

Ambient ultraviolet radiation as a cardioprotective factor: A global and regional analysis

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Abstract

Background: Ambient ultraviolet radiation (UVR) has been found to have a greater cardioprotective effect than previously believed. This study aimed to quantitatively measure the role of UVR in protecting against the progression of cardiovascular disease (CVD) in general on a global and regional scale.

Methods: Population-level data on UVR, CVD incidence, aging, economic affluence, CVD genetic background (indexed with the Biological State Index, I_{BS}), obesity prevalence, and urbanization were collected and analysed. The correlation between UVR and CVD was examined using bivariate correlations, partial correlation, and stepwise multiple linear regression. Countries were grouped to investigate regional correlations between UVR and CVD, and Fisher's *r*-to-*z* transformation was used to compare correlation coefficients.

Results: UVR showed a significant inverse correlation with CVD incidence rates in bivariate correlation analyses globally ($r = -0.775$ and $r = -0.760$, $p < 0.001$), as well as within high-income ($r = -0.704$, $p < 0.001$) and low- and middle-income countries (LMIC) ($r = -0.851$, $p < 0.001$). These correlations remained significant even after controlling for confounding variables ($r = -0.689$ to -0.812 , $p < 0.001$). In stepwise regression models, UVR was found to be the most significant predictor of CVD incidence. The inverse correlation between UVR and CVD was stronger in LMICs compared to high-income countries ($z = -1.96$, $p < 0.050$).

Conclusions: Low ambient UVR may be a significant risk factor for the progression of CVD worldwide. The protective effect of UVR appears to be stronger in LMICs than in high-income countries, suggesting a greater impact of UVR on CVD prevention in these regions. These findings emphasize the need for further research into the mechanisms underlying the cardioprotective effects of UVR and the development of public health strategies to mitigate CVD risk associated with low UVR exposure.

KEYWORDS

ambient, cardiovascular disease (CVD), latitude, ultraviolet radiation, vitamin D

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1 | INTRODUCTION

Cardiovascular diseases (CVDs) are a wide range of disorders that affect the heart and blood vessels.¹ They are the leading cause of global disease burden,¹ with 55.5 million people diagnosed in 2019 alone, accounting for approximately one-third of all deaths worldwide.² This prevalence highlights the significant challenge that CVDs pose to global health, affecting both developed and developing nations.

Preventative measures targeting behavioral risk factors such as unhealthy diet, obesity, and physical inactivity are crucial for reducing the incidence of CVD.³ The risk factors for CVD are complex, involving a combination of cardiometabolic, behavioral, environmental, and social factors.⁴ Understanding the interplay between these factors is essential for understanding the complex causes of CVD.^{5,6} Cardiometabolic factors like hypertension and dyslipidaemia directly impact cardiovascular health, while behavioral factors such as diet, physical activity, and tobacco use also play important roles.⁴ Environmental factors, including air quality and exposure to pollutants, further contribute to the risk, and social determinants such as socioeconomic status and healthcare access complicate the picture.⁵ A comprehensive understanding of these factors is necessary to design targeted strategies for preventing and managing CVD in diverse populations.

The relationship between sunlight exposure, specifically ultraviolet radiation (UVR), and cardiovascular health has been the subject of debate in observational studies.^{7,8} Despite the known negative effects of UVR, historical records spanning over 6,000 years suggest that sunlight has been used therapeutically for cardiocirculatory and cardiovascular disorders.⁹ Natural heliotherapy is mentioned in the works of Hippocrates, ancient Roman health practices, and the practices of Islamic physicians in ancient Iran.¹⁰ Ambient sunlight plays a vital role in the production of vitamin D, which is essential for human health.⁷ When UVB light converts provitamin D3 into stable vitamin D3, it has a significant impact on our well-being. The amount of UVR radiation we receive varies based on factors such as seasonal changes, skin pigmentation, and distance from the equator.¹¹ Interestingly, there is a correlation between these variations in UVR radiation and changes in blood pressure.

Studies have shown that there is an inverse relationship between exposure to UVR radiation and both blood pressure and the incidence of CVD.^{7,12} As we move further away from the equator, the levels of ambient UVR radiation decrease, coinciding with an increase in the risk of CVD. In ecological studies, researchers have observed seasonal variations in the prevalence of conditions like CVD, hypertension, and metabolic syndrome.¹³ Rates of these conditions tend to be lower in the summer but higher in the winter.¹³

This intricate connection between ambient UVR exposure, vitamin D levels, and the risk of CVD highlights the importance of understanding these factors so that we can develop effective preventive strategies.^{14,15} Previous observational studies have shown an association between UVR radiation and CVD, but it's important to note that these studies may have been influenced by other factors.

We need a more comprehensive analysis to truly understand this relationship further.⁸

Following previous observational studies with confounding residue, the UVR-CVD relationship at population-level can be depicted through the graph below¹⁶:

In the graph provided in Figure 1, we can see that UVR exposure statistically explains 63.11% of the variance in CVD, suggesting that it may play a main protective role against CVD-related health issues. However, we need to approach these findings with caution,¹⁶ as we must consider other established factors like cardiometabolic, behavioral, environmental, and social risks that also contribute significantly to CVD.

Given the biases that have been observed in previous research, the aim of this study is to determine the independent influence of UVR exposure on the pathogenesis of CVD at a population level. To address potential biases in the UVR-CVD relationship, this study first quantified the global predictive capacity of UVR exposure for CVD incidence. Then, it conducted an analysis to minimize the impact of confounding factors such as economic affluence, aging, genetic predisposition to CVD, obesity, and urbanization on the UVR-CVD relationship. The ultimate result of this study is to reveal that UVR exposure is a significant and independent predictor for CVD incidence worldwide.

2 | MATERIALS AND METHODS

2.1 | Data collection and selection

This quantitative study gathered data from reputable international organizations and focused on country-specific information. A list of WHO member countries was compiled to match the seven variables considered in the study. Throughout this article, the terms "population" and "country" are used interchangeably to refer to a geographic region or territory.¹⁷ This approach is adopted for ease of writing and to enhance readability.

1. The independent variable, the average daily ambient ultraviolet radiation (UVR) level, which encompasses solar radiation in the 280–400 nm range, was obtained as the independent variable from the WHO Global Health Observatory (GHO).¹⁸ The UVR level is quantified in joules per square meter (J/m^2).¹⁸ In this study, population-level exposure is represented by the annual ambient erythemal weighted UVR, which takes into account the biological effects on human skin by applying the Erythema Action Spectrum. This measure is expressed in joules effective per square meter (J/m^2).¹⁸ It is calculated either from satellite data or by using a proxy such as latitudinal position.¹⁸
2. The dependent variable, the most recent CVD incidence rate (new cases per 100,000 population), was collected from the Institute for Health Metrics and Evaluation (IHME) at the University of Washington.² In addition to exploring the predictive role of UVR exposure on total CVDs, this study also examined its influence on

