



Title:	Guidelines for Management of Acute Severe Asthma in Children over 2yrs		
Version:	3 PCCN 3		
	Supersedes:Version 2Summary of amendments in version 3:Aminophylline as 2nd intravenous agent (after IV magnesium sulfate bolus)Addition of potential to use short magnesium sulfate infusion (4 hours)Salbutamol dosing amended & capped (maximum equivalent to adult dose)Warning that high levels of salbutamol may cause toxicity including raised lactate & SVTClarification on how to manage bronchospasm in under 2 yearsAll children under 16 years age admitted to hospital in North West & North Wales region		
Originated /Mc &	odified By Originated By: North West (England) and North Wales Paediatric Transport Service Guideline authors:		
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Ratified by:       RMCH (Host Trust):         - Paediatric Medicines Management Committee (MMC)         - Paediatric Policies & Guidelines Committee			
Date of Ratifica			
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Responsibility of:	Clinical lead North West & North Wales Paediatric Critical Care Network & NWTS guideline lead consultant

EqIA Registration Number: 150/12
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## 1. Detail of Procedural Document

Guidelines for Management of Acute Severe Asthma in Children >2yrs.

This guideline is for use by staff working in the District General Hospitals of the North West (England) and North Wales region and NWTS team to use when caring for those over 2 years of age with an acute severe exacerbation of asthma. It focuses on acute management and potential differential diagnosis that need to be considered.

This does not replace an acute referral to NWTS team for advice on management, but is designed to help both NWTS and the referring team throughout the acute stabilisation period.

## 2. Equality Impact Assessment

EqIA Registration Number: 150/12

#### 3. Consultation, Approval and Ratification Process

This guideline was developed with input from:

- North West (England) and North Wales Paediatric Transport Service (NWTS) medical & nursing
- Paediatric Respiratory Consultants from Royal Manchester Children's Hospital and Alder Hey Children's Hospital
- Representatives from the District General Hospitals within the North West (England) & North Wales Paediatric Critical Care operational delivery network; includes medical, nursing and AHP (paediatrics, anaesthetics, and emergency medicine teams)
- Representatives from both Paediatric Critical Care Units (Royal Manchester Children's Hospital and Alder Hey Children's Hospital) medical and nursing

These guidelines were circulated amongst the North West and North Wales Paediatric Critical Care Network for comments on 25.03.2021 All comments received have been reviewed and appropriate amendments incorporated.

These guidelines were signed off by the Network's Joint Clinical Leads For ratification process see appendix 1.

## 4. References and Bibliography

See guidelines.

#### 5. Disclaimer

These clinical guidelines represent the views of the North West and North Wales Paediatric Critical Care Network and North West and North Wales Paediatric Transport Service, which were produced after careful consideration of available evidence in conjunction with clinical expertise and experience.

The guidance does not override the individual responsibility of healthcare professionals to make decisions appropriate to the circumstances of the individual patient.

# Clinical advice is always available 24/7 from NWTS on a case by case basis via the referral line: 08000 84 83 82

Please feel free to contact NWTS (01925 853 550) regarding these documents if there are any queries.



## Guidelines for Management of Acute Severe Asthma in Children OVER 2yrs



#### **SUMMON SENIOR HELP**

Give 15 L/min O<sub>2</sub> to achieve normal saturations (> 94%) Consider High Flow Humidified Nasal Cannula Oxygen Nebulised  $\beta_2$  agonist every 20 mins Nebulised Ipratropium bromide every 20 mins PO prednisolone / dexamethasone **OR** IV hydrocortisone

#### IF POOR RESPONSE AFTER 3 NEBULES:

Give IV magnesium sulfate 40 mg/kg (max 2 g) (0.16 mmol/kg, max 8 mmol) bolus over 20 mins

#### IF NOT IMPROVING RAPIDLY (within 10-20min):

Give Aminophylline loading 6 mg/kg (max 500mg) over 20 mins (if not on oral theophyllines) Then start aminophylline infusion

2-11 years @ 1 mg/kg/hr; 12-17 years @ 0.5-0.7 mg/kg/hr

#### IF NO IMPROVEMENT WITHIN 30 MINUTES OR CONTINUING TO DETERIORATE:

Give **2nd** bolus dose intravenous **magnesium sulfate OR** consider short (4 hrs) **magnesium sulfate infusion** Give **salbutamol bolus** over 5 mins & pause & reassess Blood gas + lactate

