Incorporating New Evidence into Stewardship Interventions for CAP



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Outline

- Discuss how a guideline and intervention focused on treatment selection and duration was created for CAP
- Review recent data regarding diagnosis and treatment of CAP
- Discuss how recent data and guideline recs have been incorporated into updated CAP guideline.



2016 CAP Care Pathway Development

- Develop and implement single "Best Practice" pathway for CAP
- Do away with HCAP
- Duration of therapy
- Reducing unnecessary atypical coverage
- Early IV to PO conversion

- Assess impact of Pathway on:
 - Intravenous antibiotic duration
 - Length of stay
 - Costs
 - Balancing Measures

CAP: Defining our terms

- Community Acquired Pneumonia (CAP)
 - With or without risk factors for drug resistance
 - HCAP dinical criteria did not distinguish patients with drug resistant pathogens
 - HCAP HGA Regerided at reation entridich a patiem prove mortality
- who: HCAR guided that apply did had and to over-us Healthcare-Associated Pneumonia Does Not
 - resides in a long-term care facility
 - received recent IV antibiotic therapy / chemotherapy / wound care within last 30 days
 - attended a hemodialysis center within last 30 days ٠

Accurately Identify Potentially Resistant Pathogens: A Systematic Review and Meta-Analysis

James D. Chalmers,¹ Catriona Rother,¹ Waleed Salih,¹ and Santiago Ewig²

Chalmers, Clin Infect Dis 2014

Troitino, Lung 2013 Attridge, Eur Respir J 2011 Postma, NEJM 2015

CAP: Drug resistant pathogens

- HCAP criteria did not distinguish CAP patients with drug resistant pathogens (DRP) from those with sensitive pathogens
- Recently published DRP prediction tool shows promise
- DRIP score >4 has 76% sensitivity and 91% specificity for drug resistant pathogen





Derivation and Multicenter Validation of the Drug Resistance in Pneumonia Clinical Prediction Score

[©] Brandon J. Webb,^a Kristin Dascomb,^a Edward Stenehjem,^a Holenarasipur R. Vikram,^b Neera Agrwal,^c Kenneth Sakata,^d Kathryn Williams,^e Bruno Bockorny,^f Kavitha Bagavathy,^f Shireen Mirza,^f Mark Metersky,^g Nathan C. Dean^h

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Characteristic	No. of points
Major Risk Factors	
Antibiotic use within previous 60 days	2
Residence in LTCF	2
Tube Feeding	2
Prior infection with DRP (1 year)	2
Minor Risk Factors	
Hospitalization in previous 60 days	1
Chronic pulmonary disease	1
Poor functional status	1
Gastric acid suppression	1
Wound care	1
MRSA colonization (1 year)	1

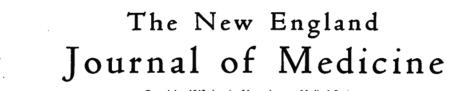
Webb BJ, Antimicrob Agents Chemother. 2016

CAP: Antibiotic duration

The Journal of the American Medical Association

Published Under the Auspices of the Board of Trustees

	RICAN MEDICAL ASSOCIATION MARCH 4, 1944
THE CLINICAL USE OF PENICILLIN OBSERVATIONS IN ONE HUNDRED CASES MARTIN HENRY DAWSON, M.D. AND GLADYS L. HOBBY, PH.D.	of cases of this disease was therefore abandoned unti- such time as larger supplies might become available. In the light of subsequent work it became obvious that the amount of penicillin given in this early group of cases was totally insufficient to secure a significant result. During this stage of the investigation 3 cases of acute pneumococcic endocarditis came under observation



Copyright, 1945, by the Massachusetts Medical Society

Volume 232

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Number 26

TREATMENT OF PNEUMOCOCCAL PNEUMONIA WITH PENICILLIN*

MANSON MEADS, M.D., † H. WILLIAM HARRIS, M.D., ‡ AND MAXWELL FINLAND, M.D.§ WITH THE TECHNICAL ASSISTANCE OF CLARE WILCOX

BOSTON

"...satisfactory results with 1.5-2 days of antibiotics"

- 2007 IDSA CAP guidelines: ~5 days until clinical stability
- One randomized controlled trial supports 3 days in select patients
- Durations < 3 days: limited data

Average duration of therapy (including patients in shock) was 4-5 days

Evidence-Based Interventions

• Atypical coverage

- Atypical infection uncommon
- NO impact on survival or clinical efficacy with empirical atypical coverage in hospitalized floor patients
- 25% of *S. pneumoniae* are macrolide-resistant
- β-lactam monotherapy recommended in CAP guidelines in other countries

IV to PO conversion

• Time to resolution of symptoms and relapse similar with IV vs. PO in nonsevere CAP and with rapid PO conversion in severe CAP

> Musher DM, et al. Clin Infect Dis 2017; 65(10);1736-44. Wiersinga WJ, et al. Neth J Med. 2018 Jan; 76(1)4-13. Eliakim-Raz N, et al. Cochrane Database Syst Rev, 2012; 26(9);CD00418 Castro-Guardiola A, et al. Am J Med 2001; 111:367-74.

