.08
Frailty						
Robust	Ref			Ref		
Prefrail	1.34	1.31	1.37	1.22	1.17	1.27
Mildly frail	1.65	1.60	1.71	1.53	1.44	1.62
Moderately-to-severely frail	1.42	1.36	1.49	1.39	1.29	1.50
Charlson Comorbidity Index Score						
0	Ref			Ref		
1	1.08	1.05	1.11	1.05	1.00	1.10
≥2	1.09	1.07	1.12	1.04	0.99	1.09
Operative indications for parathyroidectomy						
History of osteoporosis	1.67	1.64	1.71	1.41	1.36	1.46
History of kidney stones	0.96	0.93	0.98	0.92	0.88	0.97
History of stage 3 CKD	0.88	0.85	0.91	1.01	0.96	1.07
Endocrinologist care within 6 months of diagnosis	0.98	0.96	1.00	0.93	0.90	0.96
Fracture Risk Factors						
Prior fracture	2.20	2.15	2.25	1.47	1.41	1.54
Tobacco use	1.27	1.22	1.33	1.59	1.48	1.70
Alcohol use disorder	1.44	1.32	1.57	1.44	1.23	1.69
Obesity	0.93	0.90	0.96	0.75	0.70	0.80

^{*} The propensity score used for inverse probability weighting (IPW) in this table was based on all characteristics listed in this table. Abbreviations: IPW, inverse probability weighted; ADI, area deprivation index; CKD, chronic kidney disease.

eTable 2. Baseline characteristics of subgroup of older adults diagnosed with PHPT from 2006 – 2017 who have continuous enrollment in Medicare Part D (prescription) coverage for 12 months prior to diagnosis.

	Unadjusted								IPW-adjusted ^a					
	Overall cohort (n=210,206)		Parathyroidect omy (n=63,136)		Non- operatively managed (n=147,070)			Overall cohort (n=210, 206)	Parathyroidect omy (n=63,136)	Non- operatively managed (n=147,070)				
Characteristic	No.	%	No.	%	No.	%	SMD	%	%	%	SMD			
Sex														
Female	89,489	80.9%	25,917	80.6%	63,572	81.1%	0.013	81.1%	81.4%	81.0%	-0.008			
Male	21,063	19.1%	6,246	19.4%	14,817	18.9%	-0.013	18.9%	18.6%	19.0%	0.008			
Age group														
66 to 75	62,744	56.8%	21,740	67.6%	41,004	52.3%	-0.316	56.6%	56.5%	56.7%	0.004			
76 to 85	37,788	34.2%	9,396	29.2%	28,392	36.2%	0.150	34.4%	34.6%	34.2%	-0.008			
86 and above	10,020	9.1%	1,027	3.2%	8,993	11.5%	0.322	9.0%	8.9%	9.1%	0.006			
Race														
White	95,761	86.6%	28,998	90.2%	66,763	85.2%	-0.152	86.6%	86.5%	86.6%	0.003			
Black	10,012	9.1%	2,225	6.9%	7,787	9.9%	0.109	9.1%	9.2%	9.1%	-0.005			
Asian	1,388	1.3%	249	0.8%	1,139	1.5%	0.065	1.3%	1.3%	1.3%	-0.003			
Hispanic	1,783	1.6%	349	1.1%	1,434	1.8%	0.062	1.6%	1.6%	1.6%	0.001			
American Native	220	0.2%	74	0.2%	146	0.2%	-0.010	0.2%	0.2%	0.2%	-0.001			
Unknown	1,388	1.3%	268	0.8%	1,120	1.4%	0.056	1.2%	1.2%	1.3%	0.005			
ADI Group														
Most advantaged	28,396	25.7%	6,708	20.9%	21,688	27.7%	0.159	25.6%	25.5%	25.7%	0.003			
Slightly advantaged	35,104	31.8%	10,202	31.7%	24,902	31.8%	0.001	31.6%	31.5%	31.7%	0.005			
Slightly disadvantaged	34,361	31.1%	11,215	34.9%	23,146	29.5%	-0.115	31.2%	31.4%	31.1%	-0.005			
Most disadvantaged	12,691	11.5%	4,038	12.6%	8,653	11.0%	-0.047	11.5%	11.6%	11.5%	-0.004			

