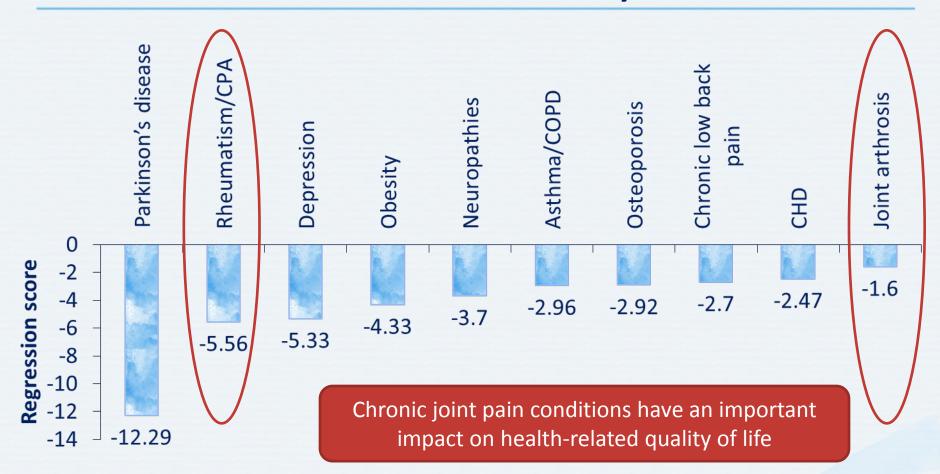
BURDEN OF ILLNESS

Overview

Impact of Chronic Conditions on Health-Related Quality of Life



Note: a larger negative score indicates a greater impact on health-related quality of life CHD = coronary heart disease; COPD = chronic obstructive pulmonary disease; CPA = chronic polyarthritis Brettschneider C et al. PLoS One 2013; 8(6):e66742.

Physical Burden

Ankylosing Spondylitis Negatively Impacts Quality of Life

- Most common concerns:
 - Stiffness
 - Pain
 - Fatigue
 - Poor sleep
 - Appearance
 - Worry about the future
 - Medication side effects
- Ankylosing spondylitis patients have:
 - Higher well-being and lower direct cost of illness than patients with fibromyalgia or chronic low back pain
 - Higher mortality rate than the general population

Rheumatoid Arthritis Negatively Impacts Daily Functioning

- Rheumatoid arthritis causes a functional burden by limiting ability to perform daily activities:¹
 - Self-care (e.g., dressing, feeding, bathing, grooming and toileting)
 - Vocational (e.g., work, school and homemaking)
 - Avocational (e.g., exercise, recreation and leisure)
- Long-term studies have found that 50% of patients with rheumatoid arthritis have had to stop working 10 years after diagnosis²

Rheumatoid arthritis results in functional burden to the patient and to caregivers

Hand Rheumatoid Arthritis Can Have Severe Impact





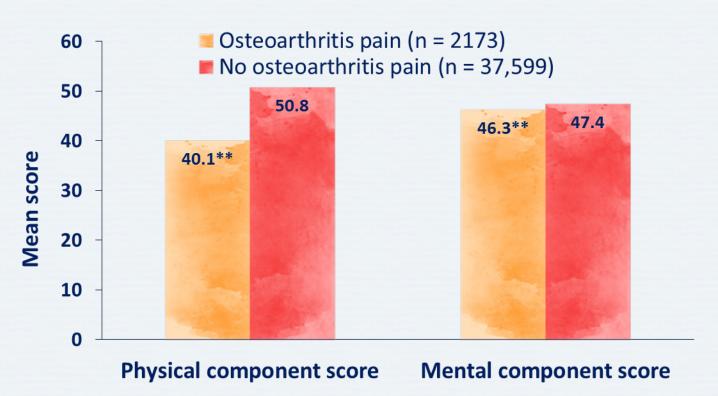
Osteoarthritis Negatively Impacts Daily Functioning

- Osteoarthritis causes a functional burden by limiting ability to perform daily activities, such as dressing, feeding, bathing, grooming and toileting¹
- 80% of osteoarthritis patients report physical limitations due to disease²

