

# Long-Term Risk of Colorectal Cancer in Patients With Prediabetes: A Systematic Review and Meta-Analysis

Praneeth Reddy Keesari, MBBS<sup>1</sup>; Yashwitha Sai Pulakurthi, MBBS<sup>2</sup>; Sreeram Pannala, MBBS<sup>3</sup>; Jnana Pramod Valluri, MBBS<sup>4</sup>; Nikhilendhar Nag Mopuru, MBBS<sup>5</sup>; Adhvithi Pingili, MBBS<sup>8</sup>; MBBS<sup>8</sup>; Manasa Ginjupalli, MBBS<sup>7</sup>; Sahas Reddy Jitta, MBBS<sup>8</sup>; Rishi Devaraja Vattikuti, MBBS<sup>9</sup>; Ummul Asfeen, DO<sup>10</sup>; Rupak Desai, MBBS<sup>11</sup>

Staten Island University Hospital, <sup>2,10</sup>New York Medical College – Saint Michael's medical Center, <sup>3,4,6,9</sup>Kamineni Academy of Medical Sciences and Research Centre, <sup>6</sup>Medstar health, <sup>7,8</sup>Osmania Medical College, <sup>11</sup>Independent researcher

Odds Ratio (log scale

## Introduction

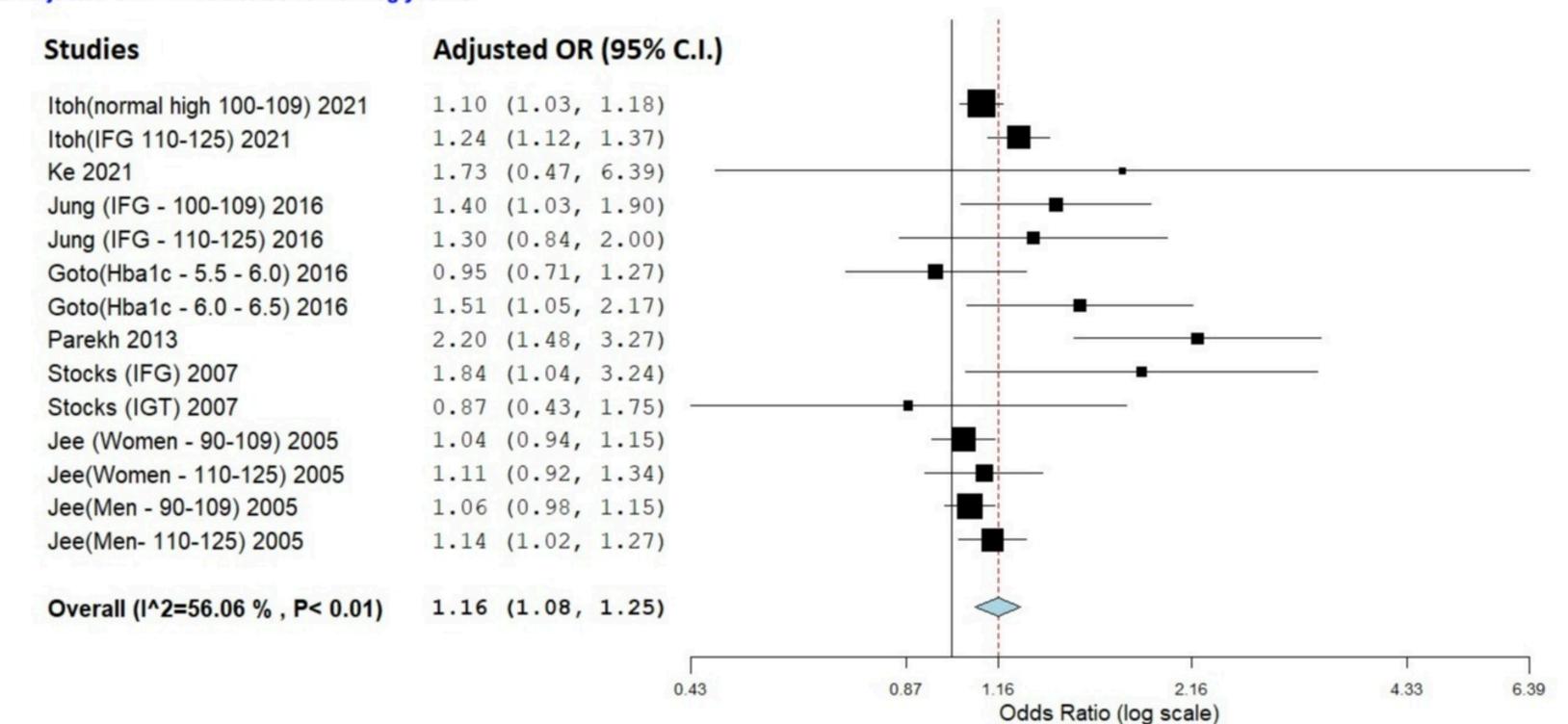
Prediabetes is often underdiagnosed and underreported due to its asymptomatic state in over 80% of individuals. Considering its role in promoting cancer incidence and limited evidence linking prediabetes and colorectal cancer (CRC), we conducted a meta-analysis to evaluate the incidence of colorectal cancer in people with prediabetes.

