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A chronic inflammatory disorder of the airways, associated with variable airflow obstruction

4 Key Components

- 1. Bronchial constriction
- 2. Inflammation
- 3. Bronchial hyper-responsiveness
- 4. Airway remodeling

Bronchoconstriction

- Predominant physiologic event --> symptoms
- Stimulus-> bronchial smooth muscle contraction -> acute narrowing of bronchi
- Allergen-induced: IgE-dependent release of mediators (histamine, tryptase, leukotrienes and prostaglandins) from mast cells -> direct contract airway smooth muscle
- Aspirin + NSAIDs: non-IgE-dependent response
- Other irritants (exercise, cold air, etc): mechanisms less well defined, intensity related to underlying inflammation



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Inflammation

- Inflammatory cell infiltration contributes to airway hyperresponsiveness, airflow limitations, symptoms and disease chronicity – patterns may differ which has implications for phenotypic differences
 - Neutrophils (esp in sudden-onset, fatal exacerbations; occupational asthma, and tobacco smoke exposure)
 - o Eosinophils
 - Lymphocytes
 - Mast cell activation
 - Resident cells
 - o Epithelial cell injury
- Mucus hypersecretion, inspissated mucus plugs and edema of airways
- Persistent inflammation in some patients will lead to structural changes, including sub-basement fibrosis, mucous hypersecretion, injury to epithelial cells, smooth muscle hypertrophy, and angiogenesis, i.e. airway remodeling





Hyperresponsiveness

- Exaggerated bronchoconstrictor response to a wide variety of stimuli
- Mechanisms: inflammation, dysfunctio nal neuroregulation and structural changes

Airway Remodeling:

- Permanent structural changes can occur associated with loss of lung function
- Thickening of subbasement membrane, subepithelial fibrosis, airway smooth muscle hypertrophy and hyperplasia, blood vessel proliferation and dilation, and mucus gland hyperplasia and hypersecretion





WHY DEX?

- Cheaper \$\$
- Palatability (kids like it more)
- 5-6 x more potent, half-life 36-72hrs -> Improved compliance with less dosing
- Fewer side effects such as vomiting



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WHAT'S THE PROOF? DEX VS PRED

- Qureshi 2001: prospective non-blinded RCT no significant different in relapse rate (6.9 v 7.4%), no difference in admission rate when stratified by asthma severity (12 v. 11%), total time spent in ED, or severity score on discharge
 - Pred: more vomiting, less compliance, More missed > 2 days of school
- Andrews 2012: More cost effective, especially if doses provided by ED
- Keeny 2014: No difference in asthma scores during ED visit, # Alb treatments needed, rates of hospitalization, or relapse visits
- Cronin 2016: mean pediatric respiratory assessment measures (PRAM), beta-agonist use, admissions, admission LOS, unscheduled return visits within 14 days, quality of life indicators not significantly different; did see difference in use of further steroids and study did not include "severe" exacerbations
- Watnick 2016: Relative risk reduction of 35% for relapse visit in a retrospective analysis
- Paniagua 2017: no difference in persistence of symptoms, quality of life, admissions, ED returns or vomiting at day 7; greater adherence with Dex
- PECARN 2022: no difference in rate of return visits for continued or worsening symptoms between 1 and 2 doses of dexamethasone
- Hoefgen 2022: initial steroid choice not associated with 30 day re-utilization after hospitalization