Osteoarthritis results in functional burden to the patient and to caregivers

Osteoarthritis Reduces Quality of Life

Physical and Mental Component Scores on the SF-12v2*



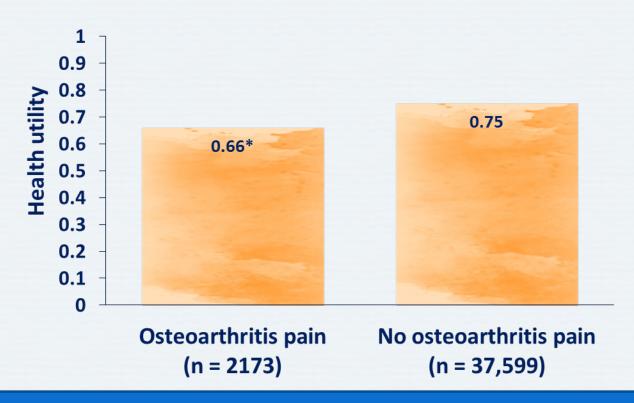
Workers with osteoarthritis pain have lower quality of life scores

SD = standard deviation; SF-12v2 = Short Form 12 version 2; US = United States

^{*}Range of 0–100, higher scores indicate better health, adjusted for covariates and normed to the US population (mean = 50, SD = 10) **p < 0.0001 vs. the comparator cohort.

Osteoarthritis Reduces Quality of Life

Health Utility Score on the SF-6D[†]



Workers with osteoarthritis pain have lower quality of life scores

Economic Burden

Health Care Costs of Ankylosing Spondylitis Are Significant*

- 241 patients enrolled in the Longitudinal Study of Outcomes in Ankylosing Spondylitis
- Indirect costs dominated the total costs associated with ankylosing spondylitis

Annual cost/patient	Amount (1999 USD)							
Annual cost/patient	Mean	Median	Range	% of total cost				
Direct	1775	1113	0–36,267	26.4				
Indirect	4945	0	0-45,800	73.6				
Total	6720	1495	0-80,017	100				

Economic Burden of Ankylosing Spondylitis

- 234 individuals* with ankylosing spondylitis (median duration = 21.4 years)
 - 84% still working after a 30-year history of the disease
 - 77% still working after a 40-year history
 - 13.2% reported work disability
 - 57 subjects (24.3%) had received work disability payments
 - 46% of these individuals received payments for ≤1 year

Economic Burden of Rheumatoid Arthritis

Cost	Amount (2000 USD)							
	With rheumatoid arthritis	With osteoarthritis	No arthritis					
Direct	5763	N/A	N/A					
Indirect	2785	1011	1002					
Lifetime [†]	61,000-122,000*	N/A	N/A					

Individuals with rheumatoid arthritis are estimated to be 6 times more likely than people without rheumatoid arthritis to incur medical charges.

*USD in 1995 † 25 years following a diagnosis of rheumatoid arthritis; lifetime costs were higher among younger people with rheumatoid arthritis N/A = not available; USD = United States Dollars

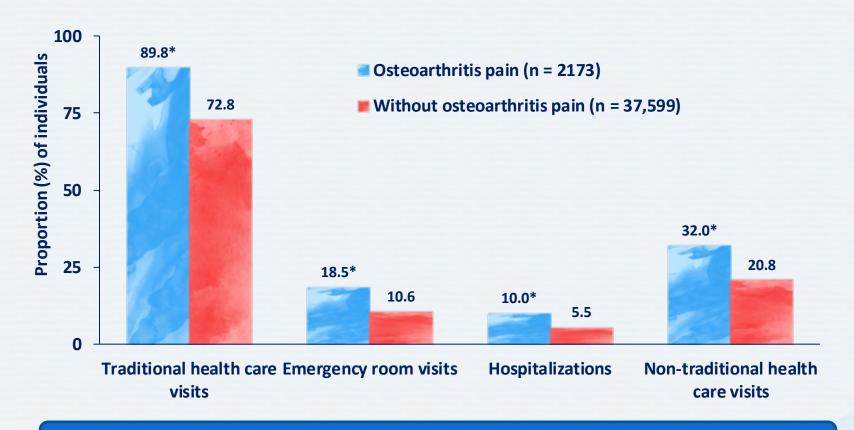
Economic Burden of Rheumatoid Arthritis: Societal Costs

Annual cost	Amount (billions) (2005 USD)				
Direct	8.4				
Indirect	10.9				

Total costs are approximately \$14,900 per patient per year.

