

## Recollection

# Impact of Comprehensive Strategy on Mortality in Heterosexually Transmitted HIV-Infected Individuals — Liangshan Prefecture, Sichuan Province, China, 2008–2024

Ruiying Liang<sup>1,2</sup>; Fangfang Chen<sup>3,†</sup>; Yan Zhao<sup>1</sup>; Ju Wang<sup>4</sup>; Decai Zhao<sup>1</sup>; Jiejun Yu<sup>1</sup>; Houlin Tang<sup>1</sup>; Jian Li<sup>1</sup>; Zhongfu Liu<sup>1,†</sup>

## ABSTRACT

A comprehensive strategy integrating universal human immunodeficiency virus (HIV) testing, real-time patient tracking, and enhanced antiretroviral therapy (ART) within a poverty-alleviation framework was implemented to curb the HIV epidemic in Liangshan Prefecture. This study aimed to quantify its impact on HIV treatment coverage and mortality among heterosexually transmitted HIV-infected individuals. Data on heterosexually transmitted HIV infections in Liangshan from 2008 to 2024 were extracted from China's HIV/acquired immunodeficiency syndrome (AIDS) Comprehensive Response Information Management System (CRIMS) database. Cox proportional hazards models were used to estimate the effectiveness of the comprehensive strategy on mortality. Interrupted time series analysis examined trends and changes in treatment coverage and mortality before and after implementation. Subgroup analysis investigated outcomes variations across different populations. Among 37,034 eligible HIV-infected individuals, 7,302 deaths occurred. Not receiving ART was the strongest risk factor for mortality [hazard ratio (HR): 17.74]. The comprehensive strategy led to a 21.62% immediate increase and a 1.48% annual increase in treatment coverage, accompanied by a 0.47% decline in mortality per year [95% confidence interval (CI): -0.73, -0.21]. By 2024, the mortality rate was 4.89% lower than the expected level (95% CI: -8.02, -1.75), representing an 81.2% reduction. Reductions were greatest among males, residents of key counties, and patients at the HIV infection stage. The comprehensive strategy in Liangshan significantly improved treatment coverage and reduced mortality among heterosexually infected individuals. Further efforts are needed to promote early diagnosis, rapid ART initiation, and tailored care support for advanced disease. This comprehensive

model offers an adaptable template for high-burden, resource-limited settings.

Human immunodeficiency virus (HIV) remains a major global public health threat. The burden is disproportionately concentrated in settings with constrained health and economic resources, where access to timely diagnosis and sustained antiretroviral therapy (ART) is limited. Liangshan Yi Autonomous Prefecture, situated in southwest China, has been a focal area for national HIV prevention efforts. Following successful harm-reduction programs that curtailed injection drug use transmission, heterosexual contact emerged as the dominant transmission route from 2014 onward, accounting for 59.0% of newly diagnosed cases in 2017 and 79.4% in 2019 (1). Although national initiatives have slowed the overall epidemic trajectory (2–3), gaps remain along the HIV care cascade. Late presentation, delayed ART initiation, and poor retention in care leave a substantial proportion of HIV-positive individuals without durable viral suppression. Consequently, Liangshan's treatment coverage and virological suppression continue to lag behind national averages, undermining the UNAIDS 95-95-95 targets and adversely affecting survival and secondary transmission (4–5).

To address these challenges, the “Action Plan for HIV Prevention and Health Poverty Alleviation in Liangshan Prefecture” was initiated in September 2017 (Phase I, 2017–2020), subsequently expanded into Phase II and hereafter referred to as the “Action Plan”. The Action Plan introduced a comprehensive strategy, aiming to expand testing and treatment coverage, reduce mortality, and improve healthcare accessibility (3,6). Phase I targeted four “key counties” (Butuo, Zhaojue, Yuexi, Meigu) with high HIV prevalence, and Phase II extended these successful measures to all

17 counties in Liangshan. The core measures of the comprehensive strategy included universal HIV screening integrated into routine health checkups, real-time monitoring of the ART treatment cascade, “Three Lines and One Network Base” technical support system and “1+M+N” community-based management model, which were specifically designed to enhance multi-sector collaboration and deliver contextually adapted HIV/AIDS services.

While early reports documented substantial increases in HIV testing and treatment coverage, the long-term impact of the comprehensive strategy on treatment outcomes, particularly mortality, has yet to be formally evaluated. Therefore, this study conducted a population-based assessment of heterosexually transmitted HIV-infected individuals diagnosed during 2008–2024, employing interrupted time series (ITS) analysis and COX proportional hazards model to quantify changes in treatment coverage and mortality and to explore disparities across subgroup populations, to optimize HIV control strategies in Liangshan and other high-burden settings.

