Chapter 10. Musculoskeletal and joint diseases

For cost information please go to the most recent cost comparison charts

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- 10.1 Drugs used in rheumatic diseases and gout
 10.2 Drugs used in neuromuscular disorders
 10.3 Drugs for the relief of soft-tissue inflammation

Key

R	Red drug see GMMMG RAG list Click on the symbols to access this list
A	Amber drug see GMMMG RAG list Click on the symbols to access this list
Gn	Green drug see GMMMG RAG list Click on the symbols to access this list
U	If a medicine is unlicensed this should be highlighted in the template as follows Drug name \ensuremath{U}
	Not Recommended
отс	Over the Counter
Order of Drug Choice	Where there is no preferred 1^{st} line agent provided, the drug choice appears in alphabetical order.



BNF chapter	10 Musculoskeletal and joint diseases		
Section	10.1 Drugs used in rheumatic diseases and gout		
Subsection	10.1.1 Non-steroidal anti-inflammatory drugs		
First choice	Ibuprofen tablets 200mg; 400mg; 600mg	MHRA DSU: High-dose ibuprofen: small increase in cardiovascular risk, June 2015	
Alternatives	Naproxen tablets 250mg; 500mg		
	Diclofenac tablets 50mg	MHRA DSU: diclofenac: new contraindications and warnings, June 2013. MHRA DSU: NSAIDs: cardiovascular risks October 2012. Further evidence that the cardiovascular risk with diclofenac is higher than other non-selective NSAIDs and similar to the selective COX-2 inhibitors	
	Etoricoxib tablets 30mg, 60mg, 90mg, 120mg	MHRA DSU: Etoricoxib (Arcoxia): revised dose recommendation for rheumatoid arthritis and ankylosing spondylitis MHRA DSU: etoricoxib: prescribing to patients with high blood pressure, July 2008.	
Grey drugs Items which are listed as Grey are deemed not suitable for routine prescribing but may be suitable for a defined patient population	Piroxicam Capsules, orodispersible tablets Systemic piroxicam should be initiated only by specialists as a second-line treatment for arthritis. Patients who currently take piroxicam should be reassessed at a routine appointment.	Gn following specialist initiation Criterion 1 (see RAG list)	
Do Not Prescribe	Naproxen with esomeprazole Modified-release tablets	Criterion 2 (see RAG list)	
General Guidance applicable to all drugs			

General Guidance applicable to all drugs

• NICE CG177: Osteoarthritis: Care and management in adults.



- NICE NG100: Rheumatoid arthritis in adults: management.
- CKS (2013) NSAID prescribing issues
- BNF: NSAID-associated ulcers
- CKS (2013): Patients on low dose aspirin
- MHRA DSU: Non-steroidal anti-inflammatory drugs: reminder on renal failure and impairment, May 2009
- MHRA DSU: Non-steroidal anti-inflammatory drugs: cardiovascular risk, October 2012
- MHRA DSU: NSAIDs and coxibs: balancing of cardiovascular and gastrointestinal risks, December 2007

Subsection 10.1.2 Corticostero	ds
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Systemic corticosteroids

The general actions, uses, and cautions of corticosteroids are described in the BNF.

Local corticosteroid injections

First choice	Methylprednisolone acetate 40mg/ml or	
	Methylprednisolone acetate 40mg, lidocaine hydrochloride 10mg/ml	
Alternatives	Dexamethasone sodium phosphate 3.8mg/ml	MHRA DSU: Dexamethasone 4 mg/ml injection (Organon Laboratories Limited): reformulation with changes in name, concentration, storage conditions and presentation, October 2014.
	Hydrocortisone acetate 25mg/ml	
	Triamcinolone acetonide 10mg/ml Triamcinolone acetonide 40mg/ml	



Subsection	10.1.3 Drugs that suppress the rheumatic dise	ase process	
DMARDS			
First Choice	Methotrexate 2.5mg tablets	À	
		NPSA Alert, June 2006. Improving compliance with oral methotrexate guidelines.	
		When prescribing oral methotrexate only ever prescribe 2.5mg tablets and pay particular attention to correct dosing instructions – weekly dosing.	
		Co-prescribe oral folic acid 5mg weekly with methotrexate (to be taken at least 24 hours after the methotrexate [often 2 -3 days after]) and increase dose if necessary dependant on folate levels.	
	Methotrexate 50mg/ml pre-filled Pen injection (Metoject® brand only amber formulation)	A Specialist initiation only	
		If moving the parenteral formulation to homecare, clinicians should ensure that responsibility for the different aspects of care is clearly defined (in particular blood monitoring.	
	Sulfasalazine e/c 500mg tablets	A	
	Hydroxychloroquine sulphate 200mg tablets	A	
Alternatives	Leflunomide tablets 10mg; 15mg; 20mg	A Specialist initiation only	
	Azathioprine tablets 25mg; 50mg	A	
	Ciclosporin capsules 10mg; 25mg; 50mg; 100mg	BNF Online Patients should be stabilised on a particular brand of oral ciclosporin because switching between formulations without close	