# Methods and Materials

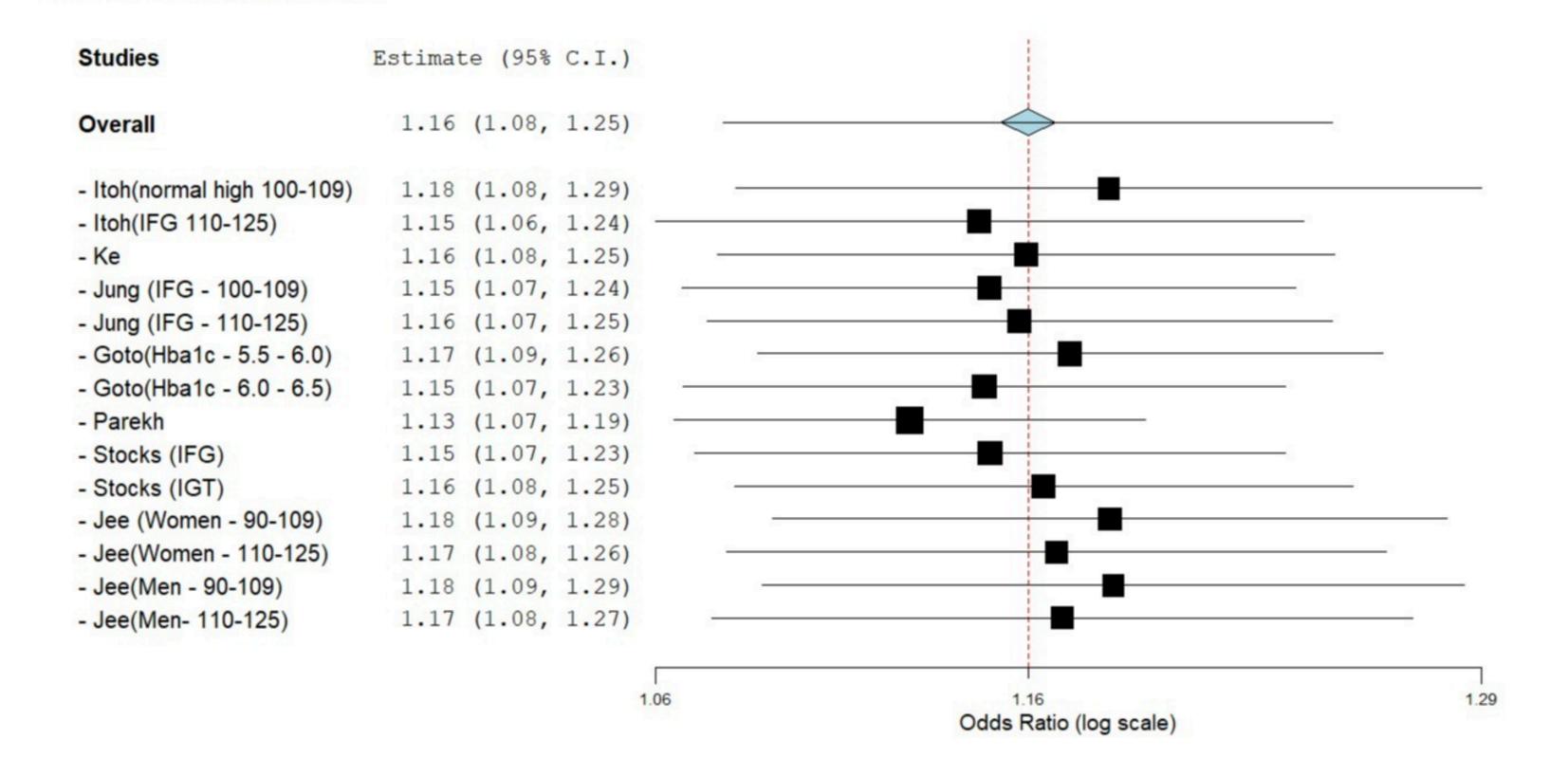
through comprehensive search PubMed/Medline, Embase, Scopus, Google Scholar was performed until June 1, screen for studies reporting 2022, to incidence/risk colorectal cancer prediabetics. Binary random-effects models were used to perform meta-analysis and subgroup analyses. Sensitivity analysis was done using leave-one-out method.