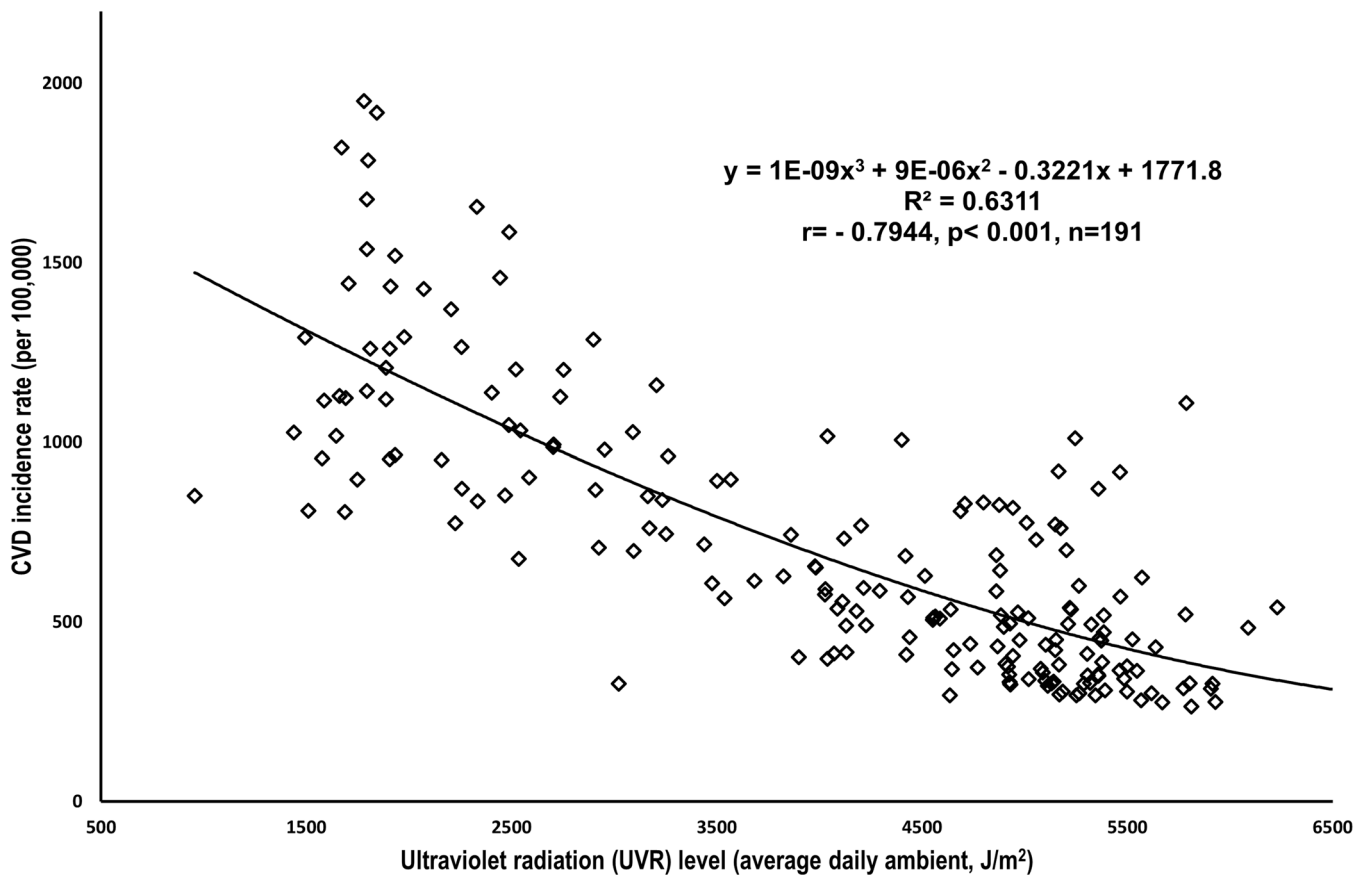


FIGURE 1 Plot to show the relationship between UV radiation and CVD incidence. Data source & definition: UV Radiation, expressed as the average daily ambient ultraviolet radiation level (in J/m^2), the World Health Organization; CVD incidence, cardiovascular disease (CVD) incidence rate, the number of new cases per 100,000, the Institute for Health Metrics and Evaluation. Raw data were incorporated into the Excel for the generating the scatterplots.

specific types of CVDs identified in previous research. The IHME provided incidence rates for all CVDs and 14 distinct types across various age groups. These types include ischemic heart disease, total stroke, ischemic stroke, intracerebral hemorrhage, subarachnoid hemorrhage, peripheral artery disease, atrial fibrillation and flutter, rheumatic heart disease, non-rheumatic valvular heart disease (including non-rheumatic degenerative mitral valve disease and non-rheumatic calcific aortic valve disease), cardiomyopathy and myocarditis (including myocarditis), and endocarditis. All available data were integrated into the study for comprehensive analysis.

In this study, we suggest that UVR generally helps protect against the development of CVDs. We have also chosen to explore the statistical predictive role of UVR because different types of CVDs have similar underlying mechanisms. While focusing on specific types of CVD could introduce bias, investigating the relationship between UVR and each individual CVD can still offer valuable evidence to support our main argument.

Previous studies have suggested that the relationship between UVR and CVD may be influenced by other factors,¹⁶ so it is important to account for these potential confounders.

This study included the following five variables that are recognized as major risk factors for CVD to isolate the independent effect of UVR exposure:

1. Aging, as indexed by life expectancy at age 65 years, was obtained from the United Nations.¹⁹ Age has been independently linked to the deterioration of cardiovascular health, significantly affecting the heart and arterial system.^{20,21}
2. Gross domestic product (GDP), converted to international dollars using purchasing power parity rates (GDP PPP), was extracted from the World Bank databank.²² Economic affluence impacts cardiovascular health through biological, behavioral, and psychosocial risk factors.²³ It is also associated with education, employment, life expectancy, and access to healthcare, which all influence the accuracy of CVD diagnosis.²⁴
3. The level of genetic background accumulation of CVD was measured using the Biological State Index (I_{bs}), which ranges from 0 to 1.0. Higher I_{bs} values indicate reduced natural selection and have been associated with the accumulation of harmful genes/mutations [54].²⁵ These genes/mutations are linked to conditions such as cancer, obesity, and type 1 diabetes. Including I_{bs} in the analysis helps address the impact of genetic predisposition on the UVR-CVD correlation.

4. The prevalence of obesity, defined as the percentage of the population aged 18+ with a BMI ≥ 30 kg/m², was obtained from the WHO Global Health Observatory.²⁶ Obesity increases the risk of CVD through factors such as atherosclerosis and comorbidities like hypertension, diabetes, and dyslipidaemia.^{27–29} It poses a multifactorial health challenge in terms of CVD prognosis.²⁷
5. Urbanization, measured as the percentage of the population living in urban areas, was gathered from the World Bank.²² Urbanization has been linked to CVD morbidity and mortality, and recently, it has also been associated with vitamin D deficiency among CVD patients.^{30,31} However, urban residents generally have better access to medical services, which aids in disease detection.

All of the above variables were extracted and matched with the WHO member list before being saved in Microsoft Excel[®] 2016 for further analysis. Each country was treated as an individual research subject, and the number of WHO member countries included for each variable may vary due to differences in data reporting.

2.2 | Statistical analyses

An ecological study was conducted to examine the relationship between UVR exposure and the incidence of CVD at the population level.^{32–37} To ensure accurate findings, several strategies were implemented: (1) different data analysis models were used for validation; (2) all seven variables were log-transformed to reduce homoscedasticity; and (3) the correlation between family size and CVD incidence rate was assessed globally and regionally.

1. Bivariate Correlations: Pearson's and nonparametric analyses were performed to determine the strength and direction of the associations between UVR exposure and the incidence rates of total CVDs as well as 14 specific types.
2. Partial Correlation Analysis: Using the Pearson moment-product approach, partial correlation was used to examine the independent relationships between UVR exposure and each of the 15 CVD incidence rates. This analysis considered confounding factors such as aging, GDP PPP, BMI, obesity, and urbanization to refine the examination of these relationships.
3. Independent Sample *t*-test: This test was conducted to compare the means of each CVD variable between countries with higher and lower UVR levels. Countries were divided into two groups based on UVR exposure, using a cut point of 4420.5 J/m², which is the world average published by IHME. The hypothesis was that significantly different UVR exposure may lead to significantly different CVD incidence rates worldwide, as well as within high- and upper-middle-income countries (HUIC) and low- and middle-income countries (LMIC).
4. Standard Multiple Linear Regression (stepwise): This analysis was used to identify variables that significantly predict CVD incidence rate. The independent variables included UVR exposure, aging,

GDP PPP, BMI, obesity, and urbanization. A total of 15 CVD incidence variables were entered as dependent variables.

5. Pearson's *r* Correlation: To explore regional associations between UVR and CVD incidence, countries were categorized based on different criteria:
 - World Bank Income Classifications: High-income, upper-middle-income, low-middle-income, and low-income. High-income countries formed one grouping, while low- and middle-income countries were combined into another (LMIC) as a new grouping. Pearson's *r* correlations were examined between UVR and CVD incidence in both groupings, and Fisher's *r*-to-*z* transformation was used to compare correlations.
 - United Nations Classifications: Developed versus developing countries. Fisher's *r*-to-*z* transformation was used to compare correlation coefficients between UVR and CVD incidence in developed versus developing countries.
 - Geographic and Socioeconomic Groupings: Seven groupings were analysed based on geographic distributions, per capita GDP levels, and cultural backgrounds: Asia Cooperation Dialog (ACD), Asia-Pacific Economic Cooperation (APEC), the Arab World, countries with English as the official language, Latin America, Latin America and the Caribbean (LAC), and the Organization for Economic Cooperation and Development (OECD).

SPSS v. 28 (SPSS Inc.) and Microsoft Excel 2016[®] were used for data analysis. The significance level was set at 0.05, with additional reporting at 0.01 and 0.001. The criteria for stepwise multiple linear regression analysis were a probability of *F* to enter ≤ 0.05 and a probability of *F* to remove ≥ 0.10 .

