Filed Pursuant to Rule 424(b)(4) Registration No. 333-221010 Registration No. 333-221716

PROSPECTUS SUPPLEMENT Dated September 20, 2021 (To Prospectus Dated November 22, 2017)



Nuwellis, Inc.

127 Warrants to Purchase Common Stock (and 50,800 Shares of Common Stock Underlying the Warrants to Purchase Common Stock)

This prospectus supplement (this "Supplement") modifies, supersedes and supplements certain information contained in, and should be read in conjunction with, that certain prospectus (including the information incorporated by reference therein, the "Prospectus") filed by Nuwellis, Inc. (f/k/a CHF Solutions, Inc.) (the "Company"), dated November 22, 2017, related to the underwritten public offering (the "November 2017 Offering") of the Company's Series F convertible preferred stock, Series 1 warrants and Series 2 warrants (each as defined in the Prospectus) and later supplemented by the prospectus supplements filed by the Company on July 3, 2018, March 13, 2019, October 25, 2019, November 7, 2019, January 29, 2020, March 24, 2020, and September 3, 2021. This Supplement is not complete without, and may not be delivered or used except in connection with, the Prospectus, as previously supplemented.

The Company's common stock is listed on the Nasdaq Capital Market under the symbol "NUWE." On September 17, 2021, the last reported sale price of the Company's common stock on the Nasdaq Capital Market was \$2.60 per share.

The information contained in this Supplement modifies and supersedes, in part, the information in the Prospectus. Any information that is modified or superseded in the Prospectus shall not be deemed to constitute a part of the Prospectus, except as modified or superseded by this Supplement.

We may amend or supplement the Prospectus from time to time by filing amendments or supplements as required. You should read the entire Prospectus and any amendments or supplements carefully before you make an investment decision.

Investing in our securities involves risks. See "Risk Factors" beginning on page 7 of the Prospectus and in the documents incorporated by reference into the Prospectus.

FORWARD-LOOKING STATEMENTS

You should carefully consider the risk factors set forth in the Prospectus, as well as the other information contained, or incorporated by reference, in this Supplement or the Prospectus. The Prospectus contains forward-looking statements about our expectations, beliefs, plans, objectives, assumptions or future events or performance that are not historical facts. You are cautioned that any forward-looking statements are not guarantees of future performance and are subject to risks and uncertainties. Actual results may differ materially from those included within the forward-looking statements as a result of various factors. Cautionary statements in the "Risk Factors" section of the Prospectus and in the documents incorporated by reference therein identify important risks and uncertainties affecting our future, which could cause actual results to differ materially from the forward-looking statements made or included in this Supplement and the Prospectus. Further, any forward-looking statement speaks only as of the date on which it is made, and we undertake no obligation to update any forward-looking statement or statements to reflect events or circumstances after the date on which such statement is made or to reflect the occurrence of unanticipated events.

REDUCTION OF CONVERSION PRICE OF SERIES F CONVERTIBLE PREFERRED STOCK

Pursuant to the terms set forth in the Certificate of Designations of Preferences, Rights and Limitations of the Company's Series F Convertible Preferred Stock (the "Series F Preferred Stock"), such Series F Preferred Stock is subject to adjustment in certain circumstances, including in connection with stock splits or if the Company sells common stock at a price lower than the then-conversion price of the Series F Preferred Stock. As a result of an underwritten public offering that closed on July 3, 2018 (the "July 2018 Offering"), the conversion price of the Series F Preferred Stock was reduced from \$4.50 to \$2.12, the per share price to the public in the July 2018 Offering, upon the closing of the July 2018 Offering on July 3, 2018. On January 2, 2019, the Company effected a 1-for-14 reverse stock split, which further adjusted the conversion price of the Series F Preferred Stock to \$29.68. As a result of an underwritten public offering that closed on March 12, 2019 (the "March 2019 Offering"), the conversion price of the Series F Preferred Stock was further adjusted to \$5.25, the per share price to the public in the March 2019 Offering, upon the closing of the March 2019 Offering on March 12, 2019. As a result of a registered direct offering that closed on October 25, 2019 (the "October 2019 Offering"), the conversion price of the Series F Preferred Stock was further adjusted to \$1.15, the per share price to the public in the October 2019 Offering, upon the closing of the October 2019 Offering on October 25, 2019. As a result of a registered direct offering with a concurrent private placement that closed on November 6, 2019 (the "November 2019 Offering"), the conversion price of the Series F Preferred Stock was further adjusted to \$0.9942, the per share price to the private investors in the November 2019 Offering upon the exercise of warrants, upon the closing of the November 2019 Offering on November 6, 2019. As a result of an underwritten public offering that closed on January 28, 2020 (the "January 2020 Offering"), the conversion price of the Series F Preferred Stock was further adjusted to \$0.55, the per share price to the public in the January 2020 Offering, upon the closing of the January 2020 Offering on January 28, 2020. As a result of a registered direct offering that closed on March 23, 2020 (the "March 2020 Offering"), the conversion price of the Series F Preferred Stock was further adjusted to \$0.30, the per share price to the public in the March 2020 Offering, upon the closing of the March 2020 Offering on March 23, 2020. On October 16, 2020, the Company effected a 1-for-30 reverse stock split, which further adjusted the conversion price of the Series F Preferred Stock to \$9.00. As a result of an underwritten public offering that closed on March 19, 2021 (the "March 2021 Offering"), the conversion price of the Series F Preferred Stock was further adjusted to \$5.50, the per share price to the public in the March 2021 Offering, upon the closing of the March 2021 Offering on March 19, 2021. As a result of an underwritten public offering that closed on September 17, 2021 (the "September 2021 Offering"), the conversion price of the Series F Preferred Stock was further adjusted to \$2.50, the per share price to the public in the September 2021 Offering, upon the closing of the September 2021 Offering on September 17, 2021. As of September 17, 2021, 127 shares of Series F Preferred Stock were outstanding and convertible into 50,800 shares of common stock, following such conversion price adjustment.

This Supplement is being filed to reflect the foregoing adjustments in the conversion price of the Series F Preferred Stock, as described avobe.

The date of this prospectus supplement is September 20, 2021.