Geography											
Urban	93,187	84.3%	26,029	80.9%	67,158	85.7%	0.127	84.2%	84.1%	84.3%	0.004
Rural	17,365	15.7%	6,134	19.1%	11,231	14.3%	-0.127	15.8%	15.9%	15.7%	-0.004
Frailty											
Robust	35,777	32.4%	11,859	36.9%	23,918	30.5%	-0.135	32.4%	32.6%	32.4%	-0.004
Prefrail	57,976	52.4%	17,039	53.0%	40,937	52.2%	-0.015	52.5%	52.6%	52.4%	-0.003
Mildly frail	11,483	10.4%	2,248	7.0%	9,235	11.8%	0.165	10.3%	10.2%	10.4%	0.007
Moderately-to- severely frail	5,316	4.8%	1,017	3.2%	4,299	5.5%	0.114	4.8%	4.7%	4.8%	0.004
Charlson Comorbidity Index Score											
0	23,753	21.5%	7,991	24.8%	15,762	20.1%	-0.114	21.6%	21.6%	21.5%	-0.002
1	23,664	21.4%	7,230	22.5%	16,434	21.0%	-0.037	21.5%	21.6%	21.4%	-0.004
≥2	63,135	57.1%	16,942	52.7%	46,193	58.9%	0.126	57.0%	56.8%	57.0%	0.005
Operative indications for parathyroidecto my											
History of osteoporosis											
No history	55,699	50.4%	16,772	52.1%	38,927	49.7%	-0.050	50.5%	50.4%	50.5%	-0.002
History based on claim diagnosis	27,180	24.6%	7,696	23.9%	19,484	24.9%	0.022	24.6%	24.2%	24.7%	-0.014
On pharmacologic treatment for osteoporosis	27,673	25.0%	7,695	23.9%	19,978	25.5%	0.036	25.0%	25.4%	24.8%	0.015
History of kidney stones	17,251	15.6%	5,985	18.6%	11,266	14.4%	-0.114	15.5%	15.3%	15.5%	0.006
History of stage 3 CKD	11,568	10.5%	2,420	7.5%	9,148	11.7%	0.141	10.4%	10.3%	10.5%	0.005
Endocrinologist care within 6	51,523	46.6%	18,367	57.1%	33,156	42.3%	-0.299	46.7%	47.0%	46.6%	-0.007

months of diagnosis											
Fracture Risk Factors											
Prior fracture	10,996	9.9%	2,535	7.9%	8,461	10.8%	0.100	10.0%	10.0%	10.0%	-0.002
Tobacco use	5,305	4.8%	1,777	5.5%	3,528	4.5%	-0.047	4.8%	4.8%	4.8%	0.002
Alcohol abuse disorder	933	0.8%	223	0.7%	710	0.9%	0.024	0.9%	0.9%	0.8%	-0.002
Obesity	14,396	13.0%	4,343	13.5%	10,053	12.8%	-0.020	13.0%	12.9%	13.0%	0.004
Pharmacologic Treatment											
Chronic steroid use	1,408	1.3%	306	1.0%	1,102	1.4%	0.021	1.3%	1.3%	1.3%	0.000
Cinacalcet	2,757	2.5%	730	2.3%	2,027	2.6%	0.042	2.6%	2.6%	2.5%	-0.008

^a The propensity score used for inverse probability weighting in this table was based on all characteristics listed in this table.

Abbreviations: IPW, inverse probability weighted; ADI, area deprivation index; CKD, chronic kidney disease.

eTable 3. Unadjusted incidence rate and adjusted hazard of any fracture and hip fracture based on inverse-probability weighted Cox proportional hazard regression among subgroup of Medicare beneficiaries with PHPT who have continuous enrollment in Medicare Part D (prescription) coverage for 12 months prior to diagnosis.