## METHODS

### Data Source and Comprehensive Strategy

This study utilized routine case reporting data retrieved from China’s HIV/AIDS Comprehensive Response Information Management System (CRIMS). The study population comprised adults aged 15 years and older who were first diagnosed with heterosexually transmitted HIV in Liangshan between 2008 and 2024. Data were de-identified prior to analysis.

The comprehensive strategy combined several key measures: 1) universal HIV screening integrated into routine health checkups, achieving up to 98% coverage among residents in key counties; 2) real-time monitoring of the ART cascade with enhanced measures such as timely patient referrals and differentiated service delivery; 3) governance through the “Three Lines and One Network Base” technical support system, which integrates prefecture- and county-level CDC, hospitals, and maternal-child-health centers (“Three Lines”) with a robust grassroots community network (“One Network Base”) to ensure comprehensive care; and 4) the “1+M+N” community-based management model, which clearly defines the responsibilities of township government (“1”), township healthcare institutions (“M”), and village-level health workers (“N”) to maintain continuity and

effectiveness of HIV care. Phase I focused resources on four HIV high-prevalence counties, while Phase II expanded to all 17 counties (3,6).

### Outcomes and Survival Analysis

The primary outcomes for assessing the comprehensive strategy’s effectiveness were treatment coverage and mortality. Follow-up time, as recorded in months, started on the date of HIV diagnosis to either the date of death, date of last follow-up visit, or December 31, 2024, whichever came first. Cox proportional hazards regression was used to identify potential risk factors for mortality, with hazard ratios (HRs) quantifying the impact of the comprehensive strategy on mortality after controlling for demographic factors (7). A detailed methodological explanation is provided in the [Supplementary Material](https://weekly.chinacdc.cn/) (available at <https://weekly.chinacdc.cn/>).

### Interrupted Time Series (ITS) Analysis

Given that randomization was infeasible or unethical in this context, ITS analysis was conducted to evaluate the comprehensive strategy’s impact on treatment coverage and mortality at the population level. Segmented linear regression models quantified both the immediate and long-term effects of the intervention, with 2018 designated as the intervention (comprehensive strategy) start point in this study. The model is as follows:

$$Y_t = \beta_0 + \beta_1 T + \beta_2 X_t + \beta_3 (T - T_0) X_t + \varepsilon$$

Where:  $Y_t$  denotes the outcome variable at time  $t$ ,  $T$  is the continuous time variable,  $\beta_0$  is the pre-intervention initial level,  $\beta_1$  represents the slope before the intervention, reflecting the annual trend of change before the intervention,  $\beta_2$  is the immediate change in the level of the outcome indicator before and after the intervention,  $\beta_3$  represents the change in the slope before and after the intervention,  $(\beta_1 + \beta_3)$  is the slope after the intervention, reflecting the annual average trend of change after the comprehensive strategy’s implementation,  $\varepsilon$  represents the random error (8). Additional detailed methodological explanations are provided in [Supplementary Material](#).

The Newey-West method (9) was employed to control for autocorrelation and heteroscedasticity, providing 95% confidence intervals (CIs) with optimal lag selection through an automatic bandwidth procedure. Subgroup analyses were conducted to examine disparities by gender (male *vs.* female), region

(key counties *vs.* non-key counties), disease stage (HIV *vs.* AIDS), and ART status (yes *vs.* no). The regional stratification of Liangshan was based on an HIV prevalence threshold of 1%. Four counties demonstrating HIV prevalence above 1% were classified as key counties (Butuo, Zhaojue, Yuexi and Meigu), while the remaining 13 counties were designated as non-key counties.

### Software

All statistical tests were two-sided, with a significance threshold set at  $P < 0.05$ . Data analyses were performed using SPSS (version 29.0; IBM Corp., Armonk, NY, USA) and R software (version 4.2.3; R Foundation for Statistical Computing, Vienna, Austria).

## RESULTS

Between 2008 and 2024, 37,034 individuals with heterosexually transmitted HIV were reported in Liangshan, with 16,326 cases reported before the comprehensive strategy implementation and 20,708 afterward. The number of reported cases peaked at 6,622 in the first year following implementation, then declined annually by 22% (Supplementary Figure S1, available at <https://weekly.chinacdc.cn/>). Most cases involved females, individuals aged 30–44 years, and residents of key counties. Compared to the pre-implementation period, the post-implementation period showed a slight increase in the proportion of cases in the 30–44 year age group, while decreases were observed among females and residents of key counties. Concurrently, significant increases occurred in the proportion of cases among individuals aged over 45 years and those with a baseline CD4 count  $< 200$  (Supplementary Table S1, available at <https://weekly.chinacdc.cn/>).