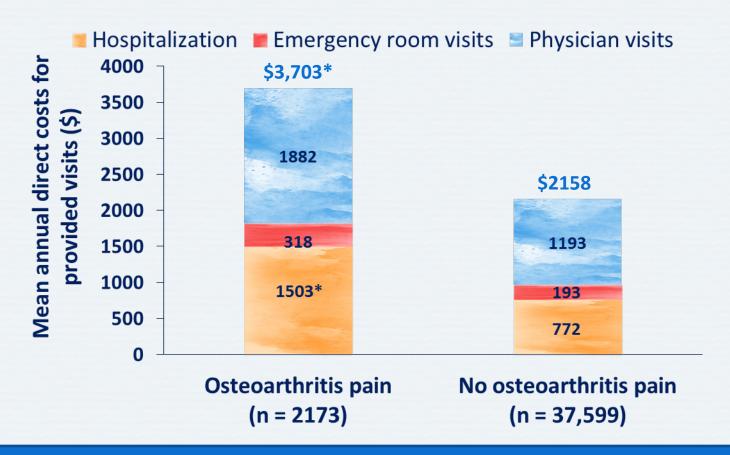
Rheumatoid arthritis is a costly, chronic disease.

Osteoarthritis Increases Health Care Utilization



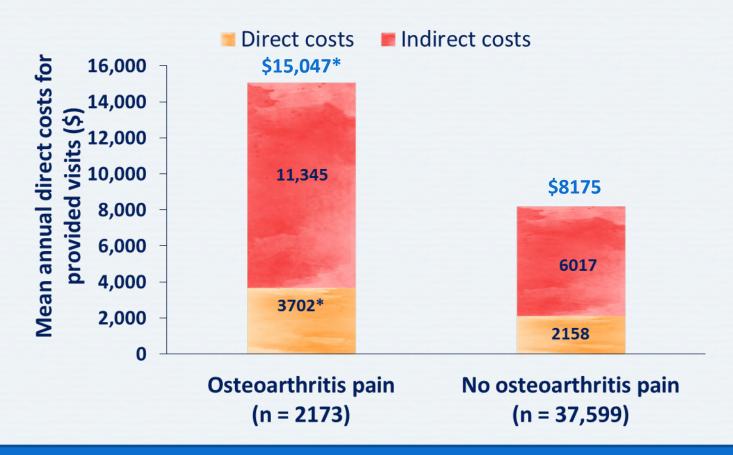
Workers with osteoarthritis pain have higher health care utilization

Osteoarthritis Increases Direct Health Care Costs



Workers with osteoarthritis pain have increased health care costs.

Osteoarthritis Increases Total Health Care Costs†



Workers with osteoarthritis pain have increased health care costs.

Economic Burden of Osteoarthritis*

Applied cost per petient (p=129,402)*	Amount (2007 USD)				
Annual cost per patient (n=128,493)*	Women	Men			
Out-of-pocket	1379	694			
Insurer	4833	4036			

- Costs associated with osteoarthritis are high
- Osteoarthritis costs are disproportionately higher for women
- Out-of-pocket expenses are substantial

Economic Burden of Osteoarthritis

- Administrative claims for about 5 million privately insured patients (aged 18–64 years) with ICD-9-CM codes for osteoarthritis (1998–2004)
- Mean annual direct and indirect per-patient costs* calculated from employer's perspective

Annual cost per patient	Amount (2005 USD)			
Direct medical	8601			
Prescription pain medications	445			
Indirect medical	4603			

Osteoarthrits is a costly, chronic disease

^{*}Mean costs were calculated from January 2003 to December 2004 and converted to annual 2005 dollars ICD-9-CM = International Classification of Diseases, 9th Revision, Clinical Modification; USD = United States Dollars White AG et al. J Occup Environ Med 2008; 50(9):998-1005.

Comorbidities

Chronic Joint Pain Conditions Have Sleep and Mental Health Comorbidities

				De	epression (%)			
Pain Cohorts	>1 sleep comorbidity (%)	>1 mental health comorbidity (%)	Mental health condition Mean (SD)	MDD	Other depressive symptoms	Total	Other psych disorders (%)	Anxiety (%)
OA	8.0	15.2	1.4 (0.7)	3.5	3.4	6.2	4.8	4.5
RA	5.7	11.7	1.3 (0.7)	3.2	2.8	5.5	3.5	3.6
AS	7.4	17.3	1.4 (0.7)	4.4	4.1	4.7	4.1	5.8
PsA	7.6	13.3	1.3 (0.6)	3.5	3.2	6.1	2.8	3.9