URGENT CONSULTANT ANAES/PAEDS REVIEW CONTACT NWTS FOR ADVICE: 08000 84 83 82

#### **STILL NO IMPROVEMENT**

Consider **3rd** bolus IV **magnesium** if not on magnesium infusion Start **salbutamol infusion** 1 microgram/kg/min (MAX 20 micrograms/min) **Nebulised adrenaline** 400 micrograms/kg (max 5 mg) Consider CXR/antibiotics/alternative diagnoses Blood gas + lactate **PREPARE FOR INTUBATION** 



#### **IMPENDING CARDIO-RESPIRATORY ARREST**

SpO2 ≤ 92% in O<sub>2</sub> PLUS any of: Cyanosis Worsening agitation / level of consciousness Poor respiratory effort or exhaustion or silent chest PRE-TERMINAL SIGNS: High CO<sub>2</sub> Acidaemia (pH ≤ 7.2) Hypotension

## **INTUBATE & VENTILATE**

#### **SEVERE ASTHMA**

SpO<sub>2</sub> <92% in air Use of accessory muscles Difficulty talking or eating Agitated HR >140 /min (< 5yr) HR >125 / min (>5yr) RR > 40 / min (<5yr) RR > 30 / min (>5yr)

#### HIGH RISK CLINICAL SIGNS Agitated or drowsy

Unable to talk or feed  $SpO_2 \le 92\%$  in air,  $pO_2 < 8$  kPa  $pCO_2$  'normal' (4.6-6 kPa) Silent chest Exhaustion/poor respiratory effort

#### INDICATIONS FOR CXR

Surgical emphysema Severe / Life-threatening asthma not responding to treatment OR to exclude: Pneumothorax, Lobar collapse/ consolidation FB or mediastinal mass

BLOOD GAS MEASUREMENTS Consider if severe/life-threatening features not responding treatment Normal or high pCO<sub>2</sub> = worsening asthma & respiratory failure Capillary blood gases give accurate pH + pCO<sub>2</sub> results ALWAYS check lactate

#### DIFFERENTIAL DIAGNOSIS

Anaphylaxis/Allergy Severe or Atypical Pneumonia Sepsis (esp. 个lactate) Hyperventilation Mediastinal Mass Foreign body Pulmonary oedema Inhalational injury





OXYGEN: Use high-flow oxygen via a tight-fitting face mask or nasal cannula at sufficient flow rates to achieve SpO2 ≥ 94%. High-flow humidified nasal cannula oxygen (eg optiflow, airvo or vapotherm) may be considered early, aiming for a flow 2 L/kg/min Max 50-60 L/min. **NEBULISERS:** Oxygen-driven nebulisation is recommended Salbutamol: **Under 5yrs** 2.5 mg Over 5yrs 5 mg Ipratropium bromide: Under 12yrs 250 micrograms **Over 12vrs** 500 micrograms Combine nebulised ipratropium bromide with  $\beta_2$  agonist to achieve significantly more bronchodilatation than  $\beta_2$  agonist alone. If poor response to initial dose of  $\beta_2$  agonist subsequent doses should be given in combination with ipratropium every 20 minutes for the first hour then 4 hourly (with ipratropium). STEROIDS: oral consider either dexamethasone OR prednisolone within one hour of presentation Dexamethasone: 0.6 mg/kg (max 16 mg) od for 3 days Prednisolone: 2 mg/kg od started within 1 hour presentation for 3-5 days Max 40 mg unless on maintenance steroids when max dose is 60 mg Hydrocortisone: 4 mg/kg 6 hourly intravenously (max 100 mg per dose) Always use iv hydrocortisone if vomiting or for those most severely affected. Benefits apparent within 3-4 hours. Oral and intravenous steroids are equivalent efficacy. Continue until clinically improved. Tapering is unnecessary unless course of steroids continues for > 14 days MAGNESIUM SULFATE BOLUS (UNLICENSED): for more information see page 8 Bolus: 40 mg/kg (max 2 grams) = 0.16 mmol/kg (max 8 mmol) intravenously over 20 min For easier prescribing / administration use banded doses according to patient weight see page 8 Do not wait for magnesium levels before giving first dose, toxicity rarely seen below level 4 mmol/L. Dose may be repeated in severe cases within 1-2 hours. SHORT INTRAVENOUS MAGNESIUM SULFATE INFUSION (UNLICENSED) : for more information see page 9 Dose: 50 mg/kg/hr for 4 hours (max. 8 gram/4 hr) may be used to treat acute severe exacerbations of asthma. NB If considering starting a magnesium infusion, please discuss with NWTS Clinically non-significant fall in BP (~ 5 mmHg) may occur following a bolus or during infusion MgSO<sub>4</sub>. If hypotensive, give 10 mL/kg fluid, ideally balanced crystalloid (eg Plasmalyte 148 or Hartmann's solution) or if not available 0.9% sodium chloride, and review. Consider if hypotension due to sepsis or hypovolaemia (2<sup>ry</sup> to increased insensible losses and poor intake). AMINOPHYLLINE: Loading: 6 mg/kg (max 500 mg) over 20 min (omit if on oral theophylline/aminophylline) Infusion: Under 12 years 1 mg/kg/hr Over 12 years 0.5-0.7 mg/kg/hr Doses should be adjusted according to plasma theophylline levels (see page 10) **INTRAVENOUS SALBUTAMOL:** Bolus (>2 years)15 microgram/kg over 5-10 minutes (max 250 microgram) See page 11 Infusion 1 microgram/kg/minute in severe refractory asthma NB MAX infusion 20 microgram/min (TOTAL dose NOT per kg) Monitor for hypokalaemia and signs toxicity. Reduce nebulised  $\beta_2$ -agonists to 4 hourly when on infusion. If evidence of salbutamol toxicity (see page 11), stop nebulised  $\beta_2$ -agonists and reduce rate infusion. Metabolic and lactic acidosis are worsened by hypovolaemia. Check blood gases + lactate min 6 hrly. MONITORING: ECG, SpO<sub>2</sub>, RR, BP, 4-6 hourly blood gases including lactate & minimum 12 hourly U&Es All patients receiving IV magnesium sulfate/ aminophylline /salbutamol must be admitted to hospital. Intravenous salbutamol and aminophylline cannot run together via same PVL.