3 | RESULTS

Table 1 displays significant and consistent correlations between UVR and the incidence of CVD worldwide. These correlations are shown through both Pearson's *r* ($r = -0.775$, $p < 0.001$) and nonparametric analyses ($r = -0.760$, $p < 0.001$). The correlations hold true for both high-income countries (HIC: $r = -0.704$, $p < 0.001$) and low- to middle-income countries (LMIC: $r = -0.851$, $p < 0.001$). Even when controlling for five confounding variables in partial correlation analysis, the strong negative correlations persist ($r = -0.689$, -0.707 , and -0.812 , respectively, all $p < 0.001$), indicating that nations with higher UVR tend to have lower incidence rates of CVD, regardless of these factors.

Regarding specific types of CVD, Table 1 demonstrates moderate to strong and statistically significant inverse relationships between UVR and each of the 13 types examined, except for intracerebral hemorrhage and rheumatic heart disease.

Overall, this bivariate correlation analysis highlights moderate to strong inverse correlations between UVR and incidence rates of CVD. These correlations remain statistically significant even after considering potential confounders such as aging, GDP per capita (PPP), obesity, smoking prevalence (*I*_{bs}), and urbanization (Table 1).

TABLE 1 Bivariate (Pearson's r , nonparametric ρ) and partial correlation analyses to examine the relationships between ultraviolet radiation exposure and cardiovascular disease incidence rates (total and 13 types).

	All countries, $n = 191$		High- and Upper-Income Countries (HUIC), $n = 113$		Low- and Middle-Income Countries (LMIC), $n = 78$											
	Pearson's r	Nonparametric ρ	Pearson's r	Nonparametric ρ	Pearson's r	Nonparametric ρ										
Cardiovascular diseases, total	-0.775	<0.001	-0.760	<0.001	-0.689	<0.001	-0.704	<0.001	-0.700	<0.001	-0.851	<0.001	-0.679	<0.001	-0.812	<0.001
Atrial fibrillation and flutter	-0.764	<0.001	-0.754	<0.001	-0.648	<0.001	-0.755	<0.001	-0.738	<0.001	-0.707	<0.001	-0.631	<0.001	-0.505	<0.001
Cardio-myopathy and myocarditis	-0.764	<0.001	-0.659	<0.001	-0.636	<0.001	-0.727	<0.001	-0.695	<0.001	-0.706	<0.001	-0.465	<0.001	-0.448	<0.001
Endocarditis	-0.502	<0.001	-0.473	<0.001	-0.083	0.298	-0.366	<0.001	-0.361	<0.001	-0.146	0.165	-0.144	0.209	0.055	0.668
Intracerebral hemorrhage	0.153	<0.050	0.143	<0.050	-0.155	<0.050	0.206	<0.050	0.231	<0.050	-0.278	<0.010	-0.243	<0.050	-0.191	0.130
Ischemic heart disease	-0.647	<0.001	-0.657	<0.001	-0.517	<0.001	-0.507	<0.001	-0.469	<0.001	-0.479	<0.001	-0.670	<0.001	-0.769	<0.001
Ischemic stroke	-0.596	<0.001	-0.574	<0.001	-0.447	<0.001	-0.477	<0.001	-0.418	<0.001	-0.603	<0.001	-0.406	<0.001	-0.385	<0.010
Myocarditis	-0.764	<0.001	-0.659	<0.001	-0.636	<0.001	-0.727	<0.001	-0.695	<0.001	-0.706	<0.001	-0.465	<0.001	-0.448	<0.001
Nonrheumatic calcific aortic valve disease	-0.751	<0.001	-0.711	<0.001	-0.587	<0.001	-0.731	<0.001	-0.719	<0.001	-0.608	<0.001	-0.561	<0.001	-0.628	<0.001
Nonrheumatic degenerative mitral valve disease	-0.684	<0.001	-0.677	<0.001	-0.531	<0.001	-0.562	<0.001	-0.551	<0.001	-0.514	<0.001	-0.593	<0.001	-0.723	<0.001
Nonrheumatic valvular heart disease	-0.791	<0.001	-0.755	<0.001	-0.651	<0.001	-0.742	<0.001	-0.729	<0.001	-0.643	<0.001	-0.625	<0.001	-0.779	<0.001
Peripheral artery disease	-0.756	<0.001	-0.711	<0.001	-0.628	<0.001	-0.737	<0.001	-0.718	<0.001	-0.727	<0.001	-0.575	<0.001	-0.396	<0.001
Rheumatic heart disease	0.585	<0.001	0.601	<0.001	0.261	<0.001	0.443	<0.001	0.431	<0.001	0.166	0.115	0.520	<0.001	0.374	<0.010
Stroke	-0.508	<0.001	-0.486	<0.001	-0.460	<0.001	-0.358	<0.001	-0.273	<0.010	-0.607	<0.001	-0.484	<0.001	-0.428	<0.001

(Continues)

TABLE 1 (Continued)

	All countries, n = 191			High- and Upper-Income Countries (HUIC), n = 113			Low- and Middle-Income Countries (LMIC), n = 78											
	Pearson's <i>r</i>	Nonparametric ρ	Partial correlation <i>r</i>	Pearson's <i>r</i>	Nonparametric ρ	Partial correlation <i>r</i>	Pearson's <i>r</i>	Nonparametric ρ	Partial correlation <i>r</i>									
	Sig. <i>p</i>	Sig. <i>p</i>	Sig. <i>p</i>	Sig. <i>p</i>	Sig. <i>p</i>	Sig. <i>p</i>	Sig. <i>p</i>	Sig. <i>p</i>	Sig. <i>p</i>									
Subarachnoid hemorrhage	-0.633	<0.001	-0.656	<0.001	-0.441	<0.001	-0.526	<0.001	-0.579	<0.001	-0.507	<0.001	-0.654	<0.001	-0.536	<0.001	-0.490	<0.001
Ageing e(65)	-0.560	<0.001	-0.550	<0.001	-	-	-0.443	<0.001	-0.455	<0.001	-	-	-0.375	<0.001	-0.315	<0.010	-	-
GDP PPP	-0.612	<0.001	-0.629	<0.001	-	-	-0.532	<0.001	-0.541	<0.001	-	-	-0.574	<0.001	-0.439	<0.001	-	-
<i>I</i> _{bs}	-0.467	<0.001	-0.630	<0.001	-	-	-0.329	<0.001	-0.568	<0.001	-	-	-0.489	<0.001	-0.497	<0.001	-	-
Obesity %	-0.294	<0.001	-0.329	<0.001	-	-	-0.008	0.939	0.029	0.767	-	-	-0.231	<0.050	-0.239	<0.050	-	-
Urbanization	-0.439	<0.001	-0.466	<0.001	-	-	-0.323	<0.001	-0.292	<0.010	-	-	-0.253	<0.050	-0.163	0.156	-	-

Note: Significance levels: * $p < 0.05$, ** $p < 0.01$, *** $p < 0.001$. Control variable: Data source & definition: Ultraviolet Radiation (UVR), expressed as the average daily ambient ultraviolet radiation level (in J/m^2), the World Health Organization; Cardiovascular disease (CVD) incidence rate, the number of newly diagnosed cases per 100,000 in 2019, the Institute for Health Metrics and Evaluation; Ageing indexed with life expectancy at 65 year old in 2014, United Nations; Per capita GDP PPP, measured with the per capita purchasing power parity (PPP) value of all final goods and services produced within a territory in 2019, the World Bank; *I*_{bs} indexing the detrimental CVD genetic background accumulation in the past decades, downloaded from previous publications.³³ Obesity prevalence, measured with the percentage of population aged 18+ with BMI equal to or over 30 kg/m^2 in 2014, the World Health Organization. Urbanization, measured with the percentage of population living in urban area in 2019, the World Bank. All the data were log-transformed for correlation analysis in SPSS v 28.

This confirms the inclusion of these variables as controls when assessing the association between UVR and CVD incidence rates.

Table 2 shows that when using a cut-off of 4420.5 J/m^2 (the global median for ultraviolet radiation), independent sample t-tests revealed significant differences in mean values between country groups exposed to higher versus lower levels of ultraviolet radiation worldwide ($t = 21.074$, $p < 0.001$). This was true for both high- and low- and middle-income countries (HUIC: $t = 15.140$, $p < 0.001$; LMIC: $t = 13.136$, $p < 0.001$). Interestingly, countries with higher ultraviolet radiation levels above the cut-off point had significantly lower overall incidence rates of CVD compared to those with lower ultraviolet radiation worldwide ($t = -11.921$, $p < 0.010$). This trend was observed within both HUIC ($t = -7.373$, $p < 0.010$) and LMIC ($t = -7.618$, $p < 0.010$). However, this disparity was not observed in cases of intracerebral hemorrhage and rheumatic heart disease. The findings presented in Table 2 suggest that variations in ultraviolet radiation exposure may significantly influence the incidence rates of different types of cardiovascular disease.