How It Works

BestPractice Advisory - Jett, Joan Care Guidance (1) $\hat{}$ If this ABx is for treatment of Pneumonia, click "Open order set" for the appropriate Orderset based on the patient's CURB score. If this antibiotic is not for Pneumonia, click "Dismiss" to continue ordering. CURB-65 Calculation Confusion Yes = 1No = 0BUN > 19 mg/dL (> 7 mmol/L)Yes = 1No = 0Respiratory Rate ≥ 30 Yes = 1No = 0SBP < 90 mmHg or DBP \leq 60 mmHg No = 0Yes = 1No = 0Age ≥ 65 Yes = 1CURB-65 Total 0 - 1: Consider Outpatient Care CURB-65 Total 2: Consider Inpatient Admission CURB-65 Total 3+: Consider Inpatient ICU Admission Remove the following orders? azithromycin (ZITHROMAX) injection Remove Keep Intravenous, Starting Today at 1059, Routine Apply the following? Do Not Open Pneumonia Orderset for Inpatient Care Preview Open Order Set Open Order Set Do Not Open Pneumonia Orderset for Outpatient Care Preview ✓ Accept Dismiss

*Fires when chest x-ray and antibiotics both ordered...embedded into workflow



- Do away with HCAP
- Duration of therapy
- Reducing unnecessary atypical coverage
- Early IV to PO conversion

CAP Treatment Options

Icor (CURB 65 score 0-2 / Drip score less than 4) - For most patients (not at an increased risk for drug-resistant pathogens)

Antibiotics

Preferred Antibiotics

azithromycin (ZITHROMAX) 500 mg in sodium chloride 0.9 % 250 mL IVPB 500 mg, Intravenous, at 250 mL/hr, Administer over 60 Minutes, Once, Today at 1000, For 1 dose, STAT

And

cefTRIAXone (ROCEPHIN) 2 g in sodium chloride 0.9% IVPB Mini-bag Plus 2 g, Intravenous, at 200 mL/hr, Once, Today at 1000, For 1 dose, STAT

And

cefuroxime axetil (CEFTIN) tablet 500 mg 500 mg, Oral, 2 times daily, First Dose Tomorrow at 1000, For 8 doses Routine

Labs

Streptococcus Pneumoniae Antigen, Urine Once - Routine - Lab First occurrence Today at 0921, Urine, Urine-General Collection Method Override: Unit Collect

Legionella Pneumophilia Antigen, Urine Once - Routine - Lab First occurrence Today at 0921, Urine Collection Method Override: Unit Collect

Procalcitonin

Once - Routine - Lab First occurrence Today at 0921 Do you want to change the specimen collection from what it shows in the banner bar? No

Culture, Blood - 1st of 2 Perpheral Draw STAT - Lab, 1st of 2 Peripheral Draw. Phleb to determine site

Culture, Blood - 2nd of 2 Peripheral Draw STAT - Lab, 2nd of 2 Peripheral Draw. Phleb to determine site

Aerobic Respiratory Culture with Gram Stain Once - Routine - Lab, Sputum, Sputum Induced *Daily ASP Prospective Audit and Feedback







Education

- Presentation to stakeholders and house staff
- Partnered with hospitalists
- Education through prospective audit and feedback

Community Acquired Pneumonia (CAP) Care Pathway



Orders are being started in the ED

- Labs: Strep Ag, legionella Ag, procalcitonin (+/- flu testing)
 - Consider sputum cultures in patients being started on broad-spectrum antibiotics (e.g. high DRIP scores) to help with deescalation
- Antibiotics for most floor patients with CAP
 - o Ceftriaxone 2 grams IV x 1 dose
 - o Azithromycin 500 mg IV x 1 dose
 - Then, Cefuroxime 500 mg PO BID x4 days (to start 24 hours after initial antibiotics)
- Most patients <u>DO NOT NEED</u> ongoing IV antibiotics or additional azithromycin
 - No need to continue IV antibiotics because of ongoing fever, leukocytosis or tachycardia. Switch to oral therapy as long as tolerating oral diet and PO meds.