	Α	ny Fracture)	Hip Fracture					
	Unadjusted incidence rate of any fracture	Cox Proportional Hazards Regression			Unadjusted incidence rate of hip fracture	Cox Proportional Hazards Regression			
Characteristic	Rate (per 100 person- years)	IPW Adjusted HR	95%	6 CI	Rate (per 100 person- years)	IPW Adjust ed HR	959	% CI	
Treatment									
Non-operative management	3.18	Ref			0.89	Ref			
Parathyroidect omy	2.05	0.77	0.74	0.80	0.51	0.73	0.69	0.78	
Sex									
Male	1.57	Ref			0.47	Ref			
Female	3.12	1.44	1.35	1.53	0.84	1.43	1.28	1.61	
Age group									
66 to 75	1.80	Ref			0.38	Ref			
76 to 85	3.87	1.67	1.60	1.74	1.14	2.47	2.28	2.68	
86 and above	6.57	2.49	2.36	2.63	2.23	4.33	3.91	4.80	
Race									
White	3.01	Ref			0.82	Ref			
Black	1.25	0.45	0.41	0.50	0.41	0.50	0.42	0.58	
Asian	2.20	0.73	0.62	0.87	0.59	0.73	0.53	1.02	
Hispanic	2.97	0.91	0.80	1.04	0.79	0.80	0.62	1.03	
American Native	3.30	1.38	0.98	1.96	0.53	1.05	0.51	2.14	
Unknown	1.80	0.62	0.51	0.77	0.51	0.74	0.51	1.08	
ADI Group									
Most advantaged	2.83	Ref			0.73	Ref			
Slightly advantaged	2.83	1.05	1.00	1.10	0.78	1.20	1.10	1.32	
Slightly disadvantage d	2.85	1.05	1.00	1.11	0.79	1.23	1.12	1.35	
Most disadvantage d	2.76	1.13	1.06	1.21	0.82	1.43	1.27	1.62	
Geography									
Urban	2.83	Ref			0.77	Ref			
Rural	2.84	1.04	0.99	1.09	0.78	0.97	0.88	1.07	

			_					
Frailty								
Robust	1.65	Ref			0.43	Ref		
Prefrail	3.02	1.42	1.36	1.50	0.83	1.44	1.32	1.58
Mildly frail	5.30	2.05	1.92	2.19	1.49	2.27	2.01	2.56
Moderately-								
to-severely	5.38	1.85	1.70	2.01	1.44	1.83	1.57	2.14
frail								
Charlson Comorbidity								
Index Score								
0	2.10	Ref			0.59	Ref		
1	2.66	1.09	1.03	1.16	0.74	1.06	0.95	1.19
≥2	3.20	1.14	1.08	1.20	0.86	1.05	0.95	1.16
Operative	3.23		11111		0.00		0.00	
indications for								
parathyroidect								
omy								
History of								
osteoporosis								
No osteoporosis	1.66	Ref			0.51	Ref		
Diagnosis code								
only	3.89	1.55	1.48	1.63	1.09	1.38	1.27	1.50
On								
pharmacologic	4.11	1.68	1.61	1.76	0.97	1.19	1.09	1.30
treatment for	4.11	1.00	1.01	1.70	0.97	1.19	1.09	1.50
osteoporosis								
History of	2.73	0.97	0.92	1.02	0.70	0.92	0.84	1.02
kidney stones								
History of stage 3 CKD	3.04	0.95	0.89	1.01	0.92	1.17	1.05	1.30
Endocrinologi								
st care within	0.77	0.00	0.04	4.04	0.70	0.04	0.07	4.00
6 months of	2.77	0.98	0.94	1.01	0.72	0.94	0.87	1.00
diagnosis								
Fracture Risk								
Factors								
Prior fracture	8.82	2.36	2.26	2.46	1.83	1.67	1.53	1.82
Tobacco use	3.63	1.43	1.32	1.55	1.07	1.70	1.47	1.97
Alcohol use	4.95	1.44	1.20	1.72	1.39	1.51	1.08	2.12
disorder	7.30	1.44	1.20	1.72		1.01	1.00	۷. ۱۷
Obesity	2.36	0.99	0.93	1.05	0.46	0.74	0.64	0.84
Pharmacologi								
c Treatment								
Chronic steroid use	6.19	1.70	1.51	1.92	1.03	1.25	0.97	1.62
Cinacalcet	3.66	0.90	0.81	1.01	1.21	1.03	0.86	1.25
2.1.4.54.561	0.00	0.00	5.5 .				0.00	

^{*} The propensity score used for inverse probability weighting in this table was based on all characteristics listed in this table.

Abbreviations: IPW, inverse probability weighted; ADI, area deprivation index; CKD, chronic kidney disease.