PAN MERSEY APC

PAN MERSEY AREA PRESCRIBING COMMITTEE
SHARED CARE FRAMEWORK
FIRST APC BOARD DATE: 27 SEP 2017
LAST APC BOARD DATE: 28 NOV 2017

NHS Pan Mersey Area Prescribing Committee

AZATHIOPRINE for patients within adult services

1. Background	Azathioprine is used as an immunosuppressant anti-metabolite either alone or, more commonly, in combination with other agents (usually corticosteroids) to influence the immune response. Therapeutic effect may be evident only after weeks or months and can include a steroid-sparing effect, thereby reducing the toxicity associated with high dosage and prolonged usage of corticosteroids. Indications, dose adjustments and monitoring requirements for disease modifying drugs (DMDs) (licensed and unlicensed indications) included in this Framework are in line with national guidance published by the British Society for Rheumatology 2017.
2. Licensed Indications	• Rheumatoid arthritis • Systemic lupus erythematosus (SLE), • Dermatomyositis and polymyositis • Pemphigus vulgaris • Auto-immune chronic active hepatitis • Polyarteritis nodosa • Auto-immune haemolytic anaemia • Chronic refractory idiopathic thrombocytopenic purpura • Inflammatory bowel disease • **Transplant indications are not included**
3. Locally agreed off-label use	• Psoriasis and psoriatic arthritis • Chronic eczema and other autoimmune skin conditions • Interstitial lung disease • Steroid sparing agent • Connective tissue diseases • Myasthenia gravis, inflammatory myopathies and neuropathies, vasculitis and other immune-mediated central and peripheral nervous system diseases • Autoimmune and inflammatory kidney conditions • Sarcoidosis • Atypical neuro-inflammatory disease •
4. Initiation and ongoing dose regime	**Transfer of monitoring and prescribing to Primary care is normally after 3 months** **The duration of treatment will be determined by the specialist based on clinical response and tolerability**
	1–3 mg/kg daily, adjusted according to response (consider withdrawal if no improvement or stabilisation within 3 months) A dose reduction of 25% may be considered for CKD 4 and 50% for CKD 5. See Table 4 in BSR monitoring guidelines.

<table>
<tr><td></td><td colspan="2">Please note for rheumatology conditions a patient may be initiated on more than one DMD.</td></tr>
<tr><td></td><td colspan="2">All dose adjustments will be the responsibility of the initiating specialist unless directions have been discussed and agreed with the primary care clinician

Dose increases should be monitored by FBC creatinine/eGFR, ALT and/or AST and albumin every 2 weeks for 6 weeks after the dose increase, then revert back to previous schedule.</td></tr>
<tr><td></td><td colspan="2">Termination of treatment will be the responsibility of the specialist.</td></tr>
<tr><td>5. Baseline investigations, initial monitoring and dose titration to be undertaken by specialist</td><td colspan="2">Baseline:
Height, weight, BP, FBC, creatinine/eGFR, ALT and/or AST and albumin.
Baseline thiopurine methyltransferase (TMPT) status
Vaccinations against pneumococcus and influenza are recommended.
Shingles vaccine (Zostavax) is recommended as per the JCVI for eligible patients.
Specialist to highlight in the first clinic letter notifying the GP of the decision to initiate DMDs that the GP will need to give the shingles vaccine if the patient is older than 69 years and the pneumococcal vaccine if this has not already been given. The GP should also be advised to add the patient to the influenza vaccine list.
Patients should be assessed for comorbidities that may influence DMD choice, including evaluation of respiratory disease and screening for occult viral infection.
Initiation:
FBC, creatinine/eGFR, ALT and /or AST and albumin every 2 weeks until on stable dose for 6 weeks;
Once on stable dose, monthly FBC, creatinine/eGFR, ALT and /or AST and albumin for 3 months
(There may be different initial monitoring for gastroenterology conditions)</td></tr>
<tr><td rowspan="2">6. Ongoing monitoring requirements to be undertaken by primary care.</td><td>Monitoring</td><td>Frequency</td></tr>
<tr><td>FBC, creatinine/eGFR, ALT and/or AST and albumin

CRP and ESR (rheumatology patients only)</td><td>After the initial monitoring period (see section 5), every 12 weeks, or more frequently in patients at higher risk of toxicity as advised by the specialist team. NB: Some of the initial monitoring (likely to be 1-2 months of monthly monitoring) may take place in primary care. The exact frequency of the monitoring to be communicated by the specialist in all cases.