Chronic Joint Pain Comorbidities

Pain cohorts	Comorbid pain conditions (%)								Maara		
	Stroke	LR	CR	Fibro	OA	LBP	Migraine	RA	PBS	IC	Mean
Osteoarthritis	6.4	4.8	2.6	3.3	100.0	19.3	1.7	2.1	2.1	0.2	1.5
Rheumatoid arthritis	5.0	3.0	1.4	3.8	21.0	12.8	1.9	100.0	1.7	0.2	1.6
Ankylosing spondylitis	3.9	10.3	2.9	6.3	20.4	31.9	2.9	3.2	2.0	0.3	1.9
Psoriatic arthritis	2.6	1.8	1.5	2.5	16.0	10.6	1.6	17.1	1.8	0.1	1.6

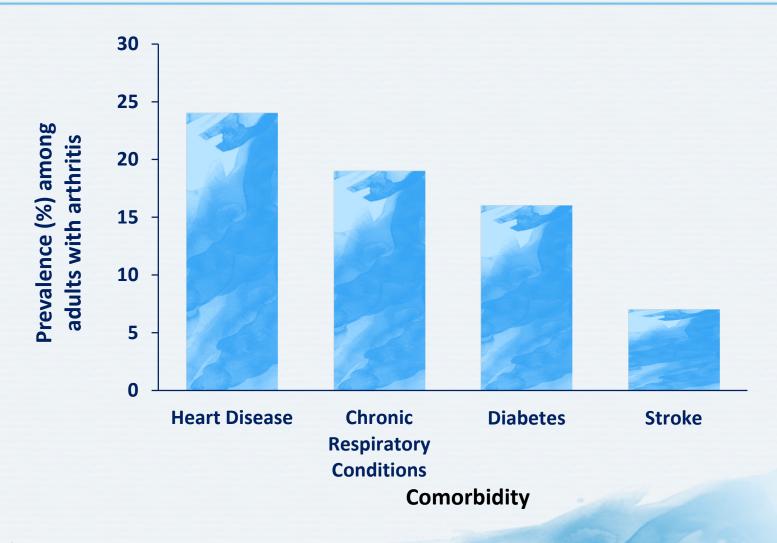
Note: infrequent comorbid conditions were omitted from the comorbid pain conditions in the table.

CR = cervical radiculopathy; HIV = human immunodeficiency virus; IC = interstitial cystitis; Fibro = fibromyalgia;

LR = lumbar radiculopathy; MS = multiple sclerosis; OA = osteoarthritis; PBS = painful bladder syndrome; RA = rheumatoid arthritis

Davis JA et al. J Pain Res 2011; 4:331-45; Dworkin RH et al. J Pain 2010; 11(4):360-8; Riley GF. Med Care 2009; 47(7 Suppl 1):S51-5.

Comorbidities Associated with Arthritis



Comorbidities Associated with Ankylosing Spondylitis

- Most common comorbidities:
 - Hypertension
 - Peptic ulcers
 - Headaches
 - Uveitis
 - Bowel inflammation
 - Psoriasis
 - Heart disease

Comorbidities Associated with Rheumatoid Arthritis

- On average, patients with established rheumatoid arthritis have ≥2 comorbidities
- Especially problematic:
 - Cardiovascular disease
 - Interstitial lung disease
 - Osteoporosis
 - Cancer
 - Depression

Rheumatoid Arthritis Increases Cardiovascular Risk

Cardiovascular risk in rheumatoid arthritis



Cardiovascular risk in type 2 diabetes

Risk score models should be adapted for patients with rheumatoid arthritis by introducing a *multiplication factor of 1.5* when ≥2 of the following conditions are met:

- Disease duration >10 years
- RF or anti-CCP positivity
- Presence of certain extra-articular manifestations

Comorbidities Associated with Osteoarthritis

- Older patients with osteoarthritis are likely to have comorbidities
 - Visual disorders, diabetes, and heart disease occur more frequently in patients with osteoarthritis
 - Result in adverse physical function
 - Associated with adverse outcomes in joint replacement
- Recognizing and treatment comorbidities is crucial in preventing or reducing related physical decline

Summary

Burden of Illness in Chronic Joint Pain: Summary

- Chronic joint pain can severely compromise functioning, quality of life and ability to work
- It also increases health care utilization costs
- In addition, patients suffering from chronic joint pain, frequently have other comorbidities, such as sleep disturbances, mood disorders, cardiovascular disease and other chronic conditions, all of which much also be addressed