

## Myeloperoxidase Deficiency: A potential cause of neutrophil anergy in chronic recurrent vulvovaginal candidiasis

Kathryn Cataldo, BS<sup>1</sup>, Christine Conageski, MD<sup>2</sup>, Kristen Desanto, MS<sup>3</sup>

<sup>1</sup>University of Colorado School of Medicine, MD Class of 2022

<sup>2</sup>Department of Obstetrics & Gynecology, Anschutz Medical Center, University of Colorado

<sup>3</sup>Strauss Health Sciences Library, Anschutz Medical Center, University of Colorado

Address for correspondence: Christine Conageski, MD, Division of Obstetrics & Gynecology, Anschutz Medical Center, University of Colorado, 12631 East 17<sup>th</sup> Ave, Aurora, CO 80045 (email: Christine.conageski@cuanschutz.edu)

## Abstract

**Background** Recurrent vulvovaginal candidiasis (RVVC) is an exceedingly common infection of the female reproductive tract that effects 75% of immunocompetent women during their childbearing years. A hallmark of RVVC is robust recruitment of neutrophils to the vaginal lumen that are unable to effectively kill the pathogen during infection, termed neutrophil anergy. Recently, significant research has been aimed at identifying the underlying pathophysiology of neutrophil anergy, of which one key factor appears to be a disabled respiratory burst. The purpose of this minireview is to highlight the significant advancements in the understanding of the pathogenesis of RVVC and neutrophil anergy, as well as to provide an overview of the current understanding of myeloperoxidase (MPO) deficiency, its role in the respiratory burst, and how it may relate to RVVC.

Methods To identify potentially relevant studies, the following electronic databases were searched on MEDLINE, Embase, Web of Science Core Collection, and Cochrane Library. Database searches were not limited by species, age group of subjects, language, or publication date. All database search results were exported to and deduplicated in EndNote<sup>™</sup> 20. The Yale MeSH Analysis Grid Generator was used to identify keywords, synonyms, and subject heading terms to develop the search strategies, based on five articles initially identified by the authors as being relevant to the topic. The relevant studies were then reviewed in detail for content related to the study question.

**Results** 719 articles were retrieved in the initial literature review. Deletion of duplicate articles yielded 397 unique articles, all of which were then reviewed for content. 137 articles remained after excluding articles based on a series of criteria (language barrier, relevance to mucocutaneous candidiasis and vulvovaginal candidiasis, relevance to OB/GYN care, as well as foundational concepts related to innate and adaptive immunity in the context of candida infection). Of all 137 articles, only two discussed MPO deficiency as a potential cause of neutrophil anergy in the context of chronic recurrent vulvovaginal candidiasis.

**Discussion** MPO deficiency is exceedingly common, heterozygosity estimated at more than 1 out of 50. RVVC is also exceedingly common, effecting 1 out of every 10 women during their reproductive years. The question remains whether the unique environment of the female reproductive tract unmasks the otherwise asymptomatic phenotype of MPO deficiency. Obstetric and Gynecological clinicians encountering patients with recurrent vulvovaginal candida infections not adequately explained by other mechanisms should consider a multidisciplinary approach to include Immunology for evaluation of MPO deficiency.

**Keywords** Myeloperoxidase, neutrophil, PMN, candida, anergy, immunology, vulvovaginal candidiasis, respiratory burst, cell immunity, chronic infection