Table 3 presents the results of the stepwise linear regression model, which highlights UVR as the most significant predictor of total CVD incidence globally. This finding holds true for both high-income and low- and middle-income countries, with adjusted R^2 values of 0.629, 0.519, and 0.748 respectively. The analysis shows a strong inverse relationship between UVR and CVD incidence rates worldwide, as well as in both high-income and low- and middle-income countries. The beta coefficients for these relationships are -0.607, -0.716, and -0.709 respectively, with p-values less than 0.010.

With the exception of endocarditis and intracerebral hemorrhage, UVR is found to be a significant predictor for all 12 types of CVD analysed. However, it is not always the most influential predictor. Additionally, there is a significant correlation between UVR and each of these 12 types of CVD.

Table 4 presents the correlation between UVR and CVD incidence across different country clusters, using Pearson's *r*. Generally, there was a consistent negative correlation between UVR and CVD incidence within each country grouping. However, the strength and significance of these correlations may vary depending on sample size and the predictive role of UVR in CVD incidence.

By applying Fisher's r-to-z transformation, it was found that the correlation between UVR and CVD incidence was significantly stronger in low- and middle-income countries (LMIC) compared to high-income countries ($z = -1.96$, $p < 0.050$). Additionally, the correlation in developing countries was significantly stronger than in developed countries ($z = -3.90$, $p < 0.001$). These results suggest that UVR may have a more pronounced impact on CVD incidence in LMIC than in high-income countries (Table 4).

4 | DISCUSSION

CVDs are a significant public health challenge influenced by various factors, including exposure to UVR. This study examines the relationship between UVR exposure and the incidence of 14 specific

TABLE 2 Independent samples t-test to compare the differences between the means in two country groups with the cut point of 4420.5 J/m² (median of ultraviolet radiations).

	All countries, n = 191, cut point 4420.5			High- and upper-income Countries (HUIC), n = 113, cut point 4420.5			Low- and middle-income Countries (LMIC), n = 68, cut point 4420.5					
	UVR			UVR			UVR					
	Mean >4420.5	Mean <4420.5	Mean difference, t	Sig. p	Mean > 4420.5	Mean <4420.5	Mean difference t	Sig. p	Mean >4420.5	Mean <4420.5	Mean difference t	Sig. p
UVR	5220.52	2791.07	21.074	<0.01	5185.66	2609.77	15.140	<0.01	5247.44	3386.76	13.136	<0.010
	CVD incidence rate			CVD incidence rate			CVD incidence rate					
	Mean > 4420.5	Mean <4420.5	Mean difference t	Sig. p	Mean > 4420.5	Mean <4420.5	Mean difference t	Sig. p	Mean >4420.5	Mean <4420.5	Mean difference t	Sig. p
Cardiovascular diseases, total	479.51	973.68	-11.921	<0.001	603.72	1031.81	-7.373	<0.001	383.63	782.68	-7.618	<0.001
Atrial fibrillation and flutter	26.82	82.02	-11.483	<0.001	37.38	91.85	-7.973	<0.001	18.66	49.73	-5.984	<0.001
Cardiomyopathy and myocarditis	12.42	18.60	-10.006	<0.001	14.04	20.19	-7.329	<0.001	11.17	13.38	-3.866	<0.001
Endocarditis	12.28	19.22	-5.448	<0.001	17.47	22.03	-2.572	<0.050	8.28	10.00	-2.082	<0.050
Intracerebral hemorrhage	43.23	38.08	1.497	0.136	44.04	34.09	2.274	<0.050	42.60	51.17	-1.385	0.170
Ischemic heart disease	172.70	424.43	-9.736	<0.001	249.20	433.21	-4.894	<0.001	113.64	395.56	-8.389	<0.001
Ischemic stroke	62.89	113.28	-7.365	<0.001	78.88	120.07	-3.985	<0.001	50.55	90.97	-5.257	<0.001
Myocarditis	12.42	18.60	-10.006	<0.001	14.04	20.19	-7.329	<0.001	11.17	13.38	-3.866	<0.001
Nonrheumatic calcific aortic valve disease	1.63	20.42	-7.923	<0.001	3.21	24.35	-6.111	<0.001	0.40	7.50	-2.406	<0.050
Nonrheumatic degenerative mitral valve disease	1.88	19.88	-6.417	<0.001	3.03	22.82	-4.274	<0.001	0.99	10.21	-4.895	<0.001
Nonrheumatic valvular heart disease	3.51	40.29	-8.174	<0.001	6.24	47.17	-5.923	<0.001	1.39	17.70	-3.506	<0.001
Peripheral artery disease	78.44	198.14	-10.738	<0.001	106.78	226.76	-7.903	<0.001	56.57	104.10	-2.105	<0.050
Rheumatic heart disease	57.03	19.28	9.946	<0.001	35.77	14.17	5.834	<0.001	73.43	36.05	-4.582	<0.001
Stroke	116.32	171.70	-5.982	<0.001	136.83	176.42	-2.875	<0.010	100.48	156.17	5.968	<0.001
Subarachnoid hemorrhage	10.20	20.34	-6.831	<0.001	13.92	22.26	-3.639	<0.001	7.33	14.03	-4.667	<0.001

Note: Significance levels: * $p < 0.05$, ** $p < 0.01$, *** $p < 0.001$. Data source and definition: Ultraviolet radiation (UVR), expressed as the average daily ambient ultraviolet radiation level (in J/m²), the World Health Organization; Cardiovascular disease (CVD) incidence rate, the number of newly diagnosed cases per 100,000 in 2019, the Institute for Health Metrics and Evaluation. All the data were not log-transformed for mean comparison in SPSS v 28.

TABLE 3 Stepwise multiple linear regression to identify the significant predictors of cardiovascular disease incidence rates (cardiovascular disease incidence rates [total and 14 types]).

	All Countries, n = 191				High- and Upper-middle Income Countries (HUIIC), n = 113				Low- and Low-middle Income Countries (LMIC), n = 68			
	Independent predictor	Adjusted R ²	Beta	Sig. p	Independent predictor	Adjusted R ²	Beta	Sig. p	Independent predictor	Adjusted R ²	Beta	Sig. p
Cardiovascular diseases, total	UV radiation	0.629	-0.607	<0.001	UV radiation	0.519	-0.716	<0.001	UV radiation	0.748	-0.709	<0.001
	l _{bs}	0.742	0.387	<0.001	Urbanization	0.561	-0.233	<0.001	l _{bs}	0.829	0.327	<0.001
	Ageing e(65)	Insignificant			l _{bs}	0.605	0.231	<0.001	Ageing e(65)	Insignificant		
	GDP PPP	Insignificant			Ageing e(65)	Insignificant			GDP PPP	Insignificant		
	Obesity %	Insignificant			GDP PPP	Insignificant			Obesity %	Insignificant		
Atrial fibrillation and flutter	Urbanization	Insignificant			Obesity %	Insignificant			Urbanization	Insignificant		
	UV radiation	0.597	-0.559	<0.001	UV radiation	0.577	-0.734	<0.001	GDP PPP	0.485	0.243	<0.050
	l _{bs}	0.747	0.443	<0.001	l _{bs}	0.628	0.255	<0.001	UV radiation	0.605	-0.384	<0.001
	Ageing e(65)	Insignificant			Urbanization	0.658	-0.193	<0.010	l _{bs}	0.664	0.347	<0.001
	GDP PPP	Insignificant			Ageing e(65)	Insignificant			Ageing e(65)	Insignificant		
Cardiomyopathy and myocarditis	Obesity %	Insignificant			GDP PPP	Insignificant			Obesity %	Insignificant		
	Urbanization	Insignificant			Obesity %	Insignificant			Urbanization	Insignificant		
	UV radiation	0.589	-0.505	<0.001	UV radiation	0.523	-0.617	<0.001	GDP PPP	0.442	0.293	<0.050
	Ageing e(65)	0.759	0.444	<0.001	Ageing e(65)	0.658	0.401	<0.001	UV radiation	0.535	-0.369	<0.001
	Obesity %	0.766	-0.138	<0.010	Obesity %	0.693	-0.177	<0.010	l _{bs}	0.573	0.335	<0.010
Endocarditis	l _{bs}	0.772	0.138	<0.050	Urbanization	0.706	-0.136	<0.050	Obesity %	0.607	-0.210	<0.050
	GDP PPP	Insignificant			GDP PPP	Insignificant			Ageing e(65)	Insignificant		
	Urbanization	Insignificant			l _{bs}	Insignificant			Urbanization	Insignificant		
	Ageing e(65)	0.616	0.697	<0.001	Ageing e(65)	0.440	5.325	<0.001	GDP PPP	0.216	0.374	<0.010
	Obesity %	0.657	0.224	<0.001	l _{bs}	0.485	3.054	0.003	Urbanization	0.253	0.241	<0.050
Intracerebral hemorrhage	UV radiation	Insignificant			UV radiation	Insignificant			UV radiation	Insignificant		
	GDP PPP	Insignificant			GDP PPP	Insignificant			Ageing e(65)	Insignificant		
	l _{bs}	Insignificant			Obesity %	Insignificant			l _{bs}	Insignificant		
	Urbanization	Insignificant			Urbanization	Insignificant			Obesity %	Insignificant		
	Ageing e(65)	0.203	-0.818	<0.001	Ageing e(65)	0.307	-0.630	<0.001	UV radiation	Insignificant		
Obesity %	l _{bs}	0.312	0.601	<0.001	Urbanization	0.384	-0.205	0.019	Ageing e(65)	Insignificant		
	l _{bs}	0.338	-0.203	<0.010	l _{bs}	0.416	0.305	0.002	GDP PPP	Insignificant		