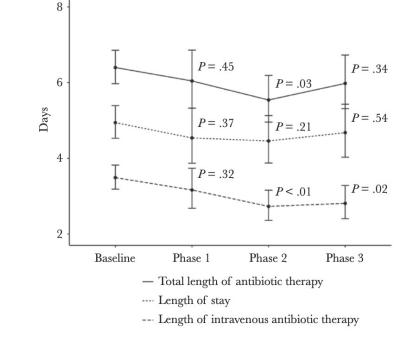
PLEASE DO NOT DISCONTINUE ED ORDERS

- Improves antibiotic stewardship
- Reduces LOS

A Pathway for Community-Acquired Pneumonia With Rapid Conversion to Oral Therapy Improves Health Care Value 3

- Median cost per case decreased by 20%
- Total length of antibiotic duration decreased by 1 day
- IV duration of antibiotics decreased by 22%
- No change in readmission rate





Ciarkowski CE, et al. Open Forum Infect Dis. 2020 Oct 19;7(11):ofaa497.

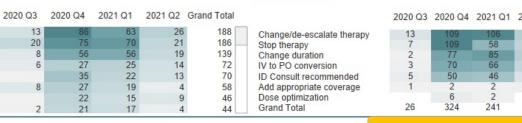
ANTIMICROBIAL STEWARDSHIP | EXECUTIVE DASHBOARD

Lâ HEALTH

Click to open MGA Knowledge Management page

9/1/2020 12:00:00 AM to 9/3.. FLOWSHEET DAT., Date Filter Sep 2020 to Jul 2021 Updated: 7/26/2021 9:36:50 AM

Date Filter



Sep 2020 to Jul 2021

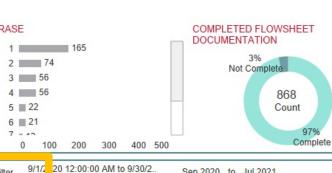
INTERVENTION



COMMUNITY-AQUIRED PNEUMONIA (CAP) Date Filter

6 21

7 - 42





MEDICATIONS REVIEWED

Piperacillin + Tazobactam

Ceftriaxone

Vancomycin

Metronidazole

Cefuroxime

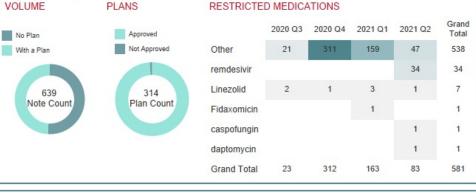
Cefepime

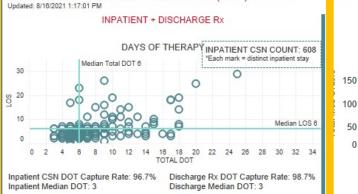
Cefazolin

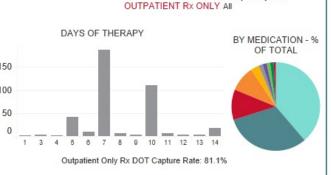
Fluconazole

RESTRICTED MEDICATIONS

9/1/2020 12:00:00 AM to 9/30/20.

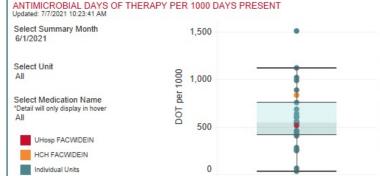






Sep 2020 to Jul 2021

NHSN DATA



STANDARDIZED ANTIMICROBIAL ADMINISTRATION RATIO (SAAR): ALL UNITS Updated: 5/26/2021 12:02:04 PM

2017 Antimicrobial Agent Category Filter Adult All Antibacterial



(MGA

OP Specialty Fitler

IDSA/ATS 2019 CAP Guidelines

- We recommend obtaining pretreatment Gram stain and culture of respiratory secretions in adults with CAP managed in the hospital setting who:
 - Have severe CAP
 - Are empirically being treated for MRSA or pseudomonas
- We recommend not routinely obtaining blood cultures in adults with CAP managed in the hospital setting
 - Exceptions for severe CAP and those empirically treated for MRSA or *Pseudomonas*
- We suggest not routinely testing urine for Legionella antigen in adults with CAP except
 - Epidemiological factors (outbreak, recent travel)
 - Severe CAP
- We suggest not routinely testing urine for pneumococcal antigen in adults with CAP except
 - Severe CAP