During the study period, 7,302 deaths were recorded (19.7% of total cases), including 1,084 AIDS-related deaths (2.9% of total cases). Cox regression analysis identified not receiving ART as the most significant risk factor for mortality (adjusted *HR*: 17.74, Table 1). After controlling for demographics and excluding ART status as a modifiable factor, the comprehensive strategy was independently associated with a 35% reduction in overall mortality (adjusted *HR*: 0.65, 95% *CI*: 0.64, 0.69) and a 60% reduction in AIDS-related mortality (adjusted *HR*: 0.40, 95% *CI*: 0.34, 0.46).

### Treatment Coverage and Mortality

From 2008 to 2024, treatment coverage demonstrated a significant upward trend ( $P < 0.05$ ), with a compound annual growth rate (CAGR) of 34.79%. The median annual coverage increased from 38.46% (IQR: 13.03–56.16) with a CAGR of 62.16% pre-implementation to 97.61% (IQR: 95.17–99.39) with a CAGR of 1.88% post-implementation. In key counties, ART coverage improved from 54.74% (IQR: 48.66–59.54) to 95.63% (IQR: 92.52–97.91), while in non-key counties it increased from 57.36% (IQR: 50.02–63.22) to 96.40% (IQR: 90.17–97.99) (Figure 1). Segmented linear regression revealed an immediate 21.62% increase (95% *CI*: 16.63, 26.62) in treatment coverage during the first implementation year, followed by a steady annual increase of 1.48%. By 2024, treatment coverage reached 99.68%, slightly exceeding the expected level (8.95%, 95% *CI*: –4.12, 22.02).

Concurrently, overall mortality exhibited a significant downward trend ( $P < 0.05$ ), decreasing from a median of 4.83% (IQR: 4.15–5.37) before implementation to 2.06% (IQR: 1.90–2.95) afterward. Key counties demonstrated a mortality reduction from 4.64% (IQR: 4.52–5.68) pre-implementation to 2.73% (IQR: 1.80–5.42) post-implementation, while non-key counties showed a decrease from 4.51% (IQR: 4.47–4.61) to 3.20% (IQR: 2.42–4.60) (Figure 1). AIDS-related mortality similarly declined significantly ( $P < 0.05$ ) from 1.09% (IQR: 0.13–1.27) pre-implementation to 0.24% (IQR: 0.09–0.50) post-implementation (Figure 2). Although no immediate change was detected in 2018, segmented linear regression showed that mortality decreased by 0.47% annually following the intervention (95% *CI*: –0.73, –0.21). By 2024, mortality was 4.89% lower than expected (95% *CI*: –8.02, –1.75), corresponding to an 81.2% reduction (Table 2 and Figure 2).

### Subgroup Analysis

Subgroup analysis revealed disparities in treatment coverage improvements and mortality reductions across gender, region, and disease stage. Greater improvements in both outcomes occurred among males, patients in key counties, and those diagnosed at the HIV infection stage. Among AIDS patients, the immediate increase in treatment coverage was small and insignificant (3.61, 95% *CI*: –5.26, 12.47), followed by a modest annual increase of 0.87%. The mortality of AIDS patients showed an insignificant

TABLE 1. Factors associated with mortality among heterosexually transmitted HIV-infected individuals in Liangshan, 2008 to 2024 (n=37,034).

Variables	Total (n)	Death (n)	Follow-up time (person-months)	Univariate analysis		Multivariate analysis	
				HR (95% CI)	P	HR (95% CI)	P
Gender							
Female	18,926	2,994	168.7	1		1	
Male	18,108	4,308	146.9	1.69 (1.62, 1.78)	<0.001	1.46 (1.38, 1.53)	<0.001
Age (years)							
15–29	12,710	1,860	172.2	1		1	
30–44	15,498	3,027	158.6	1.55 (1.46, 1.64)	<0.001	1.59 (1.49, 1.69)	<0.001
45–59	6,093	1,205	148.2	1.92 (1.78, 2.06)	<0.001	2.27 (2.09, 2.46)	<0.001
≥60	2,733	1,210	94.3	4.65 (4.32, 5.00)	<0.001	3.87 (3.53, 4.24)	<0.001
Occupation							
Farmers	31,803	6,130	159.3	0.87 (0.82, 0.93)	<0.001	1.19 (1.12, 1.28)	<0.001
Others	5,231	1,172	153.5	1		1	
Education							
Illiterate	17,614	3,689	157.5	1.24 (1.16, 1.33)	<0.001	1.16 (1.07, 1.25)	<0.001
Primary School	13,207	2,605	156.5	1.23 (1.15, 1.33)	<0.001	1.17 (1.08, 1.26)	<0.001
Junior High and Above	6,213	1,008	165.6	1		1	
Disease stage							
HIV	30,745	5,652	160.8	1		1	
AIDS	6,289	1,650	147.0	1.41 (1.34, 1.49)	<0.001	1.40 (1.30, 1.51)	<0.001
Baseline CD4 count (cell/μL)*							
<200	5,802	1,409	146.9	1		1	
200–349	10,336	1,488	167.0	0.53 (0.49, 0.57)	<0.001	0.71 (0.65, 0.77)	<0.001
350–499	9,050	1,290	170.1	0.48 (0.44, 0.51)	<0.001	0.65 (0.59, 0.71)	<0.001
≥500	9,855	1,300	173.5	0.43 (0.40, 0.46)	<0.001	0.58 (0.53, 0.63)	<0.001
Missing	1,991	1,815	13.4	16.36 (15.22, 17.59)	<0.001	2.28 (2.06, 2.52)	<0.001
Current ART status							
No	3,263	3,196	22.4	25.12 (23.91, 26.40)	<0.001	17.74 (16.54, 19.02)	<0.001
Yes	33,771	4,106	173.8	1		1	
Study phase							
Pre-Action Plan	16,326	5,062	150.5	1		1	
Post-Action Plan	20,708	2,240	76.0	0.54 (0.51, 0.57)	<0.001	1.13 (1.06, 1.19)	<0.001