This includes patients heterozygous of TMPT</td></tr>
</table>

7. Pharmaceutical aspects	Route of administration	Oral
	Formulation	Azathioprine 25mg and 50mg tablets
	Administration details	Tablets should be taken at least 1 hour before food or 3 hours after food or milk.
8. Contraindications **Please note this does not replace the Summary of Product Characteristics (SPC) and should be read in conjunction with it.**	• Hypersensitivity to azathioprine or mercaptopurine. • Azathioprine – induced pancreatitis • **Very low TPMT activity (Homozygous recessive): Avoid**. **Can be fatal**	
9. Significant drug interactions	If considering prescribing allopurinol, refer the patient back to the consultant for advice and a dose adjustment. If allopurinol is given concomitantly with azathioprine, the dose of azathioprine should be reduced to 25 % of the original dose. Monitoring will continue as above. For a comprehensive list consult the BNF or Summary of Product Characteristics. SPC Seek advice from the initiating Specialist if there are any concerns about interactions.	
10. Adverse Effects and managements	**Result**	**Action**
	Abnormal bruising or severe sore throat	Stop drug until FBC results available, contact Specialist Nurse (SN)
	Fall in WCC <3.5 x 10^9/l Fall in neutrophils <1.6 x 10^9/l Fall in platelets <140 x 10^9/l	Stop drug SN for advice and management
	Increased MCV >105fl	Check folate, B12 & TSH. Treat if abnormal, contact specialist nurse for advice if normal
	Unexplained reduction in albumin <30g/L Abnormal LFTs – AST or ALT > 100u/l Rash Mouth ulcers	Contact SN for advice and management
	Acute abdominal pain	Check serum amylase. Consider pancreatitis.

	Increase in serum creatinine >30% over period of 12 months or less OR decline in eGFR > 25%	Contact SN if there is new or unexplained renal impairment
11. Advice to patients and carers	The specialist will counsel the patient with regard to the benefits and risks of treatment and will provide the patient with any relevant information and advice, including patient information leaflets on individual drugs.	
12. Pregnancy and breast feeding	Compatible throughout pregnancy at ≤ 2 mg/kg/day after a careful assessment of risk versus benefit. Compatible with breastfeeding Compatible with paternal exposure (BSR & BHPR guideline on prescribing in pregnancy and breastfeeding)	
13. Specialist contact information	See appendix 2	
14. Additional information	**Where patient care is transferred from one specialist service or GP practice to another, a new shared care agreement must be completed.**	
15. References	BSR monitoring guidelines	
16. To be read in conjunction with the following documents.	1. Policy for Shared Care 2. Shared care agreement. When two or more DMDs are initiated, one shared care agreement form should be completed for all relevant drugs.	

Appendix 1

Policy for Shared Care

Shared care is only appropriate if it provides an optimum solution for the patient and it meets the criteria outlined in the Shared Care section of the Pan Mersey **Definitions and Criteria for Categorisation of Medicines in the Pan Mersey Formulary** document.

Before prescribing responsibilities are transferred to primary care:

- Prescribing responsibility will only be transferred when the consultant and the patient's GP agree that the patient's condition is stable.
- All information required by the shared care framework for the individual medicine has been provided to the patient's GP.
- Patients will only be referred to the GP once the GP has agreed to the Shared Care Agreement and returned signed copies.

Inherent in any shared care agreement is the understanding that participation is at the discretion of the GP, subject to the availability of sufficient information to support clinical confidence.