						DEX	-	PRE)		Risk Ratio		Risk Ratio			
Cai 2021	:				Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% C		M-H, Fixed, 95% CI			
					1.3.1 Hospital admiss	ions										
					Altamimi 2006	6	67	9	67	13.3%	0.67 [0.25, 1.77]				N	
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		- h I			Gordon 2007	17	88	19	93	27.3%	0.95 [0.53, 1.70]					
k eg im en :	S VI	abi	e		Paniagua 2017	10	281	7	276	10.4%	1.40 [0.54, 3.63]					///11/
Alternati	Ve	Ιο ·	5 D (n v r	Scarfone 1995	13	56	17	55	25.3%	0.75 [0.40, 1.39]					
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red					Total events	64		68								
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					Test for overall effect:	2 = 0.34 (1	P = 0.73)								
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kevisit O	L K C	e l a	pse	Arrer	Gries 2000	1	15	3	17	3.0%	0.38 [0.04, 3.26]			(A)		
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Jischarg	~				Paniagua 2017	13	281	9	276	9.7%	1.42 [0.62, 3.27]		- - -			
					Parikh 2015	7	642	13	642	13.9%	0.54 [0.22, 1.34]					
					Qureshi 2001	20	272	18	261	19.6%	1.07 [0.58, 1.97]					
					Scarfone 1995	0	56	9	55	10.2%	0.05 [0.00, 0.87]					
					Subtotal (95% CI)		1616		1592	100.0%	0.94 [0.71, 1.24]		•			
					Total events	89		93								
					Heterogeneity: Chi ² = 1	0.43, df =	8 (P = 0	0.24); l ² =	= 23%							
V/a mailin a					Test for overall effect: 2	Z = 0.43 (F	P = 0.67)								
vomining					1 3 3 Hospital admiss	ions after	r rolans	•								
	DEX	PRED		Risk Ratio	Risk F	Ratio	relapor	ŭ 1	67	1.5%	3 00 [0 32 28 12]					
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Cronin 2016	0 123	14	122 21.1%	0.03 [0.00, 0.57]				.4	261	66.1%	1.18 [0.82, 1.69]		-			
Gordon 2007	0 88	6	93 9.2%	0.08 [0.00, 1.42]				7	55	25.2%	0.75 [0.40, 1.39]					
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I PRATROPIUM BROMMER O Cureshi 1998: double blinded RCT adding 500mcg IB with 2nd and 3rd albuterol doses O Cureshi 1998: double blinded RCT adding 500mcg IB with 2nd and 3rd albuterol doses O Supficant reduction in admissions by 15% for severe pai O Supficant reduction in asthma scores and reduced oxygen need in severe patients O Supficant reduction in asthma scores and reduced oxygen need in severe patients O Supficant reduction in 1-3 divided doses in the ED setting leads to greater improvement in lung function (PFTs, clinical scores, need for oxygen, need for additional SABAs) and fewer side effects (nausea and tremor) O Watt 2015: Adding IB not significantly associated with reduced admissions rates; increases acheficial effects appear to decline with additional dosing (Qureshi 1998, Craven 2001, Criftins 2013) Methoded antichalmed antichalmed pratropulm on the hospitalization rates of children with asthma. New Englend Journal of Meticne, 339(15), 103-103, Methoded antichalmed antichalmed pratropulm on the hospitalization rates of children with additional dosing (Qureshi 1998, Craven 2001, Criftins 2013) Methoded antichalmed pratropulm on the hospitalization rates of children with additioned addoses and the tacting beats-agoints for initial treatment of acute asthma in children. Cedvinae Database Syst Rev. 2013 Aug 2014 Combined Instale antichalmed pratropulm on the hospitalization rates of children with additioned bases syst Rev. 2013 Aug 2014 Combined Instale antichalmed acute displaced significant for initial treatment of acute asthma in children. Cedvinae Database Syst Rev. 2013 Aug 2014 Combined Instale antichalmed reated cases and the romoria of acute asthma in children. a single-bilinded rundomised contended 2015 Combined Instale antichalmed reated acute data prestone agoints for initial treatment of acute asthma in children. Expland Aug 2014 Combined Instale antichalmed acute database in the Instale reatment of acute asthma in children. a single-bilinded rundomised combined 2015 Combined













- Evidence suggests delivers 5-6 fold greater aerosol compared to jet nebulizer
- Less volume used, less left behind
- 85% patients need only one treatment
- BUT one time use
- Dunne & Shortt: Prospective QI project. Total administered dose of albuterol in the VMN group was lower (no more than 5mg albuterol). The VMN was associated with fewer admissions to the hospital (28.1% vs 41.4%), shorter length of stay in the ED (37min reduction) and a reduction in albuterol dose. The device type was a predictor of discharge, disposition and amount of drug used.