\* The initial CD4 count at diagnosis. Action Plan refers to the comprehensive strategy implemented in 2018.

Abbreviation: ART=antiretroviral therapy; CI=confidence interval; HR=hazard ratios; HIV=human immunodeficiency virus; AIDS=acquired immunodeficiency syndrome.

immediate change (0.04%, 95% CI: –2.01, 2.09), but declined significantly thereafter. By 2024, mortality among those receiving ART was 2.69% (95% CI: –4.71, –0.67) lower than expected, whereas mortality among patients not receiving ART rose immediately after implementation (9.50, 95% CI: 0.65, 18.35) and remained elevated compared to expected levels (Table 2 and Figure 2).

## DISCUSSION

This study provides compelling evidence that the comprehensive strategy substantially accelerated progress towards universal ART coverage and produced a marked, sustained reduction in mortality among heterosexually transmitted HIV-infected individuals in Liangshan. These findings align with studies from

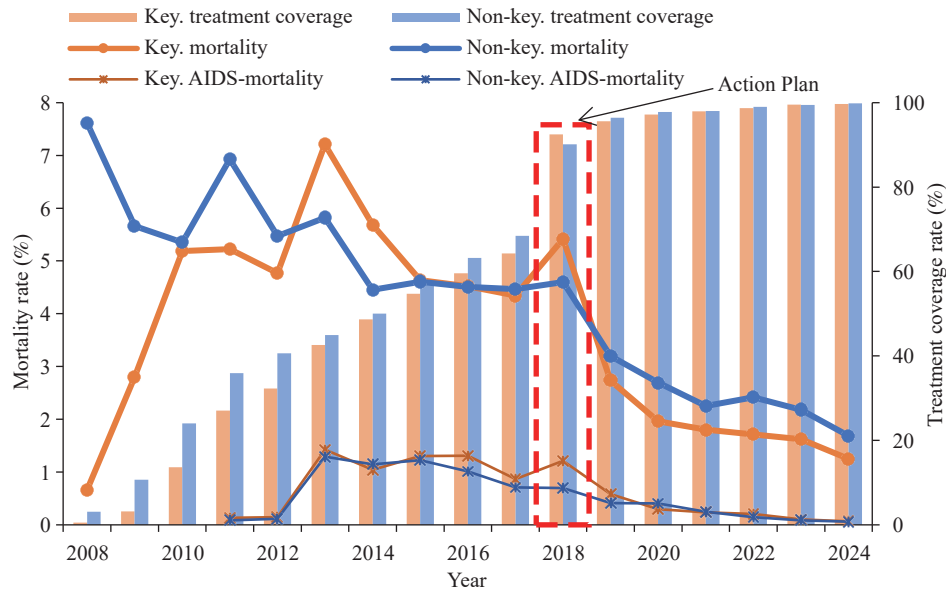


FIGURE 1. Trends in treatment coverage and mortality among heterosexually transmitted HIV-infected individuals in Liangshan Prefecture, 2008–2024. Action Plan refers to the comprehensive strategy implemented in 2018. Abbreviation: AIDS-mortality=acquired immunodeficiency syndrome-related mortality; Key.=Key counties (Butuo, Zhaojue, Yuxi, and Meigu); Non-key.=Non-key counties (all counties in Liangshan except for key counties).