Intravenous salbutamol and aminophylline cannot run together via same PVL. Either are compatible (via multi-tail connector or 3-way tap) with 0.9% sodium chloride + 5% glucose, Plasmalyte 148 or Plasmalyte 148 + 5% glucose Magnesium may be given via same PVL as salbutamol



*Guidelines for Management of Acute Severe Asthma in Children Over 2yrs* 



## **INTUBATION IN ACUTE SEVERE ASTHMA**

#### **IS A HIGH RISK PROCEDURE**

#### See NWTS intubation Guideline www.nwts.nhs.uk

Indications for Intubation	<ul> <li>Cardiac/Respiratory Arrest</li> <li>Exhaustion</li> <li>Hypoxia ie SpO<sub>2</sub> ≤ 92% despite escalation of treatment</li> <li>High CO<sub>2</sub> (&gt; 6 kPa) rare in acute asthma = sign of fatigue/exhaustion</li> <li>Acidaemia (pH ≤ 7.2)</li> <li>Altered sensorium ie agitation, confusion, decreased GCS</li> <li>Silent chest / poor air entry / inability to talk in short sentences</li> </ul>	<ul> <li>NB Asthma Severity may be difficult to assess:</li> <li>Tachycardia is universal with β<sub>2</sub> agonist</li> <li>Respiratory rate varies with respiratory drive +/- fatigue</li> <li>NB slow rate suggests fatigue</li> <li>Agitation or drowsiness may occur</li> <li>If any concerns ask for joint review by paediatric and anaesthetic consultants</li> </ul>	
	RISK	OPTIONS TO MITIGATE RISK	
	Low oxygen reserve Rapid desaturation Difficult CO <sub>2</sub> clearance	Most experienced available intubator Pre-oxygenate Apnoeic oxygenation using nasal cannulae 2L/kg/min high flow ideal or 0.2L/kg/min nasal specs Use largest fitting/cuffed ET tube SLOW respiratory rate	
Risks at Intubation	Relative hypovolaemia	Anticipate hypotension Good PVL / IO access Give 20mL/kg fluid bolus pre-induction Prepare DILUTE adrenaline IE take 0.1 mL/kg from Minijet syringe 1:10,000 adrenaline (using 3-way tap) Make this up to 10 mL 0.9% sodium chloride Use 1-2 mL aliquots to maintain BP	
	Delayed gastric emptying	Modified rapid sequence induction Eg ketamine + fentanyl + rocuronium Nasogastric tube ASAP after intubation	
Drugs for induction	Avoid histamine-releasing drugs if possible (atracurium, thiopentone, morphine) Use ketamine and/or fentanyl Volatile anaesthetic agent available for use immediately post-intubation (bronchodilator)		
Alternative bronchodilators: ONCE ventilated	Ketamine infusion Volatile anaesthetic agents Adrenaline 1:10,000 0.1mL/kg IV or via ETT can be used in extremis		
Ongoing sedation & management	Use either ketamine or fentanyl plus midazolam infusions Avoid morphine and atracurium as both cause histamine release Use low respiratory rate (with long expiratory phase) when hand ventilating to improve oxygenation + CO <sub>2</sub> clearance & reduce risk pneumothorax		



