



# Diagnostic accuracy of D-dimer for acute aortic syndromes: systematic review and meta-analysis

Kaelan Gobeil Odai<sup>1</sup> · Sacha Weill<sup>1</sup> · Robert Goulden<sup>1,2</sup>

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- Annals of Emergency Medicine (annemergmed.com)

## Article type

Systematic Review and Meta Analysis

## Rankings

Methods: 4/5

Usefulness 3/5

## Introduction

### Background

D-dimer has been proposed to be a sensitive marker to help “rule out” Acute Aortic Syndromes (AAS), but early studies were methodologically limited and led to an overestimation of sensitivity.

### Objectives

The author’s aim to update the current literature surrounding the use of D-dimer without the inclusion of case–control studies to better estimate the diagnostic accuracy of D-dimer in AAS.

## Methods

**Design:** Systematic review and meta-analysis.

**Eligibility Criteria:** All cohort studies comparing non-traumatic D-dimer to a reference imaging modality (CTA, MRI, echocardiography). Case control studies were excluded.

**Setting:** Emergency Department.

**Subjects:** Anyone presenting to emergency department (ED) with symptoms suggestive of AAS. All ages included. Studies including patients with incidental findings of AAS were excluded.

**Intervention:** D-dimer.

**Outcome:** To assess the diagnostic accuracy of D-dimer for AAS.

## Results

Twenty five studies were included for a total of 9228 patients who were assessed with a D-dimer and a reference imaging test in the setting of symptoms suggestive of AAS. Summary test characteristics were calculated for the 500 ng/mL threshold, showing the following: sensitivity of 95.7% (95% CI: 93.2%–97.5%) and specificity of 57.5% (95% CI: 50.1%–64.6%). The pooled likelihood ratio for a positive test was 2.25 (95% CI: 1.93–2.68), and the pooled likelihood ratio for a negative result was 0.08 (95% CI: 0.04–0.11). The 95% PrI of sensitivity of 86.1%–99.3%, with a 95% PrI of specificity of 25.3%–83.1%. Using QUADAS-2, most studies had a low or unclear risk of bias and high applicability.

## Appraisal

### Strengths

- clear objectives and reason to repeat a systematic review and meta-analysis;

✉ Kaelan Gobeil Odai  
dr.k.gobeilodai@outlook.com

<sup>1</sup> McGill University Health Centre, Montréal, QC, Canada

<sup>2</sup> Department of Emergency Medicine, McGill University, Montréal, QC, Canada

- methodologically sound, appraising publication bias using appropriate statistical models;
- study registered in a database;
- followed PRISMA guidelines;
- used a validated study assessment tool (QUADAS-2);
- avoids overestimation of effect by intentionally omitting case control studies;
- demonstrates encouraging results.

### Limitations

- unclear patient population in which to use D-dimer (low vs moderate risk);
- studies included have varying degrees of pre-test probability for AAS ranging from 1–60%;
- difficult to implement as there is no externally validated clinical decision rule that provides pre-test probability to guide management;
- no age adjusted dimer measurement for older individuals;
- studies were compared to any reference imaging, not gold standard ECG-Gated CTA;
- most studies had poor, moderate or unclear risk of bias;
- most studies were poor in methodological rigor;
- Significant heterogeneity between study protocols included in the study.

### Context

This is the most recent systematic review and meta-analysis meant to assess the use of D-dimer alone for ruling out AAS. The Canadian Practice Guidelines (CPG) by Ohle et al. [2] highlight the importance of patient selection. *“In those with a moderate risk for AAS, a normal result of D-dimer testing is reasonable to reduce probability of AAS; in patients with a low or high probability of the condition, the use of D-dimer is not recommended”*. The recent PROFUNDUS study [3] demonstrated that when used together, POCUS, D-dimer, and pre-test probability (PTP), are highly sensitive tools for selected PTP patients.

### Bottom line

Through this meta-analysis and others, there is robust data to support that D-dimers are highly sensitive for AAS. However, without clear guidelines on pretest probability, its isolated use is questionable especially given its specificity remains poor. Therefore, D-dimer alone, cannot be used to diagnose or rule out AAS. Clinicians should familiarize themselves with the recent CPGs to adequately assess pre-test probability, stratify the risk of AAS, determine the pertinence of D-dimers, and the necessity for further advanced imaging techniques.

### Declarations

**Conflict of interest** I, Kaelan Gobeil Odai declare that I have no conflicts of interest to the aforementioned manuscript I, Sacha Weill declare that I have no conflicts of interest to the aforementioned manuscript I, Robert Goulden declare that I have no conflicts of interest to the aforementioned manuscript.

### References

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