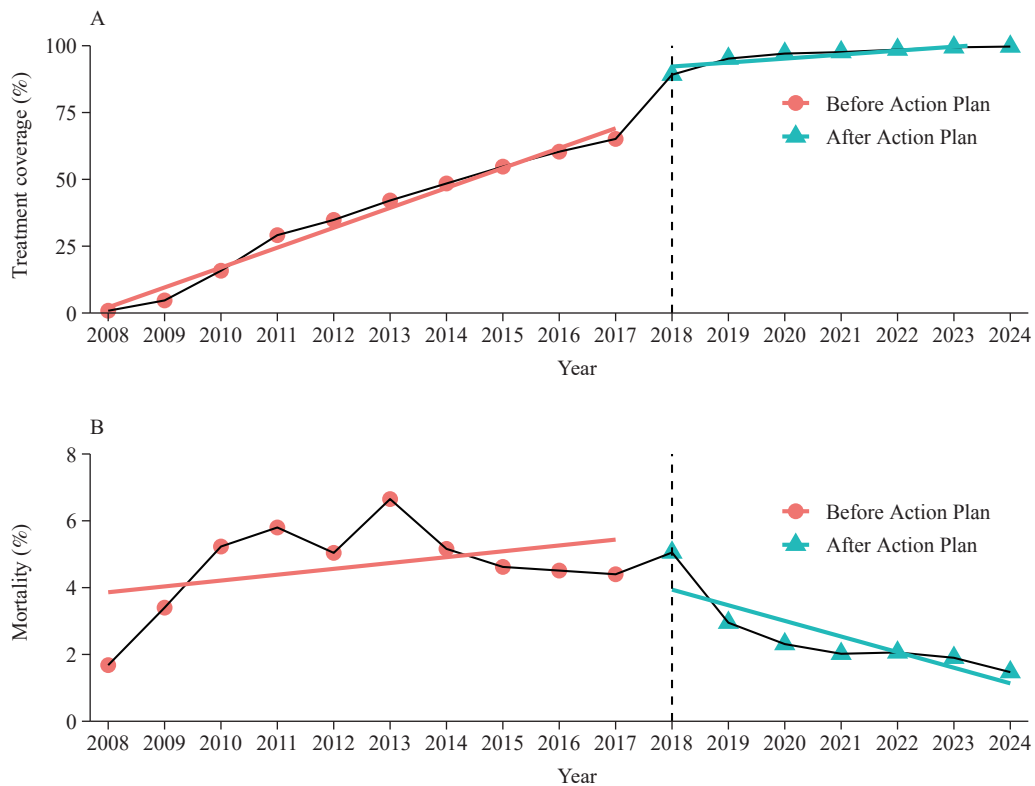


FIGURE 2. Segmented linear regression models of outcome indicators for heterosexual HIV-infected individuals in Liangshan Prefecture, 2008–2024. The solid pink and blue lines represent the model-fitted lines. Action Plan refers to the comprehensive strategy implemented in 2018. (A) Treatment coverage; (B) Mortality.

Note: Treatment coverage means treatment coverage rate per year; mortality calculated by all-cause mortality rate per year.

TABLE 2. Segmented linear regression models of the impact of comprehensive strategy on treatment coverage and mortality among heterosexually transmitted HIV-infected individuals in Liangshan from 2008 to 2024.

Variables	Level change at Action Plan	Level change in 2024	Trend before Action Plan	Trend after Action Plan
Treatment coverage rate				
Total	21.62 (16.63, 26.62)	8.95 (−4.12, 22.02)	7.44 (6.80, 8.09)	1.48 (0.73, 2.24)
Gender				
Male	21.97 (17.06, 26.88)	10.17 (−1.87, 22.21)	7.26 (6.62, 7.90)	1.72 (0.96, 2.49)
Female	21.39 (16.28, 26.51)	8.01 (−5.90, 21.91)	7.59 (6.93, 8.25)	1.29 (0.54, 2.04)
Region				
Non-key counties	20.31 (15.29, 25.33)	8.01 (−4.81, 20.83)	7.09 (6.31, 7.87)	1.30 (0.53, 2.07)
Key counties	24.18 (20.27, 28.09)	10.50 (−4.16, 25.15)	7.57 (6.82, 8.33)	1.11 (0.75, 1.46)
Disease stage				
HIV	32.50 (26.85, 38.14)	22.22 (12.28, 32.16)	6.73 (5.97, 7.49)	1.94 (1.23, 2.66)
AIDS	3.61 (−5.26, 12.47)	−5.31 (−19.77, 9.15)	5.06 (2.98, 7.13)	0.87 (0.42, 1.32)
Mortality rate				
Total	−1.03 (−2.85, 0.79)	−4.89 (−8.02, −1.75)	0.18 (−0.11, 0.46)	−0.47 (−0.73, −0.21)
Gender				
Male	−1.08 (−2.78, 0.63)	−5.22 (−7.84, −2.61)	0.18 (−0.09, 0.45)	−0.51 (−0.71, −0.32)
Female	−0.99 (−3.08, 1.10)	−4.58 (−8.05, −1.11)	0.16 (−0.16, 0.47)	−0.44 (−0.74, −0.13)
Region				
Non-key counties	−0.62 (−1.84, 0.60)	−2.88 (−5.23, −0.52)	−0.02 (−0.29, 0.26)	−0.39 (−0.53, −0.26)
Key counties	−1.27 (−4.04, 1.50)	−6.16 (−9.67, −2.66)	0.28 (−0.11, 0.67)	−0.54 (−0.85, −0.22)
Disease stage				
HIV	−1.02 (−3.12, 1.08)	−3.83 (−6.32, −1.35)	0.04 (−0.24, 0.33)	−0.42 (−0.69, −0.16)
AIDS	0.04 (−2.01, 2.09)	−2.31 (−6.47, 1.85)	−0.13 (−0.61, 0.35)	−0.52 (−0.82, −0.23)
ART				
No	9.50 (0.65, 18.35)	12.31 (6.59, 18.02)	0.51 (0.22, 0.79)	0.98 (−0.99, 2.94)
Yes	−0.90 (−1.86, 0.07)	−2.69 (−4.71, −0.67)	0.14 (−0.15, 0.43)	−0.16 (−0.22, −0.10)