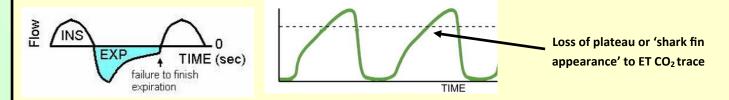


## Difficulties with Ventilation in Acute Severe Asthma REMEMBER HYPOXIA KILLS, HYPERCAPNOEA DOES NOT!

HIGH PEAK PRESSURES may cause barotrauma, pneumothorax, air leaks or  $\sqrt{}$  cardiac output

Strategies: Try PCV or square wave ventilation Limit Pmax (aim < 35-40 cmH<sub>2</sub>O) Permissive hypercapnia (ie tolerate pH ≥ 7.15; pCO<sub>2</sub> 7-14 kPa) Aim SpO<sub>2</sub> ≥ 94% Large, cuffed ETT will reduce resistance and minimal leak Keep muscle relaxed initially especially whilst high pCO<sub>2</sub>

**INCOMPLETE EXPIRATION** Slow emptying of alveoli causes poor gas exchange, progressive gas trapping and increasing residual volume



Strategies: Try slow respiratory rates (for age) + long expiratory times (I:E ratio ≥ 1:3) Allow completion of expiration ie flow to reach zero before next breath Manual decompression (disconnect ETT and manually compress chest)

**Volatile anaesthetic** eg sevoflurane can be used via anaesthetic machine/ventilator as an additional bronchodilator. Caution: watch for hypotension

**Physiotherapy** with 0.9% sodium chloride lavage may help but use a **slow** bagging rate

## Intrinsic PEEP

Aim to match extrinsic PEEP to intrinsic PEEP to reduce gas trapping ie 6-8 cmH<sub>2</sub>O

## **MUCUS PLUGGING**

Suction with 0.9% sodium chloride lavage 0.5 mL/kg max 5 mL via ETT This may make situation worse if inadequately sedated and not muscle relaxed An alternative: nebulised N-acetylcysteine

 mix 2.5 mL 20% solution N-acetylcysteine with 2.5 mL 0.9% sodium chloride, and then nebulise 2.5 mL of final concentration (or nebulise 2.5 mL 10% solution N-acetylcysteine)
 In extreme cases, Dornase Alpha (see BNFc) may improve severe mucous plugging.

**Discuss EARLY with NWTS** 



#### MANAGEMENT OF ACUTE WHEEZE IN CHILDREN UNDER 2 YEARS OF AGE

This guideline is not appropriate for use in children under age 2 years. In such children, a number of different diagnoses need to be considered and the response to treatment is highly variable. Such children should be managed on an individualised basis and early consultant involvement should be obtained.

NB Children < 2 years with clinical picture consistent with asthma / severe bronchospasm may respond better to magnesium sulfate and aminophylline rather than salbutamol (see bronchiolitis guideline)

	PATIENTS AT RISK OF NEAR-FATAL / FATAL ASTHMA
Severe asthma	Previous near-fatal asthma; History of anaphylaxis
	Previous admissions especially HDU or PICU +/- repeat attendances to A&E
	Requiring 3 or more classes of asthma medication
Plus	Adverse behavioural/psychological features; Learning difficulties
	Poor compliance; Failure to attend appointments
	Fewer GP contacts; Self-discharge from hospital
	Psychosis, depression, psychiatric illness or deliberate self harm
	Alcohol or drug abuse
	Obesity
	Looked after children
Have lower threshol	d for hospital admission + consider tertiary FU for children with above risk factors
	CRITERIA FOR REDUCING BRONCHODILATOR THERAPY
Normal	respiratory effort; Normal ability to speak; Reducing oxygen requirement
	DISCONTINUING INTRAVENOUS BRONCHODILATORS
Aminophylline:	Elimination half-life 3-5 hours. Reduce dose by 50% original dose every 6 hours
	After discontinuing infusion, aminophylline is cleared within 24 hours
Salbutamol:	Elimination half-life 4-6 hours. Reduce dose by 0.5 microgram/kg/min every 6 hours
	After discontinuing infusion, salbutamol will be cleared within 24 hours
<b>NB:</b> Substantial sys	temic absorption of salbutamol occurs via GI tract when given by inhalation or
nebulisers, so intrav	enous infusions should be discontinued before stopping nebulised salbutamol
Each patient should	receive nebulised $\beta_2$ agonists every 2 hours and nebulised ipratropium
bromide every 4 ho	urs whilst weaning off intravenous bronchodilators.
_	occur 24—48 hours after stopping either infusion so observe in hospital
Some patients with	particularly brittle asthma may require a slower weaning regime
	DISCHARGE PLANNING AFTER SEVERE ASTHMA ATTACK:
Check inhaler	technique
Start or review	v dosage of preventer treatment
Written asthm	na action plan for subsequent attacks with clear instructions on use bronchodilators