CAP Data Review (8/1/2019-1/31/2020)





Microbiology		89 Patients total	UNIVERSITY O
Respiratory culture			
Not performed		74 (83%)	
No growth	Only 1/89 patients had a respiratory	3 (3%)	
Normal resp	culture that would influence	10 (11%)	
Strep		1 (1%)	
MSSA		1 (1%)	
MRSA		0	
Pseudomonas		1 (1%)	
Other		5 (6%)	
Blood culture	<i>None</i> of the blood cultures would		
Not perform	influence management	31 (35%)	
Negative	innuchee management	56 (63%)	
True Positive		1 (1%)- strep pneum.	
Contaminant		1 (1%)	

Slide courtesy of Valerie Vaughn and Guinn Dunn

CAP Data Review (8/1/2019-1/31/2020)



Microbiology	UNIVERSIT 89 Patients total
Legionella urine antigen	
Not performed	9 (10%)
Negative	80 (90%)
Strep pneumo urine antigen	
Not performed	9 (10%)
Negative	69 (78%)
Positive	11 (12.4%)

None would influence management

Slide courtesy of Valerie Vaughn and Guinn Dunn



CAP 2.0: Recent changes to pathway for floor patients

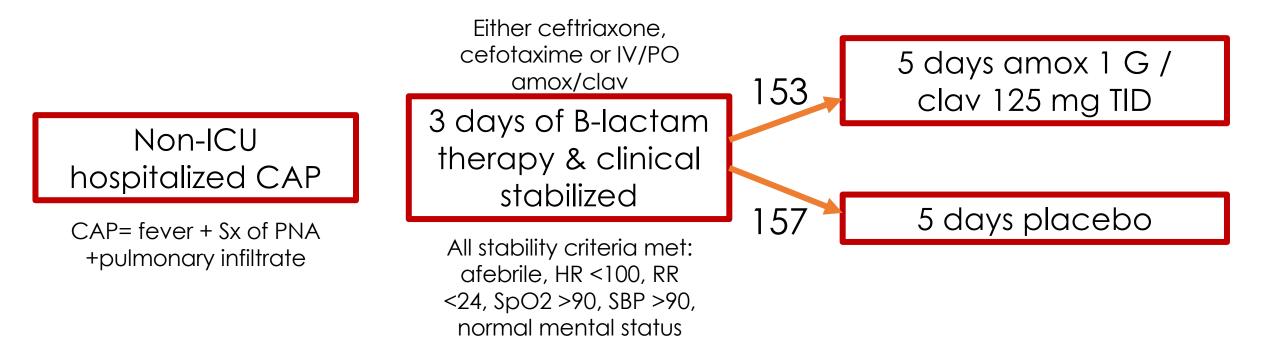
- No blood cultures
- No Streptococcus pneumoniae urine antigen
- Removed the DRIP score
- Dropped empiric azithromycin
- Default antibiotic duration of 3 days
- Amoxicillin step down oral drug

Validation of a Community-Acquired Pneumonia Score To Improve Empiric Antibiotic Selection at an Academic Medical Center



- IDSA/ATS 2019 CAP Guidelines recommend utilizing locally validated risk factors to identify patients at risk for drug-resistant pathogens (DRP)
- Validation at the SLC VA showed broad-spectrum antimicrobial use would increase by 9%
- Retrospective case-control validation study to assess predictive performance of DRIP and identify local risk factors for Drug-Resistant Pathogens
- DRIP at the U: Sensitivity 67% and specificity 73%. AUROC curve was 0.76(95% confidence interval [CI], 0.69 to 0.82)
 - Decreased performance from original published validation
- DRP within the last year predictive of current DRP

Babbel D, Sutton J, Rose R, Yarbrough P, Spivak ES. Antimicrob Agents Chemother. 2018 Feb 23;62(3):e02277-17. Oliver MB, Fong K, Certain L, Spivak ES, Timbrook TT. Antimicrob Agents Chemother. 2021 Jan 20;65(2):e01482-20. Discontinuing β-lactam treatment after 3 days for patients with community-acquired pneumonia in non-critical care wards (PTC): a double-blind, randomised, placebo-controlled, non-inferiority trial



Lancet 3 vs 8 Days for CAP

No difference in any outcomes

- Primary outcome: cure at 15 days
 - Cure was defined as afebrile, resolution of signs/symptoms, no additional antibiotics
 - 77% (3 day) vs. 68% (8 days); difference of 9·44% [95% CI −0·15 to 20·34] → non-inferior
- Adverse events (14% vs. 19%)
- Mortality (2% vs. 1%)