DEVELOPING EVIDENCE



- Prospective chart review
- 3x less drug used in order to achieve D/C with VMN (3.72 vs. 11.23mg mean dose)
- 89 minute reduction in average ED LOS in VMN vs. JN group
- Admission rates were 41% lower in the VMN group vs. JN group
- Return to ED in 48-hour rates were 42% lower in VMN group vs. JN group

children'shealth? Children's Medical Center Dallas

- Single blinded RCT, all treated until received mild assessment score and dispositioned
- Statistical significance in initial AS score with VMN group average at 9 and JN group average at 8
- 33% reduction in number of treatments needed (VMN 2 [1-3], JN 3 [2-5])
- 48% reduction in admission rates for severe AS in the VMN group vs. JN group
- Patients achieved symptom control 23 minutes faster in VMN group vs. JN group



INPATIENT MANAGEMENT

Steroids Again

- Hemani 2021: hospitalized children with mild-to-moderate exacerbations have significantly shorter LOS when starting Dex on admission opposed to Pred. Studied 1,410 children with difference found when steroids started after hospitalization. LOS did not differ when steroids started before admission. PICU transfers, ED revisits, and readmissions were uncommon.
- Tyler 2019: Using interrupted time series data, found no difference for LOS, total charges, or ICU transfers pre and post-implementation of dexamethasone protocol in ED. Slight trend in ICU transfers post.
- Cotter 2020: survey of hospitalist practices. Patient received dexamethasone in the ED, then inpatient care was variable - 44% of Hospitalists continued dex, 14% switched to pred, 2% give no additional steroids, and 40% tailors based on scenario. Hospitalists more likely to continue dex than pulmonologists (61 vs 15%, p <0.001). "Providers often revert to prednisone, especially in severe asthma exacerbations, possibly due to experience with prednisone and limited research on dexamethasone in the inpatient setting."
- Nelipovich 2022: survey hospitalist MDs and APPs in 2019 and 2021. Wide disagreement with using dex in both surveys, but self-reported increase in use in 2021. Moderate agreement with prescribing dex for poor tolerance or compliance. Moderate agreement with prescribing prednisone with higher severity of baseline asthma or current exacerbation.

Sunita Ali Hemani, Brianna Glover, Samantha Ball, Willi Rechler, Martha Wetzel, Nicole Hames, Elan Jenkins, Patricia Lantis, Anne Fitzpatrick, Sarah Varghese; Dexamethasone Versus Prednisone in Children Hospitalized for Acute Asthma Exacerbations. Hosp Pediatr November 2021; 11 (11): 1263–1272. https://doi.org/10.1542/hpeds.2020-004788 In Children Hospicalized for Acute Astimat Exactrolations. *Hosp redular November 2021*; 11:111;1263-1272. https://doi.org/10.1542/hpeds.2021-004765 Amy Tyler, Jillian M. Cotter, Angela Moss, Irina Topoz, Amanda Dempsey, Jennifer Reeses, Stanley Szeffer, Heather Hoch; Outcomes for Pediatric Asthmatic Inpatients After Implementation of a Emergency Department Dexamethasone Treatment Protocol. *Hosp Pediatr February* 2019; 9 (2): 92–99. https://doi.org/10.1542/hpeds.2018-0099 Cotter JM, Tyler A, Reese J, Ziniel S, Federico MJ, Anderson IIi WC, Kupfer O, Szeffer SJ, Kerby G, Hoch HE. Steroid variability in pediatric inpatient asthmatics: survey on provider preferences of dexamethasone versus prednisone. J Asthma. 2020 Sep;57(9):942-948. doi: 10.1080/02770903.2019.1622713. Epub 2019 Jun 12. PMID: 31113252; PMCID: PMC8086174. atients After Implementation of an vich S, Porada K, Vepraskas S, Soung P, Chou E. Current Practice and Rationale of Prescribing Dexamethasone for Pediatric Patients Hospitalized for Asthma. WMJ. 2022 Apr;121(1):30-35. PMID: 35442576













	STANDARDIZATION TOOLS
GUIDELINES	National Statements that contain recommendations based on evidence from rigorous systematic review and synthesis of published literature
P A T H W A Y S	Local Translation of guidelines, latest literature, specialty preferences and knowledge of resources into practice. Provides series of recommendations to the multidisciplinary team to reduce unnecessary variation in care.
PROTOCOLS	Role Specific Set of assessments and interventions constrained by rules or algorithms that define the care of a specific patient condition by a delegate (RT, RN, etc) initiated by a provider order