Note: Data are rate (95% CI) or trend (95% CI). Key counties: Butuo, Zhaojue, Yuexi, and Meigu. Non-key counties: all counties in Liangshan except for key counties. Action Plan: the comprehensive strategy was implemented in 2018.

Abbreviation: ART=antiretroviral therapy; CI=confidence interval; HIV=human immunodeficiency virus; AIDS=acquired immunodeficiency syndrome.

other high HIV prevalence settings (*10–11*) and underscore the value of integrating routine HIV services with locally tailored governance, real-time cascade tracking, and poverty-reduction initiatives in high-burden, resource-constrained settings.

A notable achievement is the remarkable increase in treatment coverage following implementation of the comprehensive strategy, approaching universal coverage. This achievement exceeds national and international benchmarks for the treat-all policy (*12–14*). The near-universal uptake resulted from several mutually reinforcing measures implemented under Liangshan's comprehensive strategy, including universal HIV testing integrated into health checkups, real-time cascade tracking of each patient, and two

complementary service governance models — “Three Lines and One Network Base” and “1+M+N” — that mobilize multisectoral accountability down to the grassroots community level. These measures shortened the interval from diagnosis to treatment, enhanced ART retention, and facilitated differentiated care, thereby maximizing the population-level preventive and therapeutic benefits of ART.

Our findings further confirmed the role of ART scale-up, demonstrating that Liangshan's comprehensive strategy primarily reduced mortality risk through ART expansion, with this effect being particularly pronounced in reducing the risk of AIDS-related deaths. The magnitude of mortality decline observed in Liangshan exceeds that reported from



universal test and treat programs in other settings, where reductions have ranged from 10% to 50% (5,11–12). This might be attributed to Liangshan's rapid escalation in ART coverage from 65% in 2017 to 95% within three years.

Subgroup analysis revealed larger improvements in treatment coverage and mortality among males, residents of key counties, and individuals at HIV infection stage. For males, who typically present later, exhibit poorer ART adherence, and experience higher mortality than females (4,13), the tailored peer-navigation, workplace outreach, and culturally adapted counseling embedded in the comprehensive strategy may have reduced gender-related barriers to care. The greater improvements in both outcomes and a lower proportion of cases in key counties underscore the amplifying effect of geographic prioritization and concentrated resource inputs. The transient mortality increase observed immediately after implementation, particularly among patients at AIDS stage and not receiving ART, was attributable to the surge in case detection that accompanied expanded screening in 2018 (3). Many newly diagnosed patients had advanced disease stage and therefore faced high short-term mortality despite rapid linkage to ART (15). Sustained decreases in both all-cause mortality and AIDS-related mortality over subsequent years suggest that early treatment initiation and improved care management rapidly offset this initial increase.

Notably, expanded screening also increased case detection among older patients, a demographic shift partially attributable to prolonged HIV transmission dynamics and delayed diagnosis, but also suggesting unmet prevention needs in aging populations. For AIDS patients, our findings reinforce the necessity of integrated management protocols encompassing uninterrupted ART delivery, psychosocial support, and multidisciplinary chronic care integration to mitigate mortality risks. The program's success extends prior evidence by demonstrating that scalable HIV response models can yield equivalent benefits in an ethnically diverse region with historically limited health infrastructure. Coupling HIV prevention with poverty alleviation initiatives likely enhanced health gains by addressing structural determinants that impede sustained engagement in care. To consolidate gains, Liangshan should prioritize earlier diagnosis through intensified outreach in non-key counties, expand immediate ART initiation, and develop age- and gender-specific adherence interventions. Implementation science studies should delineate which

components of the comprehensive strategy are most cost-effective and transferable to other regions.