TABLE 3 (Continued)

	All Countries, n = 191					High- and Upper-middle Income Countries (HUIC), n = 113					Low- and Low-middle Income Countries (LMIC), n = 68					
	Independent predictor	Adjusted R ²	Beta	Sig. p	Independent predictor	Adjusted R ²	Beta	Sig. p	Independent predictor	Adjusted R ²	Beta	Sig. p	Independent predictor	Adjusted R ²	Beta	Sig. p
	UV radiation	Insignificant			Obesity %	0.459	-0.262	0.001	l _{bs}	Insignificant			l _{bs}	Insignificant		
	GDP PPP	Insignificant			UV radiation	0.478	-0.238	0.007	Obesity %	Insignificant			Obesity %	Insignificant		
	Urbanization	Insignificant			GDP PPP	0.500	-0.263	<0.050	Urbanization	Insignificant			Urbanization	Insignificant		
Ischemic heart disease	UV radiation	0.460	-0.472	<0.001	UV radiation	0.280	-0.501	<0.001	UV radiation	0.714	-0.724	<0.001	UV radiation	0.714	-0.724	<0.001
	l _{bs}	0.581	0.327	<0.001	l _{bs}	0.322	0.227	<0.010	l _{bs}	0.761	0.255	<0.001	l _{bs}	0.761	0.255	<0.001
	Obesity %	0.595	0.151	<0.050	Obesity %	0.349	0.202	<0.050	Ageing e(65)	Insignificant			Ageing e(65)	Insignificant		
	Ageing e(65)	Insignificant			Urbanization	0.374	-0.185	<0.050	GDP PPP	Insignificant			GDP PPP	Insignificant		
	GDP PPP	Insignificant			Ageing e(65)	Insignificant			Obesity %	Insignificant			Obesity %	Insignificant		
	Urbanization	Insignificant			GDP PPP	Insignificant			Urbanization	Insignificant			Urbanization	Insignificant		
Ischemic stroke	UV radiation	0.367	-0.483	<0.001	UV radiation	0.235	-0.604	<0.001	UV radiation	0.354	-0.438	<0.001	UV radiation	0.354	-0.438	<0.001
	l _{bs}	0.468	0.495	<0.001	Ageing e(65)	0.302	-0.575	<0.001	l _{bs}	0.437	0.342	<0.010	l _{bs}	0.437	0.342	<0.010
	Ageing e(65)	0.483	-0.207	<0.050	l _{bs}	0.367	0.402	<0.001	Ageing e(65)	Insignificant			Ageing e(65)	Insignificant		
	GDP PPP	Insignificant			Obesity %	0.421	-0.251	0.003	GDP PPP	Insignificant			GDP PPP	Insignificant		
	Obesity %	Insignificant			GDP PPP	Insignificant			Obesity %	Insignificant			Obesity %	Insignificant		
	Urbanization	Insignificant			Urbanization	Insignificant			Urbanization	Insignificant			Urbanization	Insignificant		
Myocarditis	UV radiation	0.589	-0.505	<0.001	UV radiation	0.523	-0.617	<0.001	GDP PPP	0.442	0.293	<0.050	GDP PPP	0.442	0.293	<0.050
	Ageing e(65)	0.759	0.444	<0.001	Ageing e(65)	0.658	0.401	<0.001	UV radiation	0.535	-0.369	<0.001	UV radiation	0.535	-0.369	<0.001
	Obesity %	0.766	-0.138	<0.010	Obesity %	0.693	-0.177	<0.010	l _{bs}	0.573	0.335	<0.010	l _{bs}	0.573	0.335	<0.010
	l _{bs}	0.772	0.138	<0.050	Urbanization	0.706	-0.136	<0.050	Obesity %	0.607	-0.210	<0.050	Obesity %	0.607	-0.210	<0.050
	GDP PPP	Insignificant			GDP PPP	Insignificant			Ageing e(65)	Insignificant			Ageing e(65)	Insignificant		
	Urbanization	Insignificant			l _{bs}	Insignificant			Urbanization	Insignificant			Urbanization	Insignificant		
Nonrheumatic calcific aortic valve disease	GDP PPP	0.676	0.343	<0.001	UV radiation	0.526	-8.573	<0.001	UV radiation	0.574	-0.572	<0.001	UV radiation	0.574	-0.572	<0.001
	UV radiation	0.783	-0.405	<0.001	Ageing e(65)	0.608	4.548	<0.001	GDP PPP	0.626	0.251	<0.010	GDP PPP	0.626	0.251	<0.010
	Ageing e(65)	0.793	0.209	<0.010	GDP PPP	Insignificant			Obesity %	0.647	0.170	<0.050	Obesity %	0.647	0.170	<0.050
	Obesity %	0.800	0.106	<0.050	l _{bs}	Insignificant			Ageing e(65)	Insignificant			Ageing e(65)	Insignificant		
	l _{bs}	Insignificant			Obesity %	Insignificant			l _{bs}	Insignificant			l _{bs}	Insignificant		
	Urbanization	Insignificant			Urbanization	Insignificant			Urbanization	Insignificant			Urbanization	Insignificant		

(Continues)

TABLE 3 (Continued)

	All Countries, n = 191					High- and Upper-middle Income Countries (HUIC), n = 113					Low- and Low-middle Income Countries (LMIC), n = 68				
	Independent predictor	Adjusted R ²	Beta	Sig. p	Independent predictor	Adjusted R ²	Beta	Sig. p	Independent predictor	Adjusted R ²	Beta	Sig. p			
Nonrheumatic degenerative mitral valve disease	UV radiation	0.473	-0.528	<0.001	UV radiation	0.307	-0.575	<0.001	UV radiation	0.670	-0.686	<0.001			
	l _{bs}	0.557	0.336	<0.001	Obesity %	0.336	-0.263	<0.010	l _{bs}	0.726	0.280	<0.001			
	Ageing e(65)	Insignificant			l _{bs}	0.366	0.369	<0.001	Ageing e(65)	Insignificant					
	GDP PPP	Insignificant			Ageing e(65)	0.412	-0.305	<0.010	GDP PPP	Insignificant					
	Obesity %	Insignificant			GDP PPP				Obesity %	Insignificant					
Nonrheumatic valvular heart disease	Urbanization	Insignificant			Urbanization				Urbanization	Insignificant					
	UV radiation	0.626	-0.502	<0.001	UV radiation	0.528	-0.652	<0.001	UV radiation	0.729	-0.717	<0.001			
	GDP PPP	0.762	0.386	<0.001	l _{bs}	0.578	0.258	<0.001	l _{bs}	0.790	0.287	<0.001			
	l _{bs}	0.778	0.201	<0.001	Obesity %	0.598	-0.155	<0.050	Ageing e(65)	Insignificant					
	Urbanization	0.782	-0.102	<0.050	Ageing e(65)				GDP PPP	Insignificant					
Peripheral artery disease	Ageing e(65)	Insignificant			GDP PPP				Obesity %	Insignificant					
	Obesity %	Insignificant			Urbanization				Urbanization	Insignificant					
	UV radiation	0.584	-0.463	<0.001	UV radiation	0.555	-0.661	<0.001	l _{bs}	0.546	0.422	<0.001			
	Ageing e(65)	0.755	0.304	<0.001	Ageing e(65)	0.638	0.364	<0.001	UV radiation	0.645	-0.280	<0.010			
	l _{bs}	0.789	0.282	<0.001	Urbanization	0.669	-0.198	<0.010	GDP PPP	0.675	0.272	<0.010			
Rheumatic heart disease	GDP PPP	Insignificant			GDP PPP	Insignificant			Ageing e(65)	Insignificant					
	Obesity %	Insignificant			l _{bs}	Insignificant			Obesity %	Insignificant					
	Urbanization	Insignificant			Obesity %	Insignificant			Urbanization	Insignificant					
	GDP PPP	0.576	-0.466	<0.001	GDP PPP	0.363	-0.325	<0.001	GDP PPP	0.443	-0.280	<0.050			
	UV radiation	0.603	0.209	<0.001	l _{bs}	0.410	-0.299	<0.001	Ageing e(65)	0.501	-0.331	<0.010			
Stroke	Ageing e(65)	0.612	-0.196	<0.050	Urbanization	0.467	-0.286	<0.001	UV radiation	0.560	0.310	<0.010			
	l _{bs}	Insignificant			UV radiation	Insignificant			l _{bs}	Insignificant					
	Obesity %	Insignificant			Ageing e(65)	Insignificant			Obesity %	Insignificant					
	Urbanization	Insignificant			Obesity %	Insignificant			Urbanization	Insignificant					
	UV radiation	0.276	-6.860	<0.001	UV radiation	0.153	-0.628	<0.001	UV radiation	0.361	-0.419	<0.001			
Stroke	l _{bs}	0.359	7.092	<0.001	Ageing e(65)	0.268	-0.562	<0.001	l _{bs}	0.472	0.392	<0.001			
	Ageing e(65)	0.442	-5.021	<0.001	l _{bs}	0.353	0.477	<0.001	Ageing e(65)	Insignificant					