- and when to seek urgent medical attention if worsening symptoms.
- Contact GP to arrange primary care follow up within 48 hours
- Local paediatric (respiratory) review for those who required iv aminophylline and/or salbutamol.
- Refer all life-threatening or invasively ventilated patients to tertiary paediatric respiratory team



## **APPENDIX 1: ADDITIONAL DRUG INFORMATION MAGNESIUM**

**NEBULISED MAGNESIUM:** limited evidence benefit if used in severe asthma. No benefit in milder case. Dose: 150 mg per nebuliser (mix with salbutamol + ipratropium bromide) in 1st hour of treatment.

**INTRAVENOUS MAGNESIUM SULFATE BOLUS (UNLICENSED): first-line iv therapy**: safe and less likely to cause tachycardia. Some evidence that higher doses magnesium may be of benefit clinically but this is not currently recommended in the BTS guidelines. In practice repeating the dose 1-2 hours after initial dose is clinically safe. Discuss all patients requiring multiple doses with NWTS.

Magnesium sulfate 50% injection contains 500mg/mL magnesium sulfate.

**Dose: 40mg/kg over 20 minutes (max. dose 2g)** can be administered centrally or peripherally.

Always dilute 50% solution before administration (see below). Aim level: 1.5-2 mmol/L

A clinically non-significant fall in BP (~ 5 mmHg) may occur as magnesium sulfate causes vasodilation. If hypotensive, give 10 mL/kg fluid, ideally use balanced crystalloid (eg Plasmalyte 148 or Hartmann's solution) or if not available 0.9% sodium chloride, and then review.

Consider if hypotension may be due to sepsis or hypovolaemia (secondary to increased insensible losses and poor intake).

Contra-indications: Myasthenia gravis; Severe renal impairment

**Overdose:** Depends on size of overdose: progressive muscle weakness, significant hypotension and ultimately respiratory failure reported, but unlikely to occur if plasma levels are less than 4 mmol/L.

MAGNESIUM SULFATE BOLUS DOSE				
WEIGHT (kg)	DOSE Magnesium sulfate (40mg/kg)	VOLUME Magnesium sulfate 50%	Further DILUTION before administration	
5-5.9kg	200mg	0.4mL		
6-6.9kg	250mg	0.5mL		
7-7.9kg	300mg	0.6mL		
8-8.9kg	300mg	0.6mL		
9-9.9kg	350mg	0.7mL		
10-11.9kg	400mg	0.8mL		
12-13.9kg	500mg	1 mL		
14-15.9kg	550mg	1.1mL	ALWAYS further dilute	
16-17.9kg	600mg	1.2ml	required dose magnesium	
18-19.9kg	700mg	1.4mL	sulfate 50% up to	
20-21.9kg	800mg	1.6mL	20 mL with 0.9% sodium	
22-23.9kg	900mg	1.8mL	chloride before	
24-25.9kg	950mg	1.9mL		
26-27.9kg	1000mg	2 mL	administration	
28-29.9kg	1100mg	2.2mL		
30-34.9kg	1200mg	2.4mL		
35-39.9kg	1400mg	2.8mL		
40-44.9kg	1600mg	3.2mL		
45-49.9kg	1800mg	3.6mL		
50kg & above	2g	4mL		





## **MAGNESIUM SULFATE INFUSION**

SHORT INTRAVENOUS MAGNESIUM SULFATE INFUSION (UNLICENSED): safe and less likely to cause tachycardia than either aminophylline or salbutamol. Some evidence that high dose magnesium infusion may be of benefit clinically, and may be more effective in leading to improvement in symptoms than a bolus dose. Magnesium has a rapid onset of action (within minutes) and is rapidly eliminated (renal excretion). A high dose magnesium infusion is not currently recommended in the BTS guidelines. In practice, it has been shown to be safe and is used in other circumstances eg pre-eclampsia. Discuss all patients starting on a short magnesium infusion with NWTS.

## INFUSION DOSE: 50 MG/KG/HR FOR 4 HOURS TOTAL (MAX. 8 GRAMS/4 HR)

This can be administered peripherally or centrally.

NB adjust to ideal body weight if BMI >25 in order to avoid overdose

Always dilute 50% solution (500 mg/mL) before administration to 100 mg/mL

## Aim level: 1.5-2 mmol/L

Watch for hypotension, especially if dehydrated as magnesium sulfate causes vasodilation.

If hypotensive give 10 mL/kg fluid bolus, ideally balanced crystalloid (eg Plasmalyte 148 or Hartmann's solution). If not available use 0.9% sodium chloride. Review response following bolus.