This study has several limitations. First, concurrent national public health initiatives may have contributed to the observed trends, although our ITS modeling mitigated this risk by accounting for pre-intervention trajectories. Second, potential underreporting and misclassification of deaths, particularly before the implementation of the comprehensive strategy, could bias the actual impact of the strategy on mortality (3). Last, our findings may not generalize to settings without comparable political commitment or community governance structures.

In conclusion, our results demonstrate that the comprehensive strategy in Liangshan, as a data-driven, multisectoral approach, can achieve near-universal ART coverage and an unprecedented reduction in mortality among heterosexually transmitted HIV-infected individuals. Scaling and adapting this comprehensive model could accelerate progress towards the 95-95-95 targets and the broader goals of the Sustainable Development Agenda in similar high-burden regions.

**Conflicts of interest:** No conflicts of interest.

**Ethical statement:** Approved by the Institutional Review Board of the National Center for AIDS/STD Control and Prevention, Chinese CDC (X140121318), with informed consent waived due to the retrospective nature of aggregated data.

**Funding:** Supported by National Natural Science Foundation of China (NSFC) Project (71774150).

doi: [10.46234/ccdcw2025.138](https://doi.org/10.46234/ccdcw2025.138)

# Corresponding authors: Fangfang Chen, [chenfangfang@chinaaids.cn](mailto:chenfangfang@chinaaids.cn); Zhongfu Liu, [liuzhongfu@chinaaids.cn](mailto:liuzhongfu@chinaaids.cn).

<sup>1</sup> National Center for AIDS/STD Control and Prevention, Chinese Center for Disease Control and Prevention, Beijing, China; <sup>2</sup> Beijing Chaoyang District Center for Disease Control and Prevention, Beijing, China; <sup>3</sup> National Key Laboratory of Intelligent Tracking and Forecasting for Infectious Diseases, National Center for AIDS/STD Control and Prevention, Chinese Center for Disease Control and Prevention, Beijing, China; <sup>4</sup> Liangshan Prefecture Center for Disease Control and Prevention, Xichang City, Sichuan Province, China.

Copyright © 2025 by Chinese Center for Disease Control and Prevention. All content is distributed under a Creative Commons Attribution Non Commercial License 4.0 (CC BY-NC).

