# Effects of bariatric surgery on rapid remission of NAFLD: an exploratory metabolomics and validation study: a prospective cohort study

Mengyi Li, MD, PhD<sup>1\*</sup>, Xinyu Wang, BM<sup>2\*</sup>, Na Zeng, MSc<sup>2\*</sup>, Zhiyu Wu, MSc<sup>2</sup>, Canqing Yu. PhD<sup>2,3,4</sup>, Dianjianyi Sun, PhD<sup>2,3,4</sup>, Yang Liu, MD, PhD<sup>1</sup>, Di Cao, MD, PhD<sup>5</sup>, Peng Zhang, MD, PhD<sup>1</sup>, Ling Yang, PhD<sup>6,7</sup>, Yiping Chen, DPhil<sup>6,7</sup>, Zhengming Chen, DPhil<sup>6,7</sup>, Liming Li, MD<sup>2,3,4</sup>, Zhongtao Zhang, MD, PhD<sup>1</sup>, Jun Lv, PhD<sup>2,3,4</sup>, Yuanjie Pang, DPhil<sup>2,3</sup>

- 1. Department of General Surgery, Beijing Friendship Hospital, Capital Medical University; State Key Lab of Digestive Health; National Clinical Research Center for Digestive Diseases, Beijing 100050, China
- 2. Department of Epidemiology & Biostatistics, School of Public Health, Peking University, Beijing 100191, China
- 3. Key Laboratory of Epidemiology of Major Diseases (Peking University), Ministry of Education
- 4. Peking University Center for Public Health and Epidemic Preparedness & Response, Beijing 100191, China
- 5. Department of Radiology, Beijing Friendship Hospital, Capital Medical University, Beijing 100050, China
- 6. Medical Research Council Population Health Research Unit at the University of Oxford, Oxford, United Kingdom
  Clinical Trial Service Unit & Epidemiological Studies Unit (CTSU), Nuffield Department of Population Health, University of Oxford, United Kingdom



[ $\sqrt{\ }$ ] I have no potential conflict of interest to report.



**Objective:** To examine the associations between alteration of plasma metabolites and the effects of bariatric surgery on rapid remission of non-alcoholic fatty liver disease (NAFLD), and to further validate the results in a general population-based cohort.

**Summary of Background Data:** Bariatric surgery is a promising procedure to induce substantial weight loss and to alleviate NAFLD in short post-surgical period, but the underlying causal associations related to metabolomics are unclear.



# **Methods:**

#### Base-NAFLD cohort

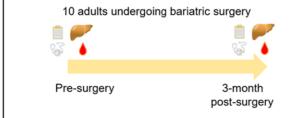
- 138 patients with NAFLD undergoing bariatric surgery between 2020-2023
- Demographics: age 17-61, female 74.6%
- BMI: 25.0-57.4 (mean 38.5 kg/m²)
- NAFLD: 100%, PDFF 5.2-54.0 (mean 19.0%)



#### Inclusion criteria

- Diagnosed with obesity according to the WHO criteria for obesity in Asian populations (BMI ≥27.5 kg/m²) and scheduled for a primary bariatric surgery
- Diagnosed with hepatic steatosis preoperatively by MRI-PDFF or intraoperative hepatic pathology
- Selected for the metabolomics sub-study





## China Kadoorie Biobank (CKB)

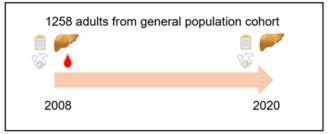
- 517 thousands participants from 10 regions in China, baseline survey 2004-2008
- Demographics: age 30-79, female 59.0%
- BMI: 17.0-32.7 (mean 23.7 kg/m<sup>2</sup>)
- NAFLD\*: 5.1%



### Inclusion criteria

- Attended the first (2008-2010) and the third resurvey (2020-2021)
- Selected for the metabolomics sub-study
- No NAFLD at the first resurvey

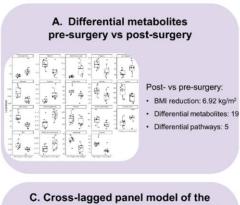


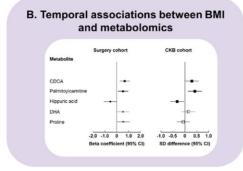


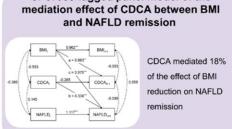


# **Results:**

Figure 2. Conceptual framework and key results of the current study







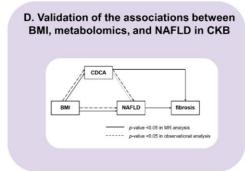
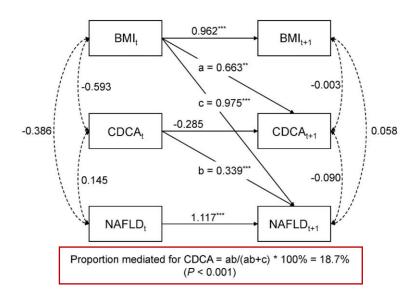


Figure 3. Cross-lagged panel model of the mediation effect of CDCA between BMI and NAFLD remission



**Conclusions:** CDCA, a bile acid metabolite, mediated causal effects of bariatric surgery on the rapid remission of NAFLD. It may also predict liver fibrosis improvement following bariatric surgery.