Contra-indications: Myasthenia gravis; Severe renal impairment

**Overdose:** Depends on size of overdose: progressive muscle weakness, significant hypotension and ultimately respiratory failure reported, but unlikely to occur if plasma levels are less than 3.5 mmmol/L

## HOW TO MAKE UP & RUN A MAGNESIUM SULFATE INFUSION:

Magnesium Sulfate 50% (500 mg/mL): draw up 10 mL and make up to 50 mL with 5% glucose Final concentration = 10% solution = 100 mg/mL

Using this concentration: dose (mg) x wt (kg)/100 = mL /hr

WEIGHT (kg)	RATE (mL/hr)
5 kg	2.5
6 kg	3
7 kg	3.5
8 kg	4
9 kg	4.5
10 kg	5
12 kg	6
15 kg	7.5
17 kg	8.5
20 kg	10
22 kg	11
25 kg	12.5
27 kg	13.5
30 kg	15
32 kg	16
35 kg	17.5
37 kg	18.5
40 kg & above	20





## **AMINOPHYLLINE infusion for PERIPHERAL administration:**

- Draw up 500 milligrams aminophylline and add to 500mL 0.9% sodium chloride
- Final concentration = 500milligrams in 500mL i.e. 1milligram/mL aminophylline
- Aminophylline is compatible with up to 40mmol/litre Potassium chloride

LOADING DOSE: 6 MG/KG OVER 20 MINS (MAX DOSE 500MG).

**OMIT LOADING DOSE IF CURRENTLY ON ORAL THEOPHYLLINES AT HOME** 

INFUSION RATE:	1 MONTH -11 YEAR	1 MG/KG/HR = 1ML/KG/HR
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**12-17 YEARS** 

0.5-0.7 MG/KG/HR = 0.5-0.7 ML/KG/HR

Therapeutic monitoring: Use local guidance, but if none available check levels every 4-6 hours until stable and then every 24 hours

## Therapeutic range 10-20mg/l

Plasma levels correlate well with clinical effect but NOT with toxicity

<5mg/L

Response to monitoring:

Increase dose by 50% and recheck in 6 hours 5-15mg/L Continue. Recheck 24 hours

15-20mg/L Half infusion rate and recheck in 6 hours

>20mg/L STOP infusion and recheck levels in 6 hours. Restart at half the previous infusion rate once levels <15mg/l

NB Using peripheral concentration aminophylline ie 1 mg / mL			
Weight	0.5 mg/kg/hr	0.7 mg/kg/hr	1mg/kg/hr
5 kg	2.5 mL/hr	3.5 mL/hr	5 mL/hr
10 kgs	5 mL/hr	7 mL/hr	10 mL/hr
15 kg	7.5 mL/hr	10.5 mL/hr	15 mL/hr
20 kg	10 mL/hr	14 mL/hr	20 mL/hr
25 kg	12.5 mL/hr	17.5 mL/hr	25 mL/hr
30 kg	15 mL/hr	21 mL/hr	30 mL/hr
35 kg	17.5 mL/hr	24.5 mL/hr	35 mL/hr
40 kg	20 mL/hr	28 mL/hr	40 mL/hr
45 kg	22.5 mL/hr	31.5 mL/hr	45 mL/hr
50 kg	25 mL/hr	35 mL/hr	50 mL/hr
55 kg	27.5 mL/hr	38.5 mL/hr	55 mL/hr
60 kg	30 mL/hr	42 mL/hr	60 mL/hr
65 kg	32.5 mL/hr	45.5 mL/hr	65 mL/hr
70 kg	35 mL/hr	49 mL/hr	70 mL/hr





## **SALBUTAMOL** infusion

**Recommendation:** always pause and reassess patients after salbutamol bolus dose, as some patients will significantly improve and not require a salbutamol infusion.

## Making up PERIPHERAL SALBUTAMOL infusion:

- Draw up 10 mg salbutamol (IE 10mL salbutamol 1 mg/mL)
- Make up to 50mL with 5% glucose or 0.9% sodium chloride
- Final concentration = 10 mg in 50 mL i.e. 200 micrograms/mL salbutamol

BOLUS infusion rate 2-17 years: 15microgram/kg (MAX 250 microgram) bolus over 5-10 mins

Draw up required dose and dilute to a final volume 5 mL with sodium chloride 0.9% or glucose 5%

BOLUS dose 1-23 months: 5microgram/kg (MAX 250 microgram) over 5-10 min

Draw up required dose and dilute to a final volume 5 mL with sodium chloride 0.9% or glucose 5%

NB less likely to effectively relieve bronchospasm than when used to treat an older child.