Reducing Variation In EDs

- ED pathway use significantly decreases time to corticosteroid administration (45 vs 29min to 20min year 2, p < 0.0001) (Lucia 2020)
- Guidelines reduce ED LOS for treat-and-release patients (3.9 v. 3.3hrs), hospital LOS (1.5 v. 1.3 days), admission rates from ED (23.5% v. 18.8%), ICU admissions (23% v. 13.2%) and total charges (\$4,457 v. \$3,651) (Johnson 2018)
- Children in General EDs compared to Pediatric EDs
 - Less likely to receive corticosteroids in compliance with NIH guidelines (Miller 2015)
 - More likely to receive unnecessary testing and treatment (CXR and antibiotics)
 - Children's ED have higher compliance with all 3 goals (42% v. 31%) but overall only 34% of visits were concordant with guidelinebased care (Hudgins 2021)
 - Guideline implementation increased utilization of steroids and decreased time to administration in community EDs (Walls 2017)

Miller AG, Breslin ME, Pineda LC, Fox JW. An Asthma Protocol Improved Adherence to Evidence-Based Guidelines for Pediatric Subjects With Status Asthmaticus in the Emergency Department. Respir Care. 2015
Dec;60(12):1759-64. doi: 10.4187/respcare.04011. Epub 2015 Jun 23. PMID: 26106203.
Hudgins JD, Neuman MI, Monuteaux MC, Porter J, Nelson KA. Provision of Guideline-Based Pediatric Asthma Care in US Emergency Departments. Pediatr Emerg Care. 2021 Oct 1;37(10):507-512. doi:
10.1097/PEC.0000000001706. PMID: 30624420
Walls TA, Hughes NT, Mullan PC, Chamberlain JM, Brown K. Improving Pediatric Asthma Outcomes in a Community Emergency Department. Pediatrics. 2017 Jan;139(1):e20160088. doi: 10.1542/peds.2016-0088.
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Lucia D, Cain J, Porter A, Sagar M, Blazovic S, Finley L, Mallett L. Pediatric asthma pathway in the emergency room. Proc (Bayl Univ Med Cent). 2020 Aug 21;34(1):40-43. doi: 10.1080/08998280.2020.1801110.
PMID: 33456142; PMCID: PMC7785189.
Johnson DP, Arnold DH, Gay JC, Grisso A, O'Connor MG, O'Kelley E, Moore PE. Implementation and Improvement of Pediatric Asthma Guideline Improves Hospital-Based Care. Pediatrics. 2018
Feb:1/11/2):e20171630. doi: 10.15/2/peds.2017-1630. PMID: 20367203

Reducing Variation In EDs

Seattle Children's (Rutman 2016): retrospective study of QI initiative comparing pre and post asthma pathway implementation

- Had Respiratory Clinical Score (CS) in place since 2002 with good interobserver agreement and pre-existing pathway
 Noted 90% of patients with high score (9-12) after 1 hr were ultimately admitted yet had variable and prolonged LOS in the ED
- Noted 90% of patients with high score (9-12) after 1 hr were ultimately admitted
- Developed respiratory score-based admission criteria after 1 hr of treatment
 Median ED LOS and time to bed request decreased by 30 mins without a change in % admissions, median inpatient LOS or PICU admissions







Wazeka A, Valacer DJ, Cooper M, Caplan DW, DiMaio M. Impact of a pediatric asthma clinical pathway on hospital cost and length of stay. Pediatr Pulmonol. 2001 Sep;32(3):211-6. doi: 10.1002/ppul.1110. PMID: 11536450. Johnson KB, Blaisdell CJ, Walker A, Eggleston P. Effectiveness of a clinical pathway for inpatient asthma management. Pediatrics 2000 Nov; 106(5): 1006-12. Kelly CS, Andersen CL, Pestian JP, Wenger AD, Finch AB, Strope GL, Luckstead EF. Improved outcomes for hospitalized asthmatic children using a clinical pathway. Ann Allergy Asthma Immunol. 2000 May;84(5):509-16. doi: 10.1016/S1081-1206(10)62514-8. PMID: 10831004. McDowell KM, Chathurn RL, Myers TR, Offsordan MA, Kercsmar CM. A cost-saving algorithm for children hospitalized for status asthmaticus. Arch Pediatr Adolesc Med. 1998 Oct;152(10):977-84. doi: 10.1001/archpedi.152.10.977. PMID: 9790607.