CAP: Antibiotic duration

- RCT of patients with mild to moderate/severe CAP
- All pts. received IV amoxicillin for 3 days
- At 3 days pts. were randomized into two groups if they had improvement, become afebrile, and were able to take oral therapy:
 - Amoxicillin 750 mg PO TID x 5 days
 - Placebo TID x 5 days

Research

BMJ

Effectiveness of discontinuing antibiotic treatment after three days versus eight days in mild to moderate-severe community acquired pneumonia: randomised, double blind study

Rachida el Moussaoui, Corianne A J M de Borgie, Peterhans van den Broek, Willem N Hustinx, Paul Bresser, Guido E L van den Berk, Jan-Werner Poley, Bob van den Berg, Frans H Krouwels, Marc J M Bonten, Carla Weenink, Patrick M M Bossuyt, Peter Speelman, Brent C Opmeer, Jan M Prins

	3 Days	8 Days
Clinical cure at 10 days	93%	93%
Clinical cure at 28 days	90%	88%
Adverse events	11%	21%

Moussaoui, BMJ 2006

New Order Set (Rolled out 11/11/21)

★ Orders

Pneumonia Orderset for Inpatient Care 🛛 🕿

Antibiotics for CAP

CAP Treatment Options

Floor (NO History of antibiotic-resistant infection in the past year)

Antibiotics

Preferred Antibiotics

cefTRIAXone (ROCEPHIN) 2 gram in sodium chloride 0.9% Mini-Bag 2 gram, Intravenous, Administer over 30 Minutes, at 200 mL/hr, Once, today at 1300, For 1 dose

Antimicrobial Use: Empiric Antimicrobial Indication: Respiratory, Pneumonia

Followed By

amoxicillin (AMOXIL) capsule 1,000 mg 1,000 mg, Oral, 3 times daily, First dose tomorrow at 1300, For 2 days

 Antimicrobial Use: Empiric Antimicrobial Indication: Respiratory, Pneumonia Routine

O Severe Beta Lactam Allergy

Oseltamivir Standard CrCl Adjustment Panel

🗸 Labs

Streptococcus Pneumoniae Antigen, Urine Once - Routine - Lab, Urine, Urine-General

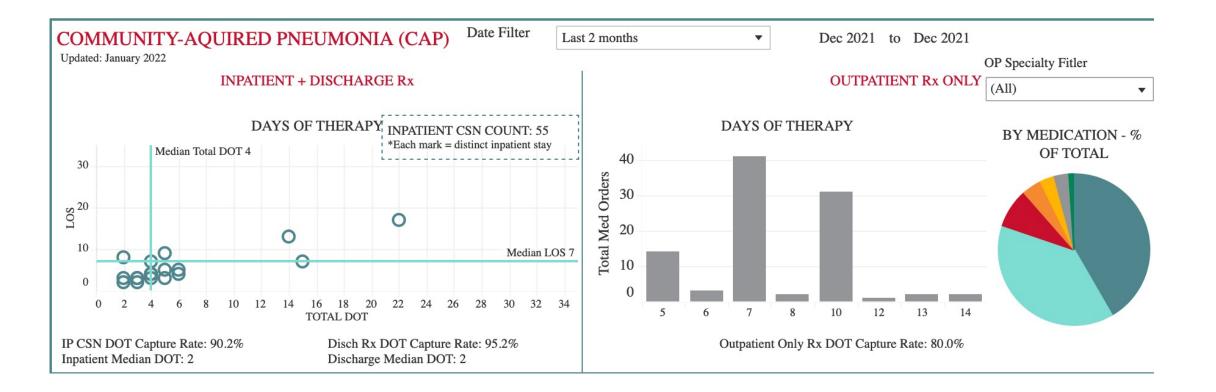
Legionella pneumophila Antigen, Urine
Once - Routine - Lab, today at 1259, For 1 occurrence
Urine, Urine-General

🗸 Procalcitonin

Aerobic Respiratory Culture with Gram Stain

Once - Routine - Lab, Sputum, Sputum Induced

Tracking Adherence



Conclusions

- Syndrome-specific stewardship interventions very impactful given evidence-based best practices X frequent diagnosis
- Standardizing practice can have significant impact
 - Embed into workflow
 - Requires ongoing audit and feedback (decreasing over time)
- Updating local guidance over time important as data evolves