# Figure 1 a. Unadjusted OR -- Prediabetes vs Normoglycemia Studies Odds Ratio (95% CI) Itoh(normal high 100-109) 2021 1. 65 (1. 55, 1. 76) Itoh(IFG 110-125) 2021 2. 26 (2. 06, 2. 48) Rentsch 2020 1. 53 (1. 29, 1. 81) Jung (IFG - 100-109) 2016 1. 40 (1. 03, 1. 90) Jung (IFG - 110-125) 2016 1. 30 (0. 84, 2. 00) Stocks (IFG) 2007 1. 87 (1. 08, 3. 25) Stocks (IGT) 2007 0. 90 (0. 45, 1. 79) Overall (I^2=85.72 %, P< 0.01) 1. 62 (1. 35, 1. 95)

### b. Adjusted OR - - Prediabetes vs Normoglycemia



### c. Leave one out Sensitivity Analysis



# Results

Seven prospective and one retrospective study comprising 15 cohorts and a pooled number of 854876 cases and 2190511 controls were included in the analysis (2 Japan, 2 Korea, 1 Sweden, 1 UK, 1 China, and 1 USA). After combining all the studies the forest plots for adjusted analysis shows a significant increase in odds of having CRC with prediabetes - (OR 1.16; 1.08-1.25, p< 0.01) (fig 1b) and unadjusted analysis also shows a significant increase in odds of having CRC with prediabetes (OR 1.62; 1.35–1.95, p< 0.01) (fig 1a). Sensitivity analysis using the Leave-one-out method did confirm equivalent results (fig 1c). Heterogeneity analysis for adjusted OR had moderate heterogeneity with an overall I^2 of 56.06% with a p value < 0.01 and for unadjusted OR had considerable heterogeneity with an overall I^2 of 85.72% with a p value less than 0.01. Subgroup analysis based on type of study, the odds of developing colorectal cancer was higher in prospective studies (OR 1.175; 1.065-1.298)(p 0.001) than retrospective studies(OR 1.162; 1.033- 1.306)(p 0.012) The odds of developing cancer was not significantly higher in ages >60(OR 1.446; 0.887-2.356)(p 0.139) compared to less than 60 years. The strongest association b/w Prediabetes and CRC was found on a median 5-10 years(aOR 1.257; 1.029-1.534)(p 0.025) follow-up compared to < 5 years and 10 years and

# Discussion

This study showed nearly 16% higher long-term risk of colorectal cancer in patients with prediabetes. Lifestyle modifications like weight loss, proper diet, and exercise are essential to control prediabetes. This study further warrants a specific prediabetes screening for patients already at high risk of colorectal cancer with other risk factors. These strides would help subsequently lower the disease burden, and associated morbidity/mortality.