TABLE 3 (Continued)

	All Countries, n = 191			High- and Upper-middle Income Countries (HUIIC), n = 113			Low- and Low-middle Income Countries (LMIC), n = 68					
	Independent predictor	Adjusted R ²	Beta	Sig. p	Independent predictor	Adjusted R ²	Beta	Sig. p	Independent predictor	Adjusted R ²	Beta	Sig. p
	GDP PPP				Obesity %	0.451	-0.340	<0.001	GDP PPP	Insignificant		
	Obesity %				GDP PPP	0.476	-0.264	<0.050	Obesity %	Insignificant		
	Urbanization				Urbanization				Urbanization	Insignificant		
Subarachnoid hemorrhage	<i>I</i> _{bs}	0.624	0.626	<0.001	<i>I</i> _{bs}	0.363	0.502	<0.001	<i>I</i> _{bs}	0.615	0.613	<0.001
	UV radiation	0.713	-0.342	<0.001	UV radiation	0.479	-0.380	<0.001	UV radiation	0.711	-0.360	<0.001
	Ageing e(65)	Insignificant			Obesity %	0.558	-0.288	<0.001	Ageing e(65)	Insignificant		
	GDP PPP	Insignificant			Ageing e(65)	Insignificant			GDP PPP	Insignificant		
	Obesity %	Insignificant			GDP PPP	Insignificant			Obesity %	Insignificant		
	Urbanization	Insignificant			Urbanization	Insignificant			Urbanization	Insignificant		

Note: Significance levels: * $p < 0.05$, ** $p < 0.01$, *** $p < 0.001$. Data source and definition: Ultraviolet radiation (UVR), expressed as the average daily ambient ultraviolet radiation level (in J/m²), the World Health Organization; Cardiovascular disease (CVD) incidence rate, the number of newly diagnosed cases per 100,000 in 2019, the Institute for Health Metrics and Evaluation; Ageing indexed with life expectancy at 65 year old in 2014, United Nations; Per capita GDP PPP, measured with the per capita purchasing power parity (PPP) value of all final goods and services produced within a territory in 2019, the World Bank; *I*_{bs} indexing the detrimental CVD genetic background accumulation in the past decades, downloaded from previous publications³⁴; Obesity prevalence, measured with the percentage of population aged 18+ with BMI equal to or over 30 kg/m² in 2014, the World Health Organization. Urbanization, measured with the percentage of population living in urban area in 2019, the World Bank. All the data were log-transformed for correlation analysis in SPSS v 28.

TABLE 4 Ultraviolet radiation (UVR) level determining total cardiovascular disease incidence rate in different country groupings.

Country groupings	Pearson <i>r</i>	<i>p</i>
World Bank income classifications		
Low Income, <i>n</i> = 30	-0.684**	<0.001
Low Middle Income, <i>n</i> = 49	-0.846**	<0.001
Upper Middle Income, <i>n</i> = 51	-0.740**	<0.001
Low- and middle-income countries (LMIC), <i>n</i> = 130	-0.785**	<0.001
High Income, <i>n</i> = 60	-0.633**	<0.001
Fisher <i>r</i> -to- <i>z</i> transformation	LMIC versus High: <i>z</i> = - 1.96,	<i>p</i> < 0.050
United Nations common practice		
Developed, <i>n</i> = 44	0.025	0.872
Developing, <i>n</i> = 146	-0.582**	<0.001
Fisher <i>r</i> -to- <i>z</i> transformation	LMIC versus High: <i>z</i> = - 3.90,	<i>p</i> < 0.001
Countries grouped with various factors		
Asia Cooperation Dialog, <i>n</i> = 33	-0.776**	<0.001
Asia-Pacific Economic Cooperation, <i>n</i> = 19	-0.900**	<0.001
Arab World, <i>n</i> = 21	-0.872**	<0.001
English as Official Language, <i>n</i> = 54	-0.441**	<0.001
Latin America, <i>n</i> = 20	-0.632**	<0.010
Latin America and Caribbean, <i>n</i> = 33	-0.124	0.494
Organization for Economic Co-operation and Development, <i>n</i> = 38	-0.598**	<0.001

Note: Pearson *r* and nonparametric correlations within country groupings were reported. Data source & definition: Ultraviolet Radiation (UVR), expressed as the average daily ambient ultraviolet radiation level (in J/m²), the World Health Organization; Cardiovascular disease (CVD) incidence rate, the number of new cases per 100,000, the Institute for Health Metrics and Evaluation. All the data were log-transformed for correlation analysis.

types of CVD [53], highlighting UVR as a notable risk factor in their development.³⁸

1. UVR exposure is significantly correlated with a reduction in the overall incidence of CVD. When accounting for confounding factors such as aging, economic status, obesity, and urbanization, UVR explains 63.11% of the variance. This statistical association underscores the protective role of UVR across different types of CVD.
2. However, after adjusting for established confounders like aging and economic status, the impact of UVR on the incidence of total CVD slightly diminishes. Stepwise linear regression reveals that UVR explains 36.84% of the variance, while partial correlation

analysis attributes 47.47% to UVR, indicating a persistent albeit reduced association.

3. Interestingly, the protective effect of UVR appears to be more pronounced in developing countries compared to developed ones, suggesting varying impacts across different geographic and economic contexts.

Throughout human evolution, there has been significant exposure to sunlight, approximately half of each day.³⁹ While research has highlighted the adverse effects of UVR, such as skin cancer and photo-aging, recent insights challenge conventional wisdom.⁴⁰ Contrary to prevailing beliefs, reduced UVR exposure, rather than increased exposure, may contribute to the development of skin cancer, prompting a re-evaluation of the complex relationship between UVR and human health.^{36,41}

Ultraviolet radiation from sunlight is believed to provide cardiovascular benefits and has been used for over 6,000 years in the treatment of cardiocirculatory disorders.^{42,43} However, there is no scientific consensus on the consistent cardioprotective effects of UVR.

One established mechanism links UVR exposure to the synthesis of vitamin D in the skin,⁴⁴ which helps protect against CVD and high blood pressure. Another suggests that UVR stimulates the conversion of skin nitrogen oxides to nitric oxide (NO), promoting coronary vasodilation and exhibiting cardioprotective and antihypertensive properties.⁸ Although UV-A radiation does not promote vitamin D synthesis,⁴⁵ it has been associated with lower blood pressure.⁴⁰ Moreover, while greater UVR exposure may degrade bioactive folate metabolites,⁴⁶ including 5-methyltetrahydrofolate, known for its cardioprotective properties, this study focuses on the role of natural ambient UVR in maintaining and improving cardiovascular health, without delving into the specifics of the UVR spectrum and its controversial impact on vitamin D and folate synthesis.