		monitoring may lead to clinically important changes in blood-ciclosporin concentration. Prescribing and dispensing of ciclosporin should be by brand name to avoid inadvertent switching.
Mycophenolate U Tablets 500mg, Capsules 250mg No. 10		MHRA DSU: Mycophenolate mofetil, mycophenolic acid: new pregnancy-prevention advice for women and men 2015 MHRA DSU: Mycophenolate mofetil: pure red cell aplasia, July 2009 MHRA DSU: Mycophenolate mofetil (CellCept) and mycophenolic acid: risk of hypogammaglobulinaemi a and risk of bronchiectasis, January 2015
	nce applicable to all drugs 1: Rheumatoid arthritis in adults: management.	
Biologics (sec	ondary care use only)	
GMMMG High	Cost Drugs Pathway for Rheumatoid Arthritis (De	ecember 2017)
First Choices (Follow relevant pathway)	Abatacept Injection or infusion	Prevents full activation of T-lymphocytes
	Adalimumab Solution for injection	TNF inhibitor NICE TA195: Adalimumab, etanercept, infliximab, rituximab and abatacept for the treatment of rheumatoid arthritis after the failure of a TNF inhibitor. NICE TA375: Adalimumab, etanercept, infliximab, certolizumab pegol, golimumab, tocilizumab and abatacept for rheumatoid arthritis not previously treated



	with DMARDs or after conventional DMARDs only have failed TA392: Adalimumab for treating moderate to severe hidradenitis suppurativa
Apremilast [▼] (Otezla®) Tablets	PDE4 Inhibitor TA433: Apremilast for treating active psoriatic arthritis MHRA: Apremilast (Otezla ▼): risk of suicidal thoughts and behaviour
Baricitinib [▼] (Olumiant®) 2mg and 4mg tablets	NICE TA466: Baricitinib for moderate to severe rheumatoid arthritis
Belimumab Infusion	Inhibits activity of B-lymphocyte stimulator TA397: Belimumab for treating active autoantibody-positive systemic lupus erythematosus
Certolizumab Solution for injection	NICE TA415: Certolizumab pegol for treating rheumatoid arthritis after inadequate response to a TNF-alpha inhibitor NICE TA445: Certolizumab pegol and secukinumab for treating active psoriatic arthritis after inadequate response to DMARDs
Etanercept Solution for injection	NICE TA35: Guidance on the use of etanercept for the treatment of juvenile idiopathic arthritis. NICE TA199: Etanercept, infliximab and adalimumab for the treatment of psoriatic arthritis.



Golimumab	R TNF inhibitor
Solution for injection	NICE TA220: Golimumab for the treatment of psoriatic arthritis.
	NICE TA225: Golimumab for the treatment of rheumatoid arthritis after the failure of previous disease-modifying anti-rheumatic drugs.
	NICE TA497: Golimumab for treating non-radiographic axial spondyloarthritis
Infliximab	R TNF inhibitor
Infusion	
Ixekizumab ▼	R Interleukin 17 antagonist
Solution for injection	NICE TA537: Ixekizumab for treating active psoriatic arthritis after inadequate response to DMARDs
Rituximab Infusion	Anti-lymphocytic - causes lysis of B lymphocytes
	MHRA DSU: Rituximab: progressive multifocal leukoencephalopathy in a patient without prior treatment for rheumatoid arthritis, December 2009.
	MHRA DSU: Rituximab: screen for hepatitis B virus before treatment, December 2013
	NICE TA308: Rituximab in combination with glucocorticoids for treating anti-neutrophil cytoplasmic antibody-associated vasculitis.
	GMMMG Rituximab biosimilar recommendation (August 2017)
Sarilumab▼	R Interleukin-6 antagonist
Solution for injection	NICE TA485: Sarilumab for moderate to severe rheumatoid arthritis



Secukinumab Solution for injection	TA407: Secukinumab for active ankylosing spondylitis after treatment with non-steroidal anti-inflammatory drugs or TNF-alpha inhibitors
Tocilizumab Injection, infusion	Interleukin-6 antagonist NICE TA238: Systemic juvenile idiopathic arthritis – tocilizumab. NICE TA247: Tocilizumab for the treatment of systemic juvenile idiopathic arthritis. NICE TA518: Tocilizumab for treating giant cell arteritis
Tofacitinib▼ 5mg and 10mg tablets	Janus Kinase Inhibitor NICE TA480: Tofacitinib for moderate to severe rheumatoid arthritis
Ustekinumab Solution for injection	NICE TA340: Ustekinumab for treating active psoriatic arthritis.