Inpatient Standardization

PATHWAYS FOR IMPROVING INPATIENT PEDIATRIC ASTHMA (PIPA)

National QI project through the AAP's Value in Inpatient Pediatrics Network, now known as the Pediatric Acute & Critical Care (PACC) Quality Network, conducted in 2017

- Provided toolkit for implementing pathways and order sets
 - Guidance on dosing MDIs
 - o Guidance on titration by RNs or RTs
 - o Hospital discharge criteria
 - o Reminders for Tobacco Smoke Exposure (TSE) screening and referrals
 - *Set aims for all participating hospitals, ED:

Inpatient:

- Decrease overall usage of CXR to 15% • Achieve 90% documented compliance with assessing
- severity of exacerbations at triage
- Increase proportion eligible children who receive systemic corticosteroids within 60 mins of arrival to 50%
- Decrease inpatient LOS by 10%
 - Increase early transition to administering bronchodilator via MDI by 50%
 - Decrease prescription Abx at discharge to 10%
- Achieve 90% compliance with screening for secondhand smoke exposure Achieve a 50% increase in documentation of caregiver referral to smoking cessation resources

AP

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Sunitha V. Kaiser, Brittany Jennings, Jonathan Rodean, Michael D. Cabana, Matthew D. Garber, Shawn L. Ralston, Bernhard Fassl, Ricardo Quinonez, Joanne C. Mendoza, Charles E. McCulloch, Kavita Parikh; Pathways for Improving Inpatient Pediatric Asthma Care (PIPA): A Multicenter, National Study. *Pediatrics* June 2020; 145 (6): e20193026. 10.1542/peds.2019-3026 Gupta N, Cattamanchi A, Cabana MD, Jennings B, Parikh K, Kaiser SV. Implementing pediatric inpatient asthma pathways. J Asthma. 2021 Jul;58(7):893-902. doi: 10.1080/02770903.2020.1741612. Epub 2020 Mar 18. PMID: 32160068.

Asiser SV, Lam R, Cabana MD, Bekmezian A, Bardach NS, Auerbach A, Rehm RS. Best practices in implementing inpatient pediatric asthma pathways: a qualitative study. J Asthma. 2019;1–11. doi:10.1080/02770903.2019.1606237.

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Value in Inpatient **Pediatrics** Network

from the revement Innovation Net ican Academy of Pediatri

STANDARDIZING IN COMMUNITY SETTINGS

MCDANIELS 2019 Key determinants of pathway

- Building implementation structure
- Engaging providers

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- Addressing organizational and resource limitations
- Devising implementation solutions with external facilitators

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High performing

- Dedicated local champions to provide consistent reminders of evidence-based practice and ongoing education
- Modify EMR tools
- o Collaborative culture receptive to practice change and firm expectations to follow evidence-
- based practices

- Utilized standardized asthma score, RT driven albuterol protocol, order sets/EMR tools, and targeted education
- aLOS decreased from 2.9 to 2.3d
- Variable direct cost savings of
- \$1,543 per case No increase in admissions

McDaniel CE, Jeske M, Sampayo EM, Liu P, Walls TA, Kaiser SV. Implementing Pediatric Asthma Pathways in Community Hospitals: A National Qualitative Study. J Hosp Med. 2020 Jan 1;15(1):35-41. doi: 10.12788/jhm.3296. Epub 2019 Sep 18. PMID: 31532746. Jaladanki, Sravya et al. "Strategies for Sustaining High-Quality Pediatric Asthma Care in Community Hospitals." *Health services research : HSR*. 57.1 (2022): 125–136. Web. Bartlett KW, Parente VM, Morales V, Hauser J, McLean HS. Improving the Efficiency of Care for Pediatric Patients Hospitalized With Asthma. Hosp Pediatr. 2017 Jan;7(1):31-38. doi: 10.1542/hpeds.2016-0108. Epub 2016 Dec 8. PMID: 27932381.