Scragg conducted an ecological study that found a possible link between seasonal variations in UVR and the observed patterns of CVD.⁴⁷ In contrast to this descriptive observational approach, this study took a quantitative approach using population-level data. As the lead author of a recent systematic review, Scragg et al. raised concerns about previous studies that linked UVR to CVD, suggesting that other influencing risk factors may have confounded the results.⁴⁷ This study addresses this concern by using partial correlation and stepwise linear regression to independently investigate the role of UVR in predicting CVD incidence globally.

Many studies have examined the statistical relationship between UVR and blood pressure.¹² However, it is important to note that while high blood pressure is a significant risk factor for CVD,^{48,49} it is not a disease in itself. Blood pressure readings can be influenced by various factors, such as medical interventions, emotional states, physical activities, and environmental conditions. Previous research has mainly focused on high blood pressure as a risk factor and has often overlooked the fact that low blood pressure can also pose risks for CVD.^{48,49} Therefore, using blood pressure as a dependent variable to explore the role of UVR exposure in predicting CVD may not provide accurate insights.

The presence of biases in previous research may have led to null or positive correlations between UVR and CVD. For example, in a study involving 17,773 American stroke patients over the age of 45,⁵⁰ Kent et al. did not observe a correlation between daily ambient solar radiation and brachial blood pressure.⁵⁰ Similarly, Scragg et al. conducted a clinical trial with 119 individuals in New Zealand who had low vitamin D levels and found that ultraviolet exposure did not lead to a reduction in blood pressure.⁵¹ These studies may have been affected by biases such as limited latitude range exposure and biased blood pressure readings from subjects who were already unhealthy.⁵¹ Similar biases were also observed in a study involving male cyclists and triathletes in the United Kingdom.⁵²

A quartile correlation analysis showed a weak positive correlation between UVR and CVD mortality in a cohort study of individuals aged 51-70 in the United States.⁵³ However, this method only measured relationships between quartiles and not individual data points, which limits its accuracy. Additionally, using CVD mortality as a measure of UVR exposure's role as a risk factor may be flawed, as CVD is preventable and involves distinct phases before and after diagnosis.⁵⁴ This improper measure may bias the findings suggesting a detrimental role of UVR in the development of CVD. The inverse correlation between UVR and CVD incidence has been observed in various countries, which has led to an examination of UVR's effects on CVD in LMICs and high-income countries (1). This study shows that the correlations between UVR and CVD incidence are much stronger in LMICs compared to high-income countries. This difference may be influenced by economic factors and disparities in healthcare education between these regions.

4.1 | Strength and limitation of this study

Population-level data, including second-hand data, are utilized in this study despite potential random errors during collection and aggregation. This enhances the repeatability of our analysis by minimizing subjective biases often found in individual-based quantitative studies. However, several limitations should be acknowledged:

First, this study is quantitative and focuses on correlating variables, which does not imply causation even if associations are observed concurrently.

Second, as a population-level study, each population serves as the research unit, potentially leading to the intrinsic ecological fallacy in data analysis. Therefore, correlations identified at the country level may not necessarily be applicable at the individual level.

Third, the incidence of CVD is influenced by the availability of formal medical diagnoses, closely tied to economic prosperity and urbanization levels. The country-specific CVD incidence rates from the University of Washington data set may be incomplete, particularly in LMICs. Economic affluence (indexed by GDP PPP) and urban advantages (measured by urbanization rates) have been integrated into our analyses to mitigate potential confounding effects on CVD incidence accuracy, yet residual confounders may persist.

Fourth, the total population-level exposure to UVR is included as an independent variable to assess its statistical role in determining CVD incidence rates. However, real-life UVR exposure is influenced by various behavioral factors, such as sunglasses usage, sunblock application, outdoor recreational activities, clothing choices, and duration of outdoor exposure. These factors significantly modify UVR exposure levels, but their exact impact has not been quantified, posing a challenge in incorporating these potential confounders into the statistical models in this study. To address this limitation, future studies could consider innovative methods, such as using Satellite Earth Observation for near real-time monitoring of UV-A solar radiation,⁵⁵ which offers more precise measurements.^{55,56} These advancements could support further research into the relationship between personal solar exposure and its implications for cardiovascular health.

5 | CONCLUSION

Ambient UVR may play a significant and independent role in protecting against CVD progression worldwide. This protective effect appears to be more prominent in LMIC. However, since there is a lack of evidence to support the cardioprotective role of vitamin D supplementation, it is worth exploring whether ambient UVR can also protect against CVD independently of vitamin D supplementation.

AUTHOR CONTRIBUTIONS

Wenpeng You: Conceptualization; Investigation; Funding acquisition; Writing—original draft; Methodology; Validation; Visualization; Writing—review and editing; Software; Formal analysis; Project administration; Data curation; Resources.

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CONFLICT OF INTEREST STATEMENT

The sole author declares that there is no conflict of interest regarding the publication of this article. The author has read and approved the final version of the manuscript (corresponding author: WY) had full access to all of the data in this study and takes complete responsibility for the integrity of the data and the accuracy of the data analysis.

DATA AVAILABILITY STATEMENT

The data used for this study were obtained from publicly available repositories hosted by United Nations (UN) Agencies and the Institute for Health Metrics and Evaluation at the University of

Washington. Our use of this data complies with the terms and conditions set by the respective UN agencies, eliminating the need for formal permission to access and analyse the datasets. More information about the data sources can be found in the "Materials and Methods" section.

ETHICS STATEMENT

All variables used in this study were collected from reputable sources, such as international organizations like United Nations agencies and the Institute for Health Metrics and Evaluation (IHME). The data set comprises solely of non-identifiable, pre-existing data about human subjects. It is not possible to trace individual persons or their communities using this data. Therefore, the study poses minimal ethical risk and does not necessitate ethical approval or written informed consent from participants.

TRANSPARENCY STATEMENT

The lead author Wenpeng You affirms that this manuscript is an honest, accurate, and transparent account of the study being reported; that no important aspects of the study have been omitted; and that any discrepancies from the study as planned (and, if relevant, registered) have been explained.