Specialist Responsibilities in Shared Care

- To initiate the medicine, prescribe, monitor for toxicity and efficacy as described by the shared care framework until the patient is stabilised.
- To ensure the patient or their carer:
    - Is counselled with regard to the risks and benefits of the medicine.
    - Provide any necessary written information to the patient with regard to the individual medicine including patient information leaflets on individual drugs.
    - Obtain and document informed consent from the patient when any medicines is prescribed for an off-label indication for any condition
- To be familiar with the shared care framework.
- To provide all information to the patient's GP as required by the shared care framework when prescribing responsibility is initially transferred and at any subsequent times as necessary for safe and effective treatment of the patient.
- To assess the patient regularly as necessary for the duration of therapy.
- To review the patient promptly if required by the GP concerned.
- To meet any additional requirements as required by the individual medicine shared care framework.

- To communicate failure of a patient to attend a routine hospital review and advise the GP of appropriate action to be taken.
- **Addition of a second DMD**: Following the addition of a new drug to an existing regime covered by a Shared Care Agreement, the Specialist must initiate, prescribe and monitor the new drug in accordance with the relevant shared care agreement including subsequent review and inform the GP of this. A new Shared Care Agreement must then be initiated for the new drug.

Primary Care Responsibilities in Shared Care

- To reply to a written request for Shared Care within 21 days ensuring both copies of the Shared Care Agreement are signed if appropriate.

If agreeing to shared care, the GP is asked to:

- To provide prescribe or manage and monitor the medicine as advised by the Specialist and in line with the individual Shared Care Framework.
- To review the patient as required by the Shared Care Framework
- To make appropriate and contemporaneous records of prescribing and/or monitoring and to note the existence of the Shared Care Agreement on the patient`s clinical record. A READ code of “6652 Shared Care- Specialist/GP” can be used.
- To be familiar with the individual Shared Care Framework.
- To report any adverse effects of treatment to the specialist team.
- To inform the Specialist of any relevant change in the patient’s circumstances.
- To seek Specialist advice as appropriate.
- To meet any additional requirements as required by the individual Shared Care Framework.
- To respond to Specialist communication relating to any change or addition to the patients treatment covered by the Shared Care Agreement.

Appendix 2: Shared Care Agreement

Disease modifying drugs (DMDs)

Request by Specialist Clinician for the patient's GP to enter into a shared care agreement

Part 1
To be signed by Consultant / Prescribing member of Specialist Team

Date _____

Name of patient_____

Address _____

_____

Patient NHS No _____

Patient hospital unit No _____

Diagnosed condition _____

If using addressograph label please attach one to each copy

Dear Dr_____

I request that you prescribe

(1) _____

(2) _____

(3) _____

(4) _____

for the above patient in accordance with the enclosed shared care framework.

Last Prescription Issued: ….. / ….. / ….. Next Supply Due: ….. / ….. / …..

Date of last blood test: ….. / ….. / ….. Date of next blood test: ….. / ….. / …..

Frequency of blood test: ……

I confirm that the patient has been stabilised and reviewed on the above regime in accordance with the Shared Care Framework and Policy. ☐

I confirm that if this is a Shared Care Agreement for a drug indication which is unlicensed or off label, informed consent has been received. ☐ N/A

Details of Specialist Clinicians

Name _____Date_____

Consultant / Associate Specialist / Prescribing member of Specialist Team ***circle or underline as appropriate***

Signature _____

In all cases, please also provide the name and contact details of the Consultant.

When the request for shared care is made by a prescriber who is not the consultant, it is the supervising consultant who takes medico-legal responsibility for the agreement.

Consultant: _____

Contact details*:*

Telephone number: _____Ext:_____

Address for return of documentation _____

_____

_____

Part 2
To be completed by Primary Care Clinician

I agree to prescribe _____for the above patient in accordance with the enclosed shared care framework.

GP signature _____Date _____

GP name _____Please print

GP: *Please sign and return a copy* ***within 21 calendar days*** *to the address above*

OR

GP- If you do not agree to prescribe, please delete the section above and provide any supporting information as appropriate below: