

# Medication in paediatric transplant

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# Medication used in transplant

- \* Immunosuppressants
- \* Antibiotics
- \* Antivirals
- \* Statins

# Immunosuppressants

- \* Ciclosporin
- \* Tacrolimus
- \* Sirolimus
- \* Azathioprine
- \* Mycophenolate mofetil
- \* Steroids

# Ciclosporin

- \* First line immunosuppressant post transplant as available intravenously.
- \* Calcineurin inhibitor – works by reduces the proliferation of T cells
- \* It is always given twice a day
- \* Levels should be taken as trough (i.e. before the next dose)

# Ciclosporin



Available as capsules 10mg,  
25mg, 50mg and 100mg  
Liquid 100mg/ml  
Injection 250mg/5ml



When giving IV or NG  
ciclosporin it must be  
given by a PVC free  
giving set

# Side effects of ciclosporin

- \* Nephrotoxicity.
- \* Decreased white blood cells and platelets
- \* Increased risk of infection
- \* Risk of malignant skin changes
- \* Increase in liver function tests
- \* Hypertension
- \* Hypomagnesaemia
- \* Hyperglycaemia
- \* Hyperkalaemia
- \* Increase in blood lipids.

# Reasons for changing to Tacrolimus

- \* Cosmetic
- \* Rejection
- \* Unstable levels.



# How we treat side effects of ciclosporin

- \* Watch for medication that can cause kidney problems
  - ibuprofen
- \* Ensure sunscreen even in may not seem sunny – factor 50
- \* Treat hypomagnesiumia
- \* Treat hypertension – amlodipine/enalapril
- \* Regular monitoring
- \* Check for interactions – ask the ward pharmacist



# Tacrolimus

- \* Second line if ciclosporin is not appropriate.
- \* Also calcineurin inhibitor and reduce proliferation of T cells
- \* Always ensure the same brand is prescribed
- \* Check for interactions – ask the ward pharmacist
- \* Can cause QT prolongation
- \* Can cause ischaemic coronary artery disorders
- \* Levels may change with diarrhoea may increase

# Preparations of Tacrolimus



We have a newcastle special liquid, tacrolimus 1mg/ml

Also capsules, generally we always use Prograf, 500 microgram, 1mg ,5mg are kept in NUTH.

# Side effects

- \* Increased risk of infection
- \* GI perforation
- \* Risk of malignant skin changes
- \* Decreased pancytopenia
- \* Hyperglycaemia
- \* Increase blood lipids
- \* Low magnesium,
- \* phosphate and sodium
- \* Insomnia
- \* Hypertension
- \* Jaundice
- \* Nephrotoxicity

# Calcineurin Inhibitor Levels

- \* Levels are generally higher the nearer the patient is from transplant
- \* Levels decrease over time
- \* Levels may need to be increased if rejection
- \* Decreased if PTLD or renal failure

# Azathioprine

- \* Generally all patients start on Azathioprine post transplant once well fed.
- \* Works on the B cells in low doses by interfering with the production of DNA and stopping cells dividing and multiplying.
- \* In higher doses also works on T cells
- \* Azathioprine is cytotoxic

# Preparations of azathioprine



- \* Also available in oral suspension 50mg/5ml
- \* Injections where possible should be made in pharmacy
- \* Tablets should not be crushed or halved – use liquid instead



# Side effects of Azathioprine

- \* Increased risk of infection
- \* Pancytopenia
- \* Pancreatitis
- \* Alopecia
- \* Liver impairment
- \* Risk of malignant skin changes
- \* Renal impairment
- \* Hypersensitivity reactions
- \* Interstitial pneumonia.
- \* Gastric ulceration

# Monitoring to prevent toxicity

- \* Monitor white blood cells, platelets and Haemoglobin
- \* Suspend if low platelets
- \* Suspend in white blood cells less than 4 (aim for WCC should be 4-6)
- \* Suspend if deranged liver function tests.



# Reasons to switch to Mycophenolate Mofetil

- \* Low blood count but want another agent
- \* Rejection
- \* Unable to tolerate azathioprine
- \* High HLA antibodies.

# Mycophenolate mofetil (MMF)

- \* An alternative to Azathioprine as second agent
- \* Inhibits immunologically mediated inflammation which inhibits proliferation of T and B cells
- \* Not to be confused with mycophenolic acid
- \* There is potential to be changed to myfortic to reduce gastric side effects

# Preparations of MMF



# Side effects of MMF

- \* Increased risk of infection
- \* Risk of malignant skin changes
- \* Increased risk of digestive adverse events
- \* Pancytopenia
- \* Acidosis
- \* Changes in electrolytes
- \* Increased blood lipids
- \* Gout
- \* Agitation
- \* Convulsions
- \* Tachycardia
- \* Changes in liver and renal function

# Sirolimus

- \* Also an immune suppressant – but works slightly differently
- \* May be used if Stomach problems with MMF or low WCC with Azathioprine
- \* To reduce the amount of ciclosporin/tacrolimus required if patients have poor renal function
- \* Potentially prevents worsening coronary artery disease
- \* Always give in the morning

# Sirolimus preparations



Tablets also available as  
0.5mg and 1mg



# Sirolimus levels

- \* As a true level if only single against aim for 15
- \* However generally aim for levels about 5 – (sirolimus is normally used with another agents
  - \* Helps to minimise side effects

# Side effects

- \* Risk of malignant skin changes
- \* Increased risk of infection
- \* Impaired wound healing
- \* Increase in blood lipids
- \* Pancytopenia
- \* Hypertension
- \* Tachycardia
- \* Pancreatitis
- \* Hyperglycaemia
- \* Increase in creatinine
- \* Oedema
- \* Pneumonitis



# Special warning sirolimus

- \* Oral solution should be stored in the fridge
- \* Clearance may be reduced in poor renal function
- \* Not to be given with patients allergic to peanuts and soya.

# Steroids

- \* All patients will receive three doses of methylprednisolone post operatively
- \* Over 5 patients will get 0.2mg/kg for 6 weeks and then gradually weaned
- \* Teenagers get 1mg/kg until first biopsy and then gradually reduced to 0.2mg/kg
- \* May change depending on rejection and renal function
- \* Always should be given in the morning.

# Antibacterials

- \* Aciclovir should always be given for at least 3 months post heart transplant to prevent viral infection
  - \* If over 2 – 200mg tds, or under 2 100mg tds, but dose should be reduced in poor renal function
- \* Chlorhexidine mouth wash – 5mls-10mls twice a day to prevent mouth infections (alcohol free is available)

# Co-trimoxazole

Weight	Body Surface Area (BNFc)	Dose	Presentation
4.5-6kg	0.28-0.36	144mg OD	3ml of 240mg/5ml
7-9kg	0.37-0.47	192mg OD	4ml of 240mg/5ml
10-16kg	0.48-0.69m <sup>2</sup>	240mg OD	5ml of 240mg/5ml
17-26kg	0.70-0.96m <sup>2</sup>	384mg OD	8ml of 240mg/5ml
27-43kg	0.97-1.3m <sup>2</sup>	480mg OD	5ml of <u>480mg/5ml</u> or 1 x 480mg tablet
44-75kg	1.4-1.9m <sup>2</sup>	768mg OD	8ml of <u>480mg/5ml</u> or 1½ x 480mg tablets
>76kg	>2m <sup>2</sup>	960mg OD	10ml of <u>480mg/5ml</u> or 1 x 960mg tablet

# Statins

- \* Statins are used post transplant to prevent coronary artery disease which can occur post transplant
- \* Generally started aged 5 when can express painful legs
- \* Patients should have CK level checked, to prevent rhabdomyolysis
- \* This should always be given at night

Any questions

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Question?