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REFERENCES

- WHO (2021). "Cardiovascular diseases (CVDs)." Retrieved December 21 2022, from [https://www.who.int/en/news-room/fact-sheets/detail/cardiovascular-diseases-\(cvds\)](https://www.who.int/en/news-room/fact-sheets/detail/cardiovascular-diseases-(cvds))
- IHME. 2020. Global Burden of Disease Collaborative Network. Global Burden of Disease Study 2019 (GBD 2019) Results. <http://ghdx.healthdata.org/gbd-results-tool>
- Elagazi A, Kachur S, Carbone S, Lavie CJ, Blair SN. A review of obesity, physical activity, and cardiovascular disease. *Curr Obes Rep.* 2020;9:571-581.
- Yusuf S, Hawken S, Ôunpuu S, et al. Effect of potentially modifiable risk factors associated with myocardial infarction in 52 countries (the INTERHEART study): case-control study. *Lancet.* 2004;364(9438):937-952.
- Brook RD, Rajagopalan S, Pope CA, et al. Particulate matter air pollution and cardiovascular disease: an update to the scientific statement from the American Heart Association. *Circulation.* 2010;121(21):2331-2378.
- Stringhini S. Association of socioeconomic position with health behaviors and mortality. *JAMA.* 2010;303(12):1159-1166.
- Juzeniene A, Moan J. Beneficial effects of UV radiation other than via vitamin D production. *Dermat Endocrinol.* 2012;4(2):109-117.
- Weller RB. Sunlight has cardiovascular benefits independently of vitamin D. *Blood Purif.* 2016;41(1-3):130-134.
- Roelandts R. The history of phototherapy: something new under the sun? *J Am Acad Dermatol.* 2002;46(6):926-930.
- Al-Ghazal SK. *The valuable contributions of al-Rāzī (Rhazes) in the history of pharmacy.* Foundation for Science Technology and Civilization; 2007.
- Brennan P, Greenberg G, Miall W, et al. Seasonal variation in arterial blood pressure. *British Med J Clin Res.* 1982;285(6346):919.
- Rostand SG. Ultraviolet light may contribute to geographic and racial blood pressure differences. *Hypertension.* 1997;30(2):150-156.
- Marti-Soler H, Gubelmann C, Aeschbacher S, et al. Seasonality of cardiovascular risk factors: an analysis including over 230 000 participants in 15 countries. *Heart.* 2014;100(19):1517-1523.
- Judd SE, Tangpricha V. Vitamin D deficiency and risk for cardiovascular disease. *Am J Med Sci.* 2009;338(1):40-44.
- Wimalawansa SJ. Vitamin D deficiency: effects on oxidative stress, epigenetics, gene regulation, and aging. *Biology.* 2019;8(2):30.
- Scragg R, Rahman J, Thornley S. Association of sun and UV exposure with blood pressure and cardiovascular disease: a systematic review. *J Steroid Biochem Mol Biol.* 2019;187:68-75.
- The World Bank (2017). "World Bank Country and Lending Groups." Retrieved 10 May 2017, from <https://datahelpdesk.worldbank.org/knowledgebase/articles/906519-world-bank-country-and-lending-groups>
- WHO (2015). "Global Health Observatory, the data repository." WHO. Retrieved 11.26.2015, from <https://www.who.int/data/gho/data/indicators/indicator-details/GHO/uv-radiation>
- United Nations-Department of Economic and Social Affairs-Population Division (2021). File MORT/05-1: Life expectancy at exact age, e(x), for both sexes combined, by region, subregion and country, annually for 1950-2100 (Estimates, 1950 - 2021). Online edition The United Nations.
- Lakatta EG, Levy D. Arterial and cardiac aging: major shareholders in cardiovascular disease enterprises: part I: aging arteries: a "set up" for vascular disease. *Circulation.* 2003;107(1):139-146.
- Rodgers JL, Jones J, Bolleddu SI, et al. Cardiovascular risks associated with gender and aging. *J Cardiovasc Develop Disease.* 2019;6(2):19.
- The World Bank (2018). "Indicators[Data]." from <https://data.worldbank.org/indicator>
- Schultz WM, Kelli HM, Lisko JC, et al. Socioeconomic status and cardiovascular outcomes: challenges and interventions. *Circulation.* 2018;137(20):2166-2178.
- Gaziano TA, Bitton A, Anand S, Abrahams-Gessel S, Murphy A. Growing epidemic of coronary heart disease in low-and middle-income countries. *Curr Probl Cardiol.* 2010;35(2):72-115.
- You W, Henneberg M. Modern medical services, a double-edged sword manages symptoms, but accumulates genetic background of cardiovascular diseases: A cross population analysis of 217 countries. *Health Sci Rep.* 2024 Jan 22;7(1):e1828. doi:10.1002/hsr2.1828
- WHO Global Health Observatory (2022). "The data repository." WHO. Retrieved November 11 2022, from <http://www.who.int/gho/database/en/>
- Powell-Wiley TM, Poirier P, Burke LE, et al. Obesity and cardiovascular disease: a scientific statement from the American Heart Association. *Circulation.* 2021;143(21):e984-e1010.
- Schwartz MW, Seeley RJ, Zeltser LM, et al. Obesity pathogenesis: an endocrine society scientific statement. *Endocr Rev.* 2017;38(4):267-296.
- Unamuno X, Gómez-Ambrosi J, Rodríguez A, Becerril S, Frühbeck G, Catalán V. Adipokine dysregulation and adipose tissue inflammation in human obesity. *Eur J Clin Invest.* 2018;48(9):e12997.
- Yusuf S, Reddy S, Ôunpuu S, Anand S. Global burden of cardiovascular diseases: part II: variations in cardiovascular disease by specific ethnic groups and geographic regions and prevention strategies. *Circulation.* 2001;104(23):2855-2864.
- Zittermann A, Schleithoff SS, Koerfer R. Putting cardiovascular disease and vitamin D insufficiency into perspective. *Br J Nutr.* 2005;94(4):483-492.
- Mahajan H, Mallinson PAC, Lieber J, et al. The Association of Total Meat Intake with Cardio-Metabolic Disease Risk Factors and Measures of Sub-Clinical Atherosclerosis in an Urbanising Community of Southern India: A Cross-Sectional Analysis for the APCAPS Cohort. *Nutrients.* 2024 Mar 5;16(5):746. doi:10.3390/nu16050746

33. You W, Henneberg M. Cereal crops are not created equal: wheat consumption associated with obesity prevalence globally and regionally. *AIMS Pub Health*. 2016;3(2):313-328.
34. You W, Henneberg M. Prostate cancer incidence is correlated to total meat intake—a cross-national ecologic analysis of 172 countries. *Asian Pacific J Cancer Preve*. 2018;19(8):2229-2239.
35. You W, Henneberg R, Henneberg M. Advanced healthcare services leading to relax natural selection may have been contributing to global and regional increase of dementia incidence. *Res Square*. 2021:1091190.
36. You W, Henneberg R, Henneberg M. Healthcare services relaxing natural selection may contribute to increase of dementia incidence. *Sci Rep*. 2022;12(1):8873.
37. You W, Henneberg R, Saniotis A, Ge Y, Henneberg M. Total meat intake is associated with life expectancy: a cross-sectional data analysis of 175 contemporary populations. *Int J Gen Med*. 2022;15:1833-1851.
38. IHME. Global Burden of Disease Collaborative Network. Global Burden of Disease Study 2019 (GBD 2019) Results. 2020. <http://ghdx.healthdata.org/gbd-results-tool>
39. Feelisch M, Kolb-Bachofen V, Liu D, et al. Is sunlight good for our heart? *Eur Heart J*. 2010;31(9):1041-1045.
40. Weller RB. The health benefits of UV radiation exposure through vitamin D production or non-vitamin D pathways. Blood pressure and cardiovascular disease. *Photochem Photobiol Sci*. 2017;16:374-380.
41. Krause R, Dobberke J, Buehring M, et al. Biologic Effects of Light. *The role of ultraviolet radiation on cardiocirculatory regulation and on cardiovascular risk*. Springer; 2002:219-229.
42. Zittermann A. Vitamin D and disease prevention with special reference to cardiovascular disease. *Prog Biophys Mol Biol*. 2006;92(1):39-48.
43. You W, Henneberg R, Coventry BJ, Henneberg M. Cutaneous malignant melanoma incidence is strongly associated with European depigmented skin type regardless of ambient ultraviolet radiation levels: evidence from Worldwide population-based data. *AIMS Public Health*. 2022;9(2):378-402.
44. de la Guía-Galipienso F, Martínez-Ferran M, Vallecillo N, Lavie CJ, Sanchis-Gomar F, Pareja-Galeano H. Vitamin D and cardiovascular health. *Clin Nutr*. 2021;40(5):2946-2957.
45. Holick MF. Sunlight and vitamin D for bone health and prevention of autoimmune diseases, cancers, and cardiovascular disease. *Am J Clin Nutr*. 2004;80(6):1678S-1688S.
46. Wolf ST, Kenney WL. Skin pigmentation and vitamin D-folate interactions in vascular function: an update. *Curr Opin Clin Nutr Metab Care*. 2021;24(6):528-535.
47. Scragg R. Seasonality of cardiovascular disease mortality and the possible protective effect of ultra-violet radiation. *Int J Epidemiol*. 1981;10(4):337-341.
48. Gu D, Kelly TN, Wu X, et al. Blood pressure and risk of cardiovascular disease in Chinese men and women. *Am J Hypertens*. 2008;21(3):265-272.
49. Stokes J, Kannel WB, Wolf PA, et al. Blood pressure as a risk factor for cardiovascular disease. *Hypertension*. 1989;13(suppl 1):113-118.
50. Kent ST, Cushman M, Howard G, et al. Sunlight exposure and cardiovascular risk factors in the REGARDS study: a cross-sectional split-sample analysis. *BMC Neurol*. 2014;14:133.
51. Scragg R, Wishart J, Stewart A, et al. No effect of ultraviolet radiation on blood pressure and other cardiovascular risk factors. *J Hypertens*. 2011;29(9):1749-1756.
52. Muggerridge DJ, Sculthorpe N, Grace FM, et al. Acute whole body UVA irradiation combined with nitrate ingestion enhances time trial performance in trained cyclists. *Nitric oxide*. 2015;48:3-9.
53. Lin S-W, Wheeler DC, Park Y, et al. Prospective study of ultraviolet radiation exposure and mortality risk in the United States. *Am J Epidemiol*. 2013;178(4):521-533.
54. Shojania KG, Forster AJ. Hospital mortality: when failure is not a good measure of success. *Can Med Assoc J*. 2008;179(2):153-157.
55. Morelli M, Michelozzi B, Simeone E, Khazova M. Validation of a satellite-based solar UV-A radiation dosimeter for mobile healthcare applications. *J Atmos Solar-Terrestrial Physics*. 2021; 215:105529.
56. Parisi A, Igoe D, Downs N, Turner J, Amar A, A Jebar M. Satellite monitoring of environmental solar ultraviolet A (UVA) exposure and irradiance: a review of OMI and GOME-2. *Remote Sens*. 2021;13(4): 752.

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