

Is it Truly Negative? The Effect of Definitions and Cancer Prevalence on Diagnostic Yield Estimates of Bronchoscopy

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### The clinical question

How does the estimated diagnostic yield of bronchoscopy change when different definitions for a diagnostic biopsy are used?

### Take home message

Diagnostic yield is highly dependent on cancer prevalence in the study population and the specific definition of diagnostic yield that is used.

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### Background

The diagnostic yield for bronchoscopy is influenced by lesion characteristics and procedural tools. However, cancer prevalence and the definition used to define a diagnostic procedure can also impact the estimated yield of bronchoscopy.

Recently, a meta-analysis of navigational bronchoscopy described that diagnostic yield estimates ranged broadly from 40% to 90%, which could be from a lack of standardization of what is considered a "true negative" in nonmalignant biopsy results.

## **Study Design**

- Type of trial: simulation-based cohort study
- Randomization, blinding, controls: N/A
- N: 1000
- Study groups: 4 groups with different definitions of diagnostic yield
- Settings: simulated patient environment
- Enrollment: simulation framework based in literature
- Treatment period: N/A
- Follow up: assumed 10% lost to follow up for calculations
- **Primary outcome:** Diagnostic yield using each definition. A change in yield >10% was considered clinically meaningful

# Population

- Inclusion criteria: hypothetical cohort of patients who underwent diagnostic bronchoscopy
- Exclusion criteria: cases who were randomly determined as lost to follow up (LTFU) in follow-up phase
- **Baseline Characteristics:** derived averages from literature: sensitivity for malignancy at index bronchoscopy of 80%, total cancer prevalence of 60%, distribution of specific benign (SPB), nonspecific benign (NSB), and nondiagnostic (ND) of 10%, 35%, and 55% respectively, and LTFU rate of 10%

### Outcomes

#### **Primary outcomes:**

- Diagnostic yield differed >10% amongst methods in 76.7% of cases.
- With base assumptions for 1,000 simulated patients undergoing bronchoscopy in a population with 60% cancer prevalence, diagnostic yield estimates were 53.2% for method 1, 71.4% for method 2, 66.8% for method 3, and 89.2% for method 4

#### Secondary outcomes:

- Variation in cancer prevalence had largest effect on DY. Changing cancer prevalence from 40% to 80% had following impact
- Method 1 38.8% to 67.6%
- Method 2 62.6% to 80.2%
- Method 3 58.5% to 74.8%
- Method 4 92.4% to 87.2%. (Inverse relation)
- Categorization of nonmalignant findings had the second largest effect on DY

#### Adverse events: N/A

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### Commentary

### Strengths

- Shows how cancer prevalence and definitions of diagnostic bronchoscopy can dramatically change estimates on utility of bronchoscopy
- Shows how biased definitions may alter expected yield of procedure
- Propose standardized definitions for future studies

### Study Limitations and Potentials for Bias:

- Many of the standardized estimates used in the calculations were based on expert opinion which may not be representative of the entire population
- Considered 4 different definitions and 3 classifications of benign diagnoses
- Simulation model may not capture all variables and relationship between them

# Funding

Johnson & Johnson Lung Cancer Initiative

# Suggested Reading

(References in Vancouver style)

1. Nadig TR, Thomas N, Nietert PJ, Lozier J, Tanner NT, Wang Memoli JS, et al. Guided bronchoscopy for the evaluation of pulmonary lesions: an updated meta-analysis. CHEST 2023;163:1589–1598.



2. Ost DE, Ernst A, Lei X, Kovitz KL, Benzaquen S, Diaz-Mendoza J, et al.; AQuIRE Bronchoscopy Registry. Diagnostic yield and complications of bronchoscopy for peripheral lung lesions. Results of the AQuIRE Registry. Am J Respir Crit Care Med 2016;193:68–77.

3. Kops S, Heus P, Korevaar D, Damen J, Idema D, Verhoeven R, et al. Diagnostic yield and safety of navigation bronchoscopy: A systematic review and metaanalysis. Lung Cancer 2023:180:107196

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# **Article citation**

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