General guidance applicable to all drugs

- Biologics in Rheumatoid Arthritis can be used, without prior funding approval, according to GMMMG approved pathways <u>High Cost Drugs Pathway for Rheumatoid Arthritis</u> and <u>Harmonised Biologics Pathway for Ankylosing Spondylitis and Psoriatic Arthritis</u>
- MHRA DSU: Tumour necrosis factor alpha inhibitors: risk of tuberculosis—screen all patients before starting treatment and monitor them closely, April 2014
- MHRA DSU: TNFa inhibitors: risk of TB screen all patients before starting treatment and monitor them closely(April 2016)

Additional NICE guidance and technology appraisals:

- NICE NG100: Rheumatoid arthritis in adults: management.
- <u>NICE TA383: TNF-alpha inhibitors for ankylosing spondylitis and non-radiographic axial</u> spondyloarthritis

Subsection	10.1.4 Gout and cytotoxic induced hyper	uricaemia
Acute attack		
First choice Alternative where NSAIDS contraindicated	Colchicine 500microgram tablets	MHRA DSU: colchicine: extremely toxic in overdose, November 2009. British Society for Rheumatology Guideline for the



		Management of Gout		
Long-term cor	Long-term control of gout			
First choice	Allopurinol tablets			
	100mg; 300mg			
Alternatives	Febuxostat tablets	Gn		
	80mg; 120mg	Use only where allopurinol is contraindicated or not tolerated. (As per NICE guidance and/or on specialist advice.)		
		NICE TA164: Febuxostat for the management of hyperuricaemia in people with gout.		
		MHRA DSU: Febuxostat (Adenuric▼): stop treatment if signs or symptoms of serious hypersensitivity occur, June 2012.		
	Sulfinpyrazone tablets			
	100mg; 200mg			
Do Not Prescribe	Lesinurad Tablets	<u>Criterion 2 (see</u> <u>RAG list)</u>		

General guidance applicable to all drugs

British Society for Rheumatology Guideline for the Management of Gout

NICE Technology Appraisals:

- NICE TA291: Pegloticase for treating severe debilitating chronic tophaceous gout
- NICE TA281: Canakinumab for treating gouty arthritis attacks and reducing the frequency of subsequent attacks (terminated appraisal)

Subsection 10.1.5 Other drugs for rheumatic diseases

Applicable guidance:

- <u>NICE CG177: Osteoarthritis: Care and management in adults.</u> section 1.4.5 Do not offer glucosamine or chondroitin products for the management of osteoarthritis.
- GMMMG 'Do Not Prescribe and Grey Lists': Glucosamine.
- GM Policy Statement (2017). Hyaluronic Acid injections for Osteoarthritis. (GM037)

NICE Technology Appraisals:

• NICE TA477: Autologous chondrocyte implantation for treating symptomatic articular cartilage



defects of the knee

• NICE TA508: Autologous chondrocyte implantation using chondrosphere for treating symptomatic articular cartilage defects of the knee

Do Not Prescribe	Glucosamine With or without chondroitin	<u>Criterion 1 (see</u> <u>RAG list)</u>
	Synovial fluid injections Including hyaluronan and sodium hyaluronate injections	<u>Criterion 1 (see</u> <u>RAG list)</u>



Section	10.2 Drugs used in neuromuscular disorders		
Subsection	10.2.1 Drugs that enhance neuromuscular transmission		
First choice	Pyridostigmine 60mg tablets		
Alternatives	Neostigmine 15mg tablets		
Subsection	10.2.2 Skeletal muscle relaxants		
First choice	Dantrolene capsules		
	25mg; 100mg		
Alternatives	Diazepam		
	tablets 2mg; 5mg; 10mg		
	oral solution 2mg/5ml		
	Baclofen 10mg tablets		
	Baclofen oral solution 5mg/5ml		
	Tizanidine tablets	Gn Following	
	2mg; 4mg	specialist initation	
Do Not Prescribe	Cannabis extract (Sativex®)	Criterion 3 (see RAG list)	
	Oromucosal spray		
Section	Miscellaneous		
Do Not	Therabite [®]	Criterion 1 (see	
Prescribe	Jaw rehabilitation system	RAG list)	



Section	10.3 Drugs for the relief of soft-tissue inflammation			
Subsection	10.3.1 Enzymes (Secondary care use)			
	Collagenase (Xiapex®)	R		
		NICE TA459: Collagenase clostridium histolyticum for treating Dupuytren's contracture		
Subsection	10.3.2 Rubefacients and other topical	antirheumatics		
Rubefacients				
Do Not Prescribe	Rubefacients		Criterion 1 (see RAG list)	
	Topical rubefacient products may contain n salicylate compounds, essential oils, capsic camphor. Topical NSAID or capsaicin preps rubefacients	um, and	RAG IISL)	
Topical NSAID	s			
Options	Ibuprofen 5% Gel			
	(Available as 30g, 50g or 100g tubes)			
	Ketoprofen 2.5% Gel		MHRA DSU: topica	
	(Available as 30g, 50g or 100g tubes)		ketoprofen: reminder on photosensitivity reactions, June 2009 and MHRA August 2010:	
	Piroxicam 0.5% Gel			
	(Available as 60g or 112g tubes)			
Capsaicin				
Options	Capsaicin 0.025% cream		NICE CG177: Osteoarthritis: Care and management in adults. GMMMG Neuropathic Pain	
	Compaigin 0.075% avenue		Guideline March 2014	
	Capsaicin 0.075% cream			
Poultices				