Magnesium sulfate or aminophylline more likely to be effective in this age group

## **PERIPHERAL infusion rate: 0.3mL/kg/hr = 1 microgram/kg/minute.**

## CONTINUOUS IV INFUSION: 0.5-1 MICROGRAM/KG/MINUTE

## MAX INFUSION RATE: 20 MICROGRAM / MINUTE (TOTAL DOSE)

NB Using peripheral concentration salbutamol ie 200 microgram/mL				
Weight	1 microgram/kg/min	2 microgram/kg/min		
10kg	3mL/hr	6 mL/hr		
15kg	4.5mL/hr	6 mL/hr		
ALL pts ≥ 20kg	6mL/hr	6 mL/hr		

Maximum dose = 20 micrograms/min = 6mL/hr

This is standard infusion rate for any patient greater than or equal to 20 kg

No clinical benefit salbutamol infusion rates greater than total dose 20 microgram / minute. NB high doses are associated with an increased risk of salbutamol toxicity.

Signs of salbutamol toxicity:

Hypokalaemia	Hyperglycaemia	Agitation
Tachycardia	Tachyarrhythmia eg SVT	
Metabolic acidosis	Raised lactate	

NB If evidence of salbutamol toxicity, stop nebulised  $\beta_2$ -agonists and reduce rate infusion.

Increasing tachypnoea whilst on salbutamol infusion may indicate toxicity and metabolic acidosis rather than acute worsening of asthma

RAISED lactate is likely to be secondary to salbutamol, but always reassess patient as it may be due to other causes eg sepsis or hypovolaemia, especially if lactate was high on admission (pre-salbutamol)

## MANAGEMENT OF SVT FOLLOWING SALBUTAMOL INFUSION

SVT has been reported in those receiving salbutamol loading doses and infusions at the higher dose range (ie more than 20 microgram/min total or 2 microgram/kg/min).

Adenosine may cause bronchospasm in known asthmatics: only use with caution in acute severe asthma.





## **References**

Magnesium sulfate for treating exacerbations of acute asthma in the emergency department (Review) Griffiths B, Kew KM. Cochrane Database Syst Rev. 2016 Apr 29;4

**Inhaled magnesium sulfate in the treatment of acute asthma (Review).** Knightly R, Milan SJ. Cochrane Database Syst Rev 2017, Issue 11. CD003898

MAGNEsium Trial In Children (MAGNETIC): a randomised, placebo-controlled trial and economic evaluation of nebulised magnesium sulphate in acute severe asthma in children. Powell CV, Kolamunnage-Dona R. MAGNETIC study group. Health Technol Assess. 2013 Oct;17(45):v-vi, 1-216.

**Clinical pharmacokinetics of magnesium sulphate in the treatment of children with severe acute asthma.** Rower JE, Liu X, Yu T, Mundorff M, Sherwin CM, Johnson MD. Eur J Clin Pharmacol. 2017 Mar;73(3):325-331. doi: 10.1007/s00228-016-2165-3. PMID:27909740

**BTS/SIGN British Guideline on the Management of Asthma—a National Clinical Guideline** May 2008, Revised September 2019

Management of Acute Severe Asthma in children (aged >2years) version 4 Royal Manchester Children's Hospital, CMFT. September 2015. Originated by Rachael Barber, PICU Consultant, Clare Murray, Respiratory Consultant.

**Intravenous salbutamol for childhood asthma: evidence-based medicine?** Starkey E, Mulla H, Pandya H, Archives of Disease in Childhood 2014;99:873-877.

**High-Dose Magnesium Sulfate Infusion for Severe Asthma in the Emergency Department: Efficacy Study.** Irazuzta JE, Paredes F, Pavlicich V, Domínguez SL. Pediatr Crit Care Med. 2016 Feb;17(2):e29-33.

Magnesium Sulfate infusion for acute asthma in the emergency department Enriquerazuzta J, Chiriboga N. Jornal de Pediatria 93(S1) 2017: 19-25

**Emergency presentation and management of acute severe asthma in children** Oymar K, Halvorsen T Scand J Trauma Resusc Emerg Med 2009; 17:40: 5-11

Adrenaline (epinephrine) compared to selective beta-2-agonist in adults or children with acute asthma: a systematic review and meta-analysis. Baggott, C., Hardy, J., Sparks, J., Sabbagh, D., Beasley, R., Weatherall, M. and Fingleton, J. *MedRxiv* 2021.