Submitted: April 21, 2025

Accepted: June 05, 2025

Issued: June 13, 2025

## REFERENCES

1. Yu G, Li TL, Wang ZH, Nengge AJ, Shama BD, Jike CN, et al.

- Analysis of the characteristics of HIV/AIDS patients reported in Liangshan Prefecture between 2017 and 2019. *Chin J AIDS STD* 2021;27(10):1152-3. <https://link.cnki.net/doi/10.13419/j.cnki.aids.2021.10.24>. (In Chinese).
2. Deng L, Liu ZF, Zhang SZ, Dou ZH, Wang QX, Ma Y, et al. Survival time and related influencing factors of AIDS patients in Liangshan prefecture, Sichuan province, during 2008-2013. *Chin J Epidemiol* 2015;36(6):569 – 75. <https://doi.org/10.3760/cma.j.issn.0254-6450.2015.06.007>.
  3. Liu ZF, Tang XF, Liu YF, Zhang LL, Yang YH, Zheng YL, et al. HIV prevention and health poverty alleviation - Liangshan prefecture, Sichuan Province, China, 2017-2020. *China CDC Wkly* 2021;3(48): 1031 – 5. <https://doi.org/10.46234/ccdcw2021.250>.
  4. Xu JL, Jike C, Ma Y, Yu G, Wang J, Wang K, et al. Effects and associated factors of HIV/AIDS anti-retroviral therapy in Liangshan Yi Autonomous Prefecture, Sichuan Province. *Chin J Prev Med* 2018;52(6):668 – 72. <https://doi.org/10.3760/cma.j.issn.0253-9624.2018.06.018>.
  5. Wu ZY, Zhao Y, Ge XM, Mao YR, Tang ZZ, Shi CX, et al. Simplified HIV Testing and Treatment in China: analysis of mortality rates before and after a structural intervention. *PLoS Med* 2015;12(9):e1001874. <https://doi.org/10.1371/journal.pmed.1001874>.
  6. Liu YF, Tang XF, Han MJ, Jike CN, Yang YH, Qu MX, et al. Achievements and prospects of AIDS Prevention and Health Poverty Alleviation in Liangshan Prefecture. *Chin J AIDS STD* 2022;28(12): 1339-44. <https://link.cnki.net/doi/10.13419/j.cnki.aids.2022.12.01>. (In Chinese).
  7. Zhang FJ, Dou ZH, Ma Y, Zhang Y, Zhao Y, Zhao DC, et al. Effect of earlier initiation of antiretroviral treatment and increased treatment coverage on HIV-related mortality in China: a national observational cohort study. *Lancet Infect Dis* 2011;11(7):516 – 24. [https://doi.org/10.1016/S1473-3099\(11\)70097-4](https://doi.org/10.1016/S1473-3099(11)70097-4).
  8. Bernal JL, Cummins S, Gasparrini A. Interrupted time series regression for the evaluation of public health interventions: a tutorial. *Int J Epidemiol* 2017;46(1):348 – 55. <https://doi.org/10.1093/ije/dyw098>.
  9. Dorward J, Khubone T, Gate K, Ngobese H, Sookrajh Y, Mkhize S, et al. The impact of the COVID-19 lockdown on HIV care in 65 South African primary care clinics: an interrupted time series analysis. *Lancet HIV* 2021;8(3):e158 – 65. [https://doi.org/10.1016/S2352-3018\(20\)30359-3](https://doi.org/10.1016/S2352-3018(20)30359-3).
  10. Cao W, Hsieh E, Li TS. Optimizing treatment for adults with HIV/AIDS in China: successes over two decades and remaining challenges. *Curr HIV/AIDS Rep* 2020;17(1):26 – 34. <https://doi.org/10.1007/s11904-019-00478-x>.
  11. Shu R, Lyu H, Ma GY, Chen HY, Zhou Y, Zhou JJ, et al. Trends in HIV/AIDS-related mortality and the impact of antiretroviral treatment strategies in Lu'an City: a comprehensive analysis. *Med Sci Monit* 2024;30(1):e944727. <https://doi.org/10.12659/MSM.944727>.
  12. Mugenyi L, Nanfuka M, Byawaka J, Agaba C, Mijumbi A, Kagimu D, et al. Effect of universal test and treat on retention and mortality among people living with HIV-infection in Uganda: an interrupted time series analysis. *PLoS One* 2022;17(5):e0268226. <https://doi.org/10.1371/journal.pone.0268226>.
  13. Wu XS, Wu GH, Ma P, Wang RG, Li LH, Sun YH, et al. Immediate and long-term outcomes after treat-all among people living with HIV in China: an interrupted time series analysis. *Infect Dis Poverty* 2023;12(1):73. <https://doi.org/10.1186/s40249-023-01119-7>.
  14. Ross J, Sinayobye JD, Yotebieng M, Hoover DR, Shi QH, Ribakare M, et al. Early outcomes after implementation of *treat all* in Rwanda: an interrupted time series study. *J Int AIDS Soc* 2019;22(4):e25279. <https://doi.org/10.1002/jia2.25279>.
  15. Edwards JK, Cole SR, Breger TL, Filiaireau LM, Zalla L, Mulholland GE, et al. Five-year mortality for adults entering human immunodeficiency virus care under universal early treatment compared with the general US population. *Clin Infect Dis* 2022;75(5):867 – 74. <https://doi.org/10.1093/cid/ciab1030>.



## SUPPLEMENTARY MATERIAL

### Definitions and Outcomes

**Disease stage** (HIV vs. AIDS refer to HIV infection and AIDS clinical stage) was clinically determined and reported within CRIMS by attending clinicians based on established diagnostic criteria.

**AIDS-related Death** was coded and reported within CRIMS by attending clinicians, classified according to the International Classification of Diseases, 10th Revision (ICD-10) coding framework (B20-B24) as the underlying cause of death, adhering to the WHO's surveillance case definition for AIDS. Since 2011, Liangshan has gradually implemented standardized reporting data for AIDS-related deaths.

**The CAGR** represents the mean annual growth rate of an outcome over a period longer than one year, calculated as  $(\text{Ending value} / \text{Beginning Value})^{(1/n)} - 1$ , where  $n$  is the number of years.

**Follow-up Time** refers to the interval from HIV diagnosis to death, last follow-up, or the cutoff date (for survivors). Outcome events include all-cause mortality and AIDS-related mortality. Censored events refer to patients still active or lost to follow-up as of December 31, 2024.

**Treatment Coverage Rate** was calculated as the proportion of patients receiving ART among all patients diagnosed in a given year  $\{[\text{Number of surviving cases at the beginning of a year} + (\text{Number of newly reported cases in the current year} - \text{Number of deaths in the current year}) / 2] * 100\}$ .