



HAL
open science

The Association of Weight Loss, Weight Status, and Abdominal Obesity with All-Cause Mortality in Older Adults

Tagrid Abdullah Alharbi, Joanne Ryan, Rosanne Freak-Poli, Danijela Gasevic, Jacqueline Scali, Karen Ritchie, Marie-Laure Ancelin, Alice Owen

► **To cite this version:**

Tagrid Abdullah Alharbi, Joanne Ryan, Rosanne Freak-Poli, Danijela Gasevic, Jacqueline Scali, et al.. The Association of Weight Loss, Weight Status, and Abdominal Obesity with All-Cause Mortality in Older Adults. *Gerontology*, 2022, pp.1-9. 10.1159/000522040 . hal-03630605

HAL Id: hal-03630605

<https://hal.umontpellier.fr/hal-03630605>

Submitted on 7 Jun 2022

HAL is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers.

L'archive ouverte pluridisciplinaire **HAL**, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.



Distributed under a Creative Commons Attribution - NonCommercial | 4.0 International License

The Association of Weight Loss, Weight Status, and Abdominal Obesity with All-Cause Mortality in Older Adults

Tagrid Abdullah Alharbi^a Joanne Ryan^a Rosanne Freak-Poli^{a, b}
Danijela Gasevic^{a, c} Jacqueline Scali^d Karen Ritchie^{d, e} Marie-Laure Ancelin^d
Alice J. Owen^a

^aSchool of Public Health and Preventive Medicine, Monash University, Melbourne, VIC, Australia; ^bDepartment of Epidemiology, Erasmus Medical Centre, Rotterdam, The Netherlands; ^cUsher Institute, University of Edinburgh, Edinburgh, UK; ^dINM, Univ Montpellier, INSERM, Montpellier, France; ^eCentre for Clinical Brain Sciences, University of Edinburgh, Edinburgh, UK

Keywords

Weight loss · Obesity · Abdominal obesity · Mortality

Abstract

Objectives: The objectives of this study were to examine whether weight loss, weight status (based on body mass index [BMI] categories), and abdominal obesity (based on waist circumference [WC]) were associated with a 17-year mortality risk in community-dwelling older adults. **Methods:** Participants were 2,017 community-dwelling adults aged 65 years or above in the longitudinal Enquête de Santé Psychologique-Risques, Incidence et Traitement study. Self-reported weight loss was collected at baseline during face-to-face interviews. Bodyweight (kg), height (m), and WC (cm) were independently measured at the baseline. BMI was categorized as follows: underweight (BMI <18.5 kg/m²), normal weight (18.5–24.9 kg/m²), overweight (25–29.9 kg/m²), and obese (≥30 kg/m²). Abdominal obesity was defined by a WC of ≥102 cm in men and ≥88 cm in women. Adjusted Cox proportional hazards models were used to examine associations of weight loss, weight status, and abdominal obesity with all-cause mortality. **Results:** Over 17 years of follow-up (median 15.5 years), 812 participants died. Ab-

dominal obesity compared to nonabdominal obesity was associated with a 49% increased mortality risk (95% confidence interval (CI): 1.22–1.83). However, being overweight (but not obese) was associated with a 20% decreased risk (95% CI: 0.66–0.97) compared to a normal BMI. Gender did not affect these associations. In the whole cohort, self-reported weight loss at baseline was not associated with an increased mortality risk after adjusting for health and lifestyle factors. However, in men, a baseline self-reported recent weight loss of >3 kg was associated with a 52% increase in mortality risk (95% CI: 1.05–2.18) in a fully adjusted model. **Conclusion:** In community-dwelling adults aged ≥65 years, abdominal obesity was strongly associated with increased mortality risk. Being overweight appeared, however, to be protective against mortality. Modest self-reported weight loss was not associated with all-cause mortality in community-dwelling older adults after adjusting for health and lifestyle factors. However, men reporting recent weight loss of more than 3 kg may be at increased risk. The findings of this study support the use of WC, rather than BMI, as a predictor of mortality risk in older adults.

© 2022 The Author(s).
Published by S. Karger AG, Basel

Introduction

Obesity has emerged as a major public health issue, with the global prevalence having tripled since 1975 [1]. While obesity affects all age-groups, it is increasingly common in older adults [2]. While there is extensive evidence that obesity is associated with an increased risk of cardiovascular disease, cancer, and mortality in younger and middle-aged adults [3], the relationship between obesity and mortality in older adults requires further research [2]. A meta-analysis by Flegal et al. [4] found that a body mass index (BMI) in the overweight range (25–30 kg/m²) was associated with a lower risk of all-cause mortality in older adults compared to those of a healthy weight (BMI between 18.5 kg/m² and <25 kg/m²), while grade 1 obesity (BMI between 30 kg/m² and ≤35 kg/m²) had a similar mortality risk to those in the healthy BMI category. BMI is a crude measure of body composition, and a BMI of ≥30 kg/m² could encompass a wide range of proportions of lean mass to fat mass. Abdominal obesity, which is defined as abdominal fat accumulation measured by the waist circumference (WC) [5], may be a more appropriate measure of risk, particularly in older adults. Notably, the significant growth of the world's aging population means that health care systems face significant challenges regarding aging-related chronic diseases that are either associated with or exacerbated by obesity.

A meta-analysis of 17 studies reported a 67% increase in the risk of mortality with weight loss among older adults (≥60 years) [6]. Another recent systematic review and meta-analysis of 35 studies examining the association between weight change and all-cause mortality in older adults found that weight loss was associated with a 59% increase in the all-cause mortality risk [7]. However, these previous studies did not include consideration of whether baseline weight or abdominal obesity influenced the mortality risks associated with weight loss in community-dwelling older adults. As obesity and abdominal obesity could differentially impact the association between weight loss and mortality in later life, this requires further investigation.

As identified in the most recent systematic review [7], only four studies [8–11] investigated the association of weight loss with mortality stratified by the baseline BMI. The findings from this limited number of studies suggested that there may be a stronger association between weight loss and mortality in older adults with an underweight/normal BMI compared to older adults with an overweight/obese BMI. However, the studies used differ-

ent BMI cutoffs, as follows: one used self-reported weight in early adulthood as the baseline weight [8] and another included only women [10]. It is also unknown whether there are gender-specific differences in the association between weight loss and mortality risk.

The aims of the current study were to examine the association of self-reported weight loss, weight status (based on BMI categories), and abdominal obesity (based on WC) with all-cause mortality in a large cohort of community-dwelling older adults and to determine whether baseline BMI, WC, or gender affected the association between weight loss and mortality.

Materials and Methods

Study Population

This study utilized data from the population-based, longitudinal Enquête de Santé Psychologique-Risques, Incidence et Traitement (ESPRIT) study [12]. Noninstitutionalized French-speaking, older adults aged ≥65 years who resided in the Montpellier district in southern France were eligible for enrolment in ESPRIT. The individuals who met the criteria were randomly drawn from 15 electoral rolls across the Montpellier district between March 1999 and February 2001. The ESPRIT study protocol was approved by the Ethical Committee of the University Hospital of Kremlin-Bicêtre. Further details of the ESPRIT study have been reported elsewhere [12, 13]. Of the 2,270 participants who were recruited for the ESPRIT study, 241 were excluded from this analysis because they were missing data on recent weight loss, and a further 12 were missing data related to BMI and WC. This left 2,017 participants, who constitute the sample for this study.

Weight Loss, BMI, and WC Measurements

Self-reported weight loss in the last 3 months was collected as a baseline during face-to-face interviews. Bodyweight (kg), height (m), and WC (cm) [14] were independently measured at baseline in lightly dressed participants. BMI was computed as weight (kg) divided by height (m) squared. WC was measured between the lower rib margin and the iliac crest following a normal expiration. BMI was categorized according to the World Health Organization's definitions as follows: underweight (BMI <18.5 kg/m²), normal weight (18.5–24.9 kg/m²), overweight (25–29.9 kg/m²), and obese (≥30 kg/m²) [15]. Abdominal obesity was defined by a WC of ≥102 cm in men and ≥88 cm in women [15].

All-Cause Mortality

All-cause mortality was ascertained for all participants over a maximum of 17 years of follow-up. Date and cause of death were determined from medical records and death registries, as detailed previously [16, 17].

Covariates

Sociodemographic characteristics, lifestyle, and physical health factors were collected at the baseline using a standardized health questionnaire. This included age, gender, smoking status (never smoked, past/current smoker), alcohol consumption (<24 g/day,

≥24 g/day), education level attained (completion of secondary school education), and living situation (alone or with others). Through face-to-face interviews, self-reported information on the history of ischemic pathologies (angina pectoris, myocardial infarction, stroke, cardiovascular surgery, arthritis, and vascular disease), chronic conditions (high blood pressure, high cholesterol, diabetes, thyroid problems, and asthma), recent cancer, and the diagnosis of current major depressive disorder, according to DSM-IV criteria [12], was collected using the Mini-International Neuropsychiatric Interview (French version 5.00). The depressive symptom severity was assessed using the 20-item inventory by the Centre for Epidemiologic Studies-Depression scale [18], with a cutoff of 16 indicating moderate to severe depressive symptoms.

Statistical Analysis

Summary statistics were reported to describe the frequencies or means (\pm standard deviation) of the baseline characteristics. The χ^2 or ANOVA tests were applied to compare the distributions of categorical and continuous variables, respectively, across the weight loss groups. Cox proportional hazard regression models were used to examine the hazard ratio (HR), 95% confidence interval (CI), and p values for the associations of weight loss, weight status (based on BMI categories), and abdominal obesity (based on WC) with all-mortality. As the mortality risk increases with age, age was used as the time scale, and a baseline age was used as the time origin [19]. The following series of models were constructed: model 1 was adjusted for gender; model 2 was additionally adjusted for education, living alone, smoking, alcohol consumption, depression, ischemic pathologies, chronic conditions, and recent cancer; and model 3 was additionally adjusted for the baseline BMI when analyzing self-reported weight loss and abdominal obesity and for baseline WC when analyzing the weight status. We performed a stratified analysis to examine whether the association between self-reported weight loss and mortality varied by gender or baseline BMI or WC categories.

Sensitivity analyses were undertaken using the final models and repeated with the following changes: (1) participants who died within the first year of the follow-up were excluded to ensure these early deaths did not affect the observed associations and (2) the 5-year follow-up analysis was restricted to test whether shorter follow-up periods influenced the associations. The statistical data analysis was conducted using STATA version 16 (StataCorp, College Station, TX, USA).

Results

There were 2,017 participants (Table 1) with an average age of 73.3 (standard deviation 5.7) years, and the median follow-up time was 15 years (IQR: 11.2–15.7). During this time, 812 (40.3%) participants died. At the baseline, there were 114 (5.6%) participants who reported a recent weight loss of >3 kg, while 219 (10.9%) participants reported having lost between 1 kg and 3 kg.

Participants who lost between 1 kg and 3 kg had a lower baseline weight compared to the participants who lost >3 kg or did not lose any weight. There was a similar pat-

tern for the BMI where the participants who lost between 1 kg and 3 kg had a lower BMI than the participants who lost >3 kg or did not report any weight loss.

The proportion of participants with chronic conditions or cardiovascular ischemic pathologies significantly differed according to the weight loss category, as did depressive symptoms. In all cases, the proportions of individuals with these conditions were highest among individuals with a recent weight loss of >3 kg (Table 1).

Association of BMI and WC with All-Cause Mortality

Compared to a “normal” baseline BMI, being underweight or obese was associated with an increased mortality risk in the minimally adjusted analysis only (Table 2, Model 1). After adjusting for lifestyle, health factors, and baseline WC, there was no significant association between being underweight or obese and the mortality risk. On the other hand, in the fully adjusted models, being overweight was associated with a 20% lower mortality risk (HR: 0.80, 95% CI: 0.66–0.97, $p = 0.02$) compared to a normal weight. The gender did not affect this result ($p = 0.71$). Abdominal obesity was significantly associated with an increased all-cause mortality risk (HR: 1.37, 95% CI: 1.17–1.61, $p < 0.001$), even after the multivariate analysis (HR: 1.49, 95% CI: 1.22–1.83, $p < 0.001$) in both women and men (Table 2).

Association of Self-Reported Weight Loss and Mortality

Compared to no weight loss, a self-reported weight loss of 1–3 kg was not associated with all-cause mortality in either the minimally adjusted or fully adjusted multivariate models (Table 3). In contrast, the participants who self-reported a recent weight loss of >3 kg at the baseline compared to those who reported no weight loss had a 44% increased risk of mortality (HR: 1.44, 95% CI: 1.09–1.88, $p = 0.008$) when adjusting for age and gender. However, following additional adjustments for other characteristics, including education, living alone, smoking, alcohol consumption, depression, ischemic pathologies chronic conditions, and a recent cancer diagnosis, the association between weight loss and all-cause mortality was no longer significant ($p = 0.09$). A further adjustment for the baseline BMI categories did not change this finding ($p = 0.07$).

However, the association between weight loss and mortality was affected by gender with some evidence of interaction (Table 3). In the fully adjusted multivariate models, male participants who lost >3 kg had a 52% higher risk of mortality (HR: 1.52, 95% CI: 1.05–2.18, $p = 0.03$) compared to those who reported no weight loss. No sig-

Table 1. Baseline characteristics of the participants according to baseline self-reported weight loss in 2,017 older adults

Characteristic	Total	Weight loss			p value
		lost >3 kg	lost 1–3 kg	no weight loss	
Total, <i>n</i> (%)	2,017	114 (5.6)	219 (10.9)	1,684 (83.5)	
Gender <i>n</i> (%)					
Women	1,183 (58.7)	67 (58.8)	135 (61.6)	981 (58.3)	0.63
Men	834 (41.3)	47 (41.2)	84 (38.4)	703 (41.8)	
Age, years, mean (±SD)	73.3±5.7	74.1±6.1	73.1±6.0	73.3±5.6	0.27
Age-group, <i>n</i> (%)					
65–74	1,276 (63.3)	68 (59.7)	148 (67.6)	1,060 (62.9)	0.29
≥75	741 (36.7)	46 (40.4)	71 (32.4)	624 (37.1)	
Weight, kg, mean (±SD)	67.3±12.2	67.6±11.9	65.0±11.2	67.5±12.3	0.01
BMI mean (±SD)	25.1±3.7	25.1±3.7	24.4±3.1	25.1±3.7	0.03
BMI group, kg/m ² , <i>n</i> (%)					
Underweight <18.5	43 (2.2)	3 (4.6)	3 (7.01)	37 (2.21)	0.26
Normal weight 18.5–24.9	1,067 (53.3)	57 (50.4)	130 (59.9)	880 (52.6)	
Overweight 25–29.9	718 (35.8)	40 (35.4)	73 (33.6)	605 (36.1)	
Obese ≥30	175 (8.7)	13 (11.5)	11 (5.07)	151 (9.03)	
WC, mean (±SD)	88.9±12.1	89.6±12.8	87.5±11.2	88.9±12.1	0.24
WC group					
Nonabdominal obesity	1,274 (70.7)	70 (70)	147 (75.8)	1,057 (70.1)	0.26
Abdominal obesity	528 (29.3)	30 (30)	47 (24.2)	451 (29.9)	
Smoking, <i>n</i> (%)					
Never smoked	1,179 (59.6)	64 (56.6)	133 (61.6)	982 (59.6)	0.68
Past/current smoker	799 (40.4)	49 (43.4)	83 (38.4)	667 (40.5)	
Alcohol consumption, <i>n</i> (%)					
Low-moderate (≤24 g/day)	1,608 (81.4)	92 (80.7)	173 (80.5)	1,343 (81.6)	0.91
High (≥24 g/day)	367 (18.6)	22 (19.3)	42 (19.5)	303 (18.4)	
Education level, <i>n</i> (%)					
Not completed at least secondary school	1,347 (66.9)	80 (70.2)	135 (61.9)	1,132 (67.3)	0.21
Completed at least secondary school	668 (33.1)	34 (29.8)	83 (38.1)	551 (32.7)	
Living alone, <i>n</i> (%)					
Lives with others	1,462 (72.6)	87 (76.3)	161 (73.5)	1,214 (72.2)	0.59
Lives alone	553 (27.4)	27 (23.7)	58 (26.5)	468 (27.8)	
Chronic conditions, ^a <i>n</i> (%)					
No	1,245 (61.7)	57 (50)	133 (60.7)	1,055 (62.7)	0.03
Yes	772 (38.3)	57 (50)	86 (39.3)	629 (37.4)	
Recent cancer, <i>n</i> (%)					
No	1,962 (97.3)	108 (94.7)	216 (98.6)	1,638 (97.3)	0.12
Yes	55 (2.7)	6 (5.3)	3 (1.4)	46 (2.7)	
Cardiovascular ischemic pathologies, ^b <i>n</i> (%)					
No	1,710 (84.8)	85 (74.5)	190 (86.8)	1,435 (85.2)	0.006
Yes	307 (15.2)	29 (25.4)	29 (13.2)	249 (14.8)	
Depressive symptoms (CES-D ≥ 16 or MDD), <i>n</i> (%)					
No	1,378 (69.1)	60 (52.6)	143 (65.9)	1,175 (70.6)	<0.001
Yes	618 (30.9)	54 (47.4)	74 (34.1)	490 (29.4)	

BMI categories: underweight (<18.5), normal (18.5–24.9), overweight (25–29.9), obese (≥30). WC groups: nonabdominal obesity (low WC: <102 cm for men and <88 cm for women), abdominal obesity (high WC: ≥102 cm for men and ≥88 cm for women). CES-D, Centre for Epidemiologic Studies Depression; MDD, major depressive disorder; *N*, number of observations; SD, standard deviation. *p* values are from ANOVA (continuous variables) and χ^2 tests (categorical variables). ^aMore than one chronic illness (high blood pressure, high cholesterol, diabetes, thyroid problems, asthma). ^bAt least one of the following: angina pectoris, myocardial infarction, stroke, cardiovascular surgery, arthritis, vascular disease.

Table 2. The association between baseline self-reported weight loss and all-cause mortality in 2,017 older adults

Weight loss	n/Events	Model 1			Model 2			Model 3*			Model 3				
		HR (95% CI)	p value	HR (95% CI)	p value	HR (95% CI)	p value	HR (95% CI)	p value	women n/ events	HR (95% CI)	p value	men n/ events	HR (95% CI)	p value
No weight loss	1,684/669	Reference	Reference	Reference	Reference	Reference	Reference	Reference	Reference	981/323	Reference	Reference	703/346	Reference	Reference
Loss 1–3 kg	219/86	1.04 (0.83–1.30)	0.71	1.07 (0.85–1.35)	0.56	1.09 (0.87–1.38)	0.45	1.14 (0.86–1.61)	0.29	135/48	1.14 (0.86–1.61)	0.29	84/38	1.01 (0.71–1.44)	0.96
Loss >3 kg	114/57	1.44 (1.09–1.88)	0.008	1.27 (0.97–1.68)	0.09	1.29 (0.98–1.70)	0.07	1.09 (0.72–1.68)	0.66	67/23	1.09 (0.72–1.68)	0.66	47/34	1.52 (1.05–2.18)	0.03

Model 1: gender (and age as the time scale in the model). Model 2: model 1 + adjusted for education, living status, smoking, alcohol consumption, depression, ischemic pathologies, chronic conditions, and recent cancer. Model 3: model 2 + baseline BMI categories. *p value for the interaction term between gender and weight loss was <0.001.

Table 3. Association of BMI and WC with all-cause mortality in 2,017 older adults

BMI categories	n/Events	Model 1			Model 2			Model 3			Model 3			Model 3		
		HR (95% CI)	p value	HR (95% CI)	p value	HR (95% CI)	p value	HR (95% CI)	p value	women n/ events	HR (95% CI)	p value	men n/ events	HR (95% CI)	p value	
Underweight	43/20	1.55 (0.99–2.43)	0.06	1.40 (0.88–2.23)	0.15	1.30 (0.77–2.19)	0.31	1.36 (0.77–2.40)	0.30	38/16	1.36 (0.77–2.40)	0.30	5/4	0.90 (0.21–3.75)	0.90	
Normal	1,067/400	Reference	Reference	Reference	Reference	Reference	Reference	Reference	Reference	701/221	Reference	Reference	366/179	Reference	Reference	
Overweight	718/299	1.01 (0.87–1.18)	0.90	0.96 (0.82–1.12)	0.64	0.80 (0.66–0.97)	0.02	0.77 (0.57–1.06)	0.11	325/108	0.77 (0.57–1.06)	0.11	393/191	0.82 (0.64–1.06)	0.13	
Obesity	175/83	1.33 (1.05–1.70)	0.02	1.20 (0.94–1.54)	0.15	0.78 (0.56–1.09)	0.15	0.67 (0.42–1.08)	0.10	110/42	0.67 (0.42–1.08)	0.10	65/41	0.89 (0.56–1.40)	0.62	

WC	n/ events	Model 1			Model 2			Model 3			Model 3			Model 3		
		HR (95% CI)	p value	HR (95% CI)	p value	HR (95% CI)	p value	HR (95% CI)	p value	women n/ events	HR (95% CI)	p value	men n/ events	HR (95% CI)	p value	
Nonabdominal obesity	1,247/441	Reference	Reference	Reference	Reference	Reference	Reference	Reference	Reference	727/193	Reference	Reference	547/248	Reference	Reference	
Abdominal obesity	528/232	1.37 (1.17–1.61)	<0.001	1.31 (1.11–1.55)	0.001	1.49 (1.22–1.83)	<0.001	1.63 (1.21–2.20)	0.001	325/119	1.63 (1.21–2.20)	0.001	203/113	1.41 (1.06–1.88)	0.01	

BMI categories: underweight (<18.5), normal (18.5–24.9), overweight (25–29.9), obese (≥30). WC groups: nonabdominal obesity (low WC: <102 cm for men and <88 cm for women), abdominal obesity (high WC: ≥102 cm for men and ≥88 cm for women). Model 1: gender (and age as the timescale in the model). Model 2: model 1 + adjusted for education, living alone, smoking, alcohol consumption, depression, ischemic pathologies, chronic conditions, and recent cancer. Model 3: model 2 + adjusted for WC or BMI, respectively.

nificant association between weight loss and all-cause mortality was observed in women (Table 3).

Effect of Weight Loss and BMI/WC on Mortality Risk

The next step was to determine whether baseline BMI or WC modified the association between baseline recent weight loss and mortality risk. In the Cox regression models, neither interaction term between the baseline BMI and weight loss ($p = 0.53$) nor between baseline WC and weight loss ($p = 0.39$) was significant. Furthermore, no significant findings were noted in the stratified analyses (online suppl. Table S1; for all online suppl. material, see www.karger.com/doi/10.1159/000522040). Based on the gender differences in Table 3, gender-specific analyses were undertaken, indicating the association between self-reported recent weight loss and mortality risk was only significant for men. Moreover, the baseline BMI and WC did not affect the association between weight loss and mortality in men (online suppl. Table S1).

In the sensitivity analysis, excluding the deaths that occurred within the first year of the follow-up, the result did not vary in both the overall sample and women. However, it was no longer significant in men who lost >3 kg (HR: 1.17, 95% CI: 0.79–1.73, $p = 0.42$; online suppl. Table S2). Moreover, by limiting the duration of the follow-up to 5 years and adjusting for all covariates, we found statistically significant results (self-reported weight loss of 1–3 kg: HR: 1.76, 95% CI: 1.13–2.73, $p = 0.01$; and lost >3 kg: HR: 1.81, 95% CI: 1.03–3.20, $p = 0.04$; online suppl. Table S3).

Discussion

In this large population-based study of community-dwelling older adults, for the men, a self-reported weight loss of more than 3 kg was associated with a 52% increased mortality risk compared to those who reported no weight loss. However, this was not true for the women, and there was no effect on the baseline BMI or WC categories. Abdominal obesity was associated with a 49% increased mortality risk, while being overweight (but not obese) was associated with a 20% reduced risk compared to a normal weight.

In the present study, we found no overall association between self-reported weight loss and all-cause mortality after considering lifestyle and health factors. This contrasts with the findings of several previous studies and meta-analyses, which reported a significant association between weight loss and all-cause mortality [6, 7]. This

could arise from several factors. First, it may be related to the follow-up period, as a previous meta-analysis demonstrated stronger associations between weight loss and mortality in studies with shorter follow-up periods [7]. Indeed, this is supported by the findings of our study. In the sensitivity analysis, when we limited the follow-up period to 5 years, we observed that a >3 kg weight loss was associated with an 81% increased risk of mortality. In addition, we observed a lower increased risk for individuals who lost between 1 kg and 3 kg. Therefore, weight loss appears to be a short-term risk factor for mortality. Second, weight loss, particularly if unintentional, can be an indication of impaired health and an underlying disease, which explains the association with an increased risk of mortality. This emphasizes the importance of adjusting for health factors and comorbid conditions, as well as obtaining information on whether the weight loss was intentional or unintentional. Our study, similar to the vast majority of the previous ones, did not distinguish between intentional and unintentional weight loss [7]. However, there is some evidence that only unintentional weight loss could increase the risk [20]. Our analysis adjusted for a range of comorbid conditions, including cancer [21, 22], unlike several previous studies. It is also possible that adjusting for health condition factors in our multivariate models may have impacted the association, as some of these factors may be biological intermediates in the causal pathway between body weight and mortality rather than confounders [23, 24]. Finally, the age of the participants may also account for differences in the findings between studies.

Our analysis indicated that gender affected the association between self-reported weight loss and all-cause mortality, which was only significant in men. Otsuka et al. [25] examined weight changes in Japanese community-dwelling adults aged 40–79 years and found that the weight of men started to decline when they were aged in their mid-50s, and women's weight declined when they were in their late-40s [25]. This suggests that there may be gender differences in age-related weight change. However, previous meta-analyses [6, 26] found that of the studies that considered gender differences, the association between weight loss and all-cause mortality was similar in both men and women. We did not find any effect in the initial baseline BMI categories or WC categories; however, we assessed these categories after the weight loss occurred. Due to the lack of previous research on this subject [7], further research is required to fully assess whether BMI or WC categories prior to weight loss influence the relationship between weight loss and mortality.

The associations observed in our study between BMI and all-cause mortality in older adults suggest that being overweight (but not obese) compared to a normal weight was associated with a decreased all-cause mortality risk. This aligns with meta-analyses on the BMI-mortality relationship [4, 27], which reported reduced risks of all-cause mortality in older adults classified as overweight compared to a normal BMI. Additionally, a recent scoping review of observational studies in a community-dwelling population aged 65 years or over reported a reduced risk of mortality for overweight status compared with normal weight status [28]. The World Health Organization BMI classification for body weight (underweight, normal, overweight, obese) is based on data from individuals 18 years and over. The relevance of these classifications for older adults may differ, given the well-known physiological changes observed in body composition with aging, such as loss of muscle and bone mass and fat gain [29]. This may have implications for weight management clinical guidelines and suggests the need for careful consideration when using above “normal” BMI to inform health decision-making for community-dwelling adults aged over 65 years.

In our study, abdominal obesity was associated with an increased risk of all-cause mortality. Our finding was consistent with previous studies that examined WC and mortality risk [30]. A meta-analysis of 29 cohort studies by de Hollander et al. [31] reported a greater mortality risk for a higher WC (>88 cm women and >102 cm men) among adults aged between 65 and 74 years. With aging, body fat tends to increase, especially in the abdominal area, which may be linked to an increased metabolic disease risk [32]. This supports mounting evidence that indicates that WC may be a better predictor of all-cause mortality risk compared to BMI in both men and women [33, 34].

The strengths of this study include a large, population-based, prospective study with 17-year follow-up of both clinical and environmental factors, which enables extensive covariate adjustment. An additional strength was the validity of the mortality status of the participants, which was obtained using death registries and medical records. In addition, the present study used objectively assessed obesity measures of BMI and WC.

The limitations of this study include self-reporting of weight loss occurring prior to enrolment, which is subject to recall error and an underestimation of the association. Studies have reported that there may be variations between the measured weight and self-reported weight [35, 36]. In addition, the participants were not asked about

whether their weight loss was intentional or unintentional, which may have influenced the strength of the association. Unintentional weight loss in later life may reflect an undiagnosed disease or undernutrition, which is linked to mortality. Intentional weight loss may be beneficial for comorbidities, disabilities, and overall health. Physical activity, which has been identified as an important potential confounder in a recent meta-analysis [7], was not available in this study. Furthermore, there were very few underweight participants (3.15% [$n = 41$] women and 0.53% [$n = 5$] men); therefore, the study may be underpowered to examine associations with the underweight status. Finally, a general limitation of cohort studies involving older adults, as in ESPRIT, is selection bias, as study participants reached a minimum of 65 years and needed to be in sufficiently good health to participate in the study. Participants in ESPRIT were also community-dwelling when they were recruited, which precludes generalization of the findings to frail or institutionalized individuals.

In conclusion, by conducting this study of community-dwelling adults aged ≥ 65 years, we found that self-reported weight loss was not associated with all-cause mortality after adjusting for health and lifestyle factors. However, men who reported a weight loss of more than 3 kg had an increased mortality risk. Although abdominal obesity was associated with a greater mortality risk in both genders, being overweight, compared to a normal weight, reduced the risk. These findings highlight the importance of healthcare providers to monitor the weight status in older adults to identify early individuals at risk of weight loss, which would initiate support strategies to maintain a stable weight. The findings of this study also support the use of central obesity to examine the mortality risk rather than a high BMI among older adults.

Acknowledgments

The authors thank the other investigators, staff, and participants of the ESPRIT trial for their valuable contributions.

Statement of Ethics

This secondary analysis of data was approved by the Monash University Human Research Ethics Committee, reference number: 22290. Written informed consent to participate in ESPRIT was obtained by ESPRIT investigators.

Conflict of Interest Statement

The authors have no conflicts of interest to declare.

Funding Sources

The ESPRIT project is financed by the regional government of Languedoc-Roussillon, the Agence Nationale de la Recherche Project 07 LVIE 004, and an unconditional grant from Novartis. The funders had no role in the design and conduct of the study and in data collection, management, analysis, or interpretation of the data and were not involved with the writing, preparation, review, or approval of the article. Joanne Ryan is funded by a fellowship (APP1135727 from the National Health & Medical Research Council (NHMRC), Australia. Rosanne Freak-Poli is funded by an Australian Heart Foundation post-doctoral fellowship (101927).

References

- 1 World Health Organization. Obesity and overweight 2020 (online). 2021. Available from: <https://www.who.int/news-room/fact-sheets/detail/obesity-and-overweight> Accessed 2020 Dec 15.
- 2 Peralta M, Ramos M, Lipert A, Martins J, Marques A. Prevalence and trends of overweight and obesity in older adults from 10 European countries from 2005 to 2013. *Scand J Public Health*. 2018 Jul;46(5):522–9.
- 3 Adams KF, Schatzkin A, Harris TB, Kipnis V, Mouw T, Ballard-Barbash R, et al. Overweight, obesity, and mortality in a large prospective cohort of persons 50 to 71 years old. *N Engl J Med*. 2006 Aug 24;355(8):763–78.
- 4 Flegal KM, Kit BK, Orpana H, Graubard BI. Association of all-cause mortality with overweight and obesity using standard body mass index categories: a systematic review and meta-analysis. *JAMA* 2013;309:71–82.
- 5 National Institute of Health. Clinical guidelines on the identification, evaluation, and treatment of overweight and obesity in adults: the evidence report. National Institutes of Health. *Obes Res*. 1998;6(Suppl 2):51S–209S.
- 6 Cheng FW, Gao X, Jensen GL. Weight change and all-cause mortality in older adults: a meta-analysis. *J Nutr Gerontol Geriatr*. 2015; 34(4):343–68.
- 7 Alharbi TA, Paudel S, Gasevic D, Ryan J, Freak-Poli R, Owen AJ. The association of weight change and all-cause mortality in older adults: a systematic review and meta-analysis. *Age Ageing*. 2021;50(3):697–704.
- 8 Corrada MM, Kawas CH, Mozaffar F, Paganini-Hill A. Association of body mass index and weight change with all-cause mortality in the elderly. *Am J Epidemiol*. 2006;163:938–49.
- 9 Haugsgjerd TR, Dierkes J, Vollset SE, Vinknes KJ, Nygård OK, Seifert R, et al. Association between weight change and mortality in community living older people followed for up to 14 years. The hordaland health study (HUSK). *J Nutr Health Aging*. 2017;21(8):909–17.
- 10 Reynolds MW, Fredman L, Langenberg P, Magaziner J. Weight, weight change, mortality in a random sample of older community-dwelling women. *J Am Geriatr Soc*. 1999;47: 1409–14.
- 11 Schaap LA, Quirke T, Wijnhoven HAH, Visser M. Changes in body mass index and mid-upper arm circumference in relation to all-cause mortality in older adults. *Clin Nutr*. 2018;37:2252–9.
- 12 Ritchie K, Artero S, Beluche I, Ancelin ML, Mann A, Dupuy AM, et al. Prevalence of DSM-IV psychiatric disorder in the French elderly population. *Br J Psychiatry*. 2004 Feb; 184:147–52.
- 13 Ritchie K, Jaussent I, Stewart R, Dupuy AM, Courtet P, Ancelin ML, et al. Association of adverse childhood environment and 5-HTTLPR genotype with late-life depression. *J Clin Psychiatry*. 2009 Sep;70(9):1281–8.
- 14 3C Study Group. Vascular factors and risk of dementia: design of the three-city study and baseline characteristics of the study population. *Neuroepidemiology*. 2003;22:316–25.
- 15 World Health Organization. *Obesity: preventing and managing the global epidemic: report of a WHO consultation*. Geneva, Switzerland: World Health Organization; 2000. xii.
- 16 Ryan J, Carrière I, Ritchie K, Stewart R, Toulemonde G, Dartigues JF, et al. Late-life depression and mortality: influence of gender and antidepressant use. *Br J Psychiatry*. 2008 Jan; 192(1):12–8.
- 17 Carrière I, Ryan J, Norton J, Scali J, Stewart R, Ritchie K, et al. Anxiety and mortality risk in community-dwelling elderly people. *Br J Psychiatry*. 2013 Sep;203(3):303–9.
- 18 Radloff LS. The CES-D scale: a self-report depression scale for research in the general population. *Appl Psychol Meas*. 1977;1:385–401.
- 19 Thiébaud AC, Bénichou J. Choice of time-scale in Cox's model analysis of epidemiologic cohort data: a simulation study. *Stat Med*. 2004;23:3803–20.
- 20 Harrington M, Gibson S, Cottrell RC. A review and meta-analysis of the effect of weight loss on all-cause mortality risk. *Nutr Res Rev*. 2009;22:93–108.
- 21 Hernández JL, Riancho JA, Matorras P, González-Macías J. Clinical evaluation for cancer in patients with involuntary weight loss without specific symptoms. *Am J Med*. 2003;114:631–7.
- 22 Metalidis C, Knockaert DC, Bobbaers H, Vanderschueren S. Involuntary weight loss. Does a negative baseline evaluation provide adequate reassurance? *Eur J Intern Med*. 2008;19:345–9.
- 23 Ajani UA, Lotufo PA, Gaziano JM, Lee IM, Spelsberg A, Buring JE, et al. Body mass index and mortality among US male physicians. *Ann Epidemiol*. 2004 Nov;14(10):731–9.
- 24 Manson JE, Bassuk SS, Hu FB, Stampfer MJ, Colditz GA, Willett WC. Estimating the number of deaths due to obesity: can the divergent findings be reconciled? *J Womens Health*. 2007;16:168–76.
- 25 Otsuka R, Kato Y, Nishita Y, Tange C, Tomida M, Nakamoto M, et al. Age-related changes in energy intake and weight in community-dwelling middle-aged and elderly Japanese. *J Nutr Health Aging*. 2016 Apr;20(4):383–90.
- 26 Karahalios A, English DR, Simpson JA. Change in body size and mortality: a systematic review and meta-analysis. *Int J Epidemiol*. 2017;46:526–46.
- 27 Winter JE, MacInnis RJ, Wattanapenpaiboon N, Nowson CA. BMI and all-cause mortality in older adults: a meta-analysis. *Am J Clin Nutr*. 2014;99:875–90.

Author Contributions

Tagrid Abdullah Alharbi: formal analysis, investigation, writing – original draft. Joanne Ryan, Rosanne Freak-Poli, Danijela Gasevic: supervision, writing – review and editing. Jacqueline Scali, Karen Ritchie, Marie-Laure Ancelin: resources, funding acquisition, writing – review and editing. Alice J. Owen: conceptualization, supervision, writing – review and editing.

Data Availability Statement

Any requests for data can be sent to the corresponding author.

- 28 Javed AA, Aljied R, Allison DJ, Anderson LN, Ma J, Raina P. Body mass index and all-cause mortality in older adults: a scoping review of observational studies. *Obes Rev*. 2020 Aug; 21(8):e13035.
- 29 Pearson MB, Bassey EJ, Bendall MJ. The effects of age on muscle strength and anthropometric indices within a group of elderly men and women. *Age Ageing*. 1985 Jul;14(4): 230–4.
- 30 Petursson H, Sigurdsson JA, Bengtsson C, Nilsen TI, Getz L. Body configuration as a predictor of mortality: comparison of five anthropometric measures in a 12 year follow-up of the Norwegian HUNT 2 study. *PLoS One*. 2011;6(10):e26621.
- 31 de Hollander EL, Bemelmans WJ, Boshuizen HC, Friedrich N, Wallaschofski H, Guallar-Castillón P, et al. The association between waist circumference and risk of mortality considering body mass index in 65- to 74-year-olds: a meta-analysis of 29 cohorts involving more than 58,000 elderly persons. *Int J Epidemiol*. 2012 Jun;41(3):805–17.
- 32 Atlantis E, Martin SA, Haren MT, Taylor AW, Wittert GA; Florey Adelaide Male Aging Study. Lifestyle factors associated with age-related differences in body composition: the Florey Adelaide Male Aging Study. *Am J Clin Nutr*. 2008 Jul;88(1):95–104.
- 33 Lahmann PH, Lissner L, Gullberg B, Berglund G. A prospective study of adiposity and all-cause mortality: the Malmö Diet and Cancer Study. *Obes Res*. 2002 May;10(5):361–9.
- 34 Jacobs EJ, Newton CC, Wang Y, Patel AV, McCullough ML, Campbell PT, et al. Waist circumference and all-cause mortality in a large US cohort. *Arch Intern Med*. 2010 Aug 9;170(15):1293–301.
- 35 Phimphasone-Brady P, Dorflinger LM, Ruser C, Bullock A, Godfrey KM, Hernandez D, et al. Self-report versus objective measurement of weight history: implications for pre-treatment weight gain. *J Behav Med*. 2019 Dec; 42(6):1142–7.
- 36 Dahl AK, Reynolds CA. Accuracy of recalled body weight: a study with 20-years of follow-up. *Obesity*. 2013 Jun;21(6):1293–8.