## Lexicomp 18th Edition

## FOR DRUG DOSES:

British National Formulary for Children 2019-2020

www.crashcall.net



Guidelines for Management of Acute Severe Asthma in Children Over 2yrs

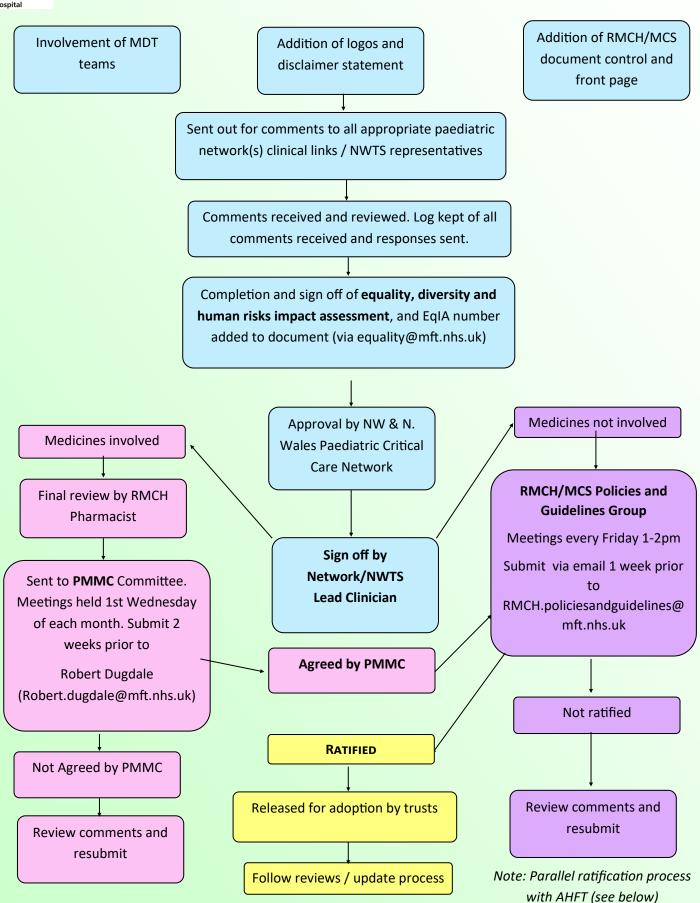
Appendix 2

Ratification of Guidelines with Host Organisation (MFT)





## Alder Hey Children's MHS NHS Foundation Trust





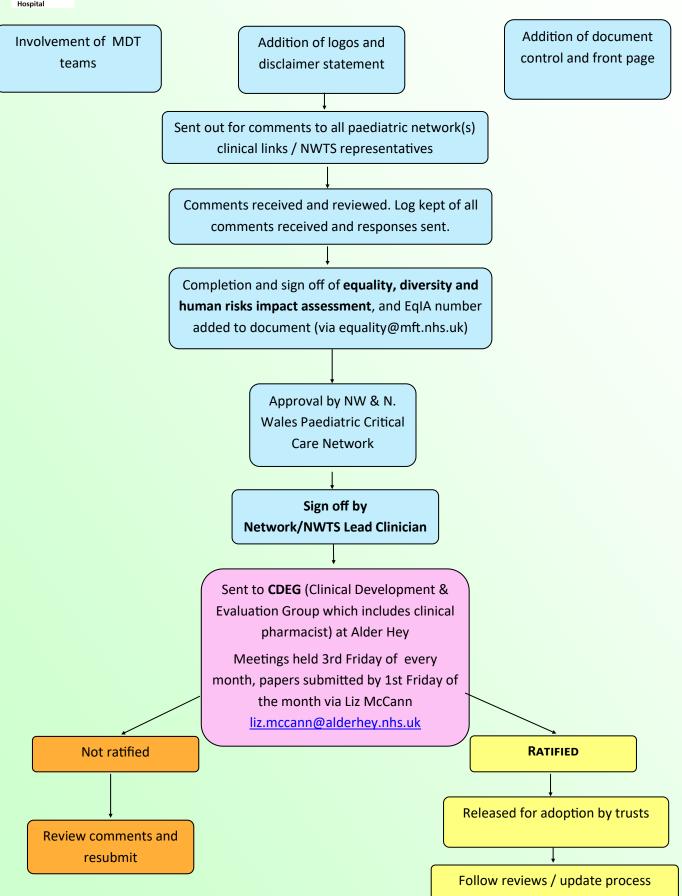
Guidelines for Management of Acute Severe Asthma in Children Over 2yrs

**Ratification of Guidelines with Alder Hey** 





Alder Hey Children's MHS NHS Foundation Trust





*Guidelines for Management of Acute Severe Asthma in Children Over 2yrs* 



## **Resources**

www.crashcall.net - for intubation drugs / sedation regime

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Please visit NWTS website for the most up to date version of this guideline: www.nwts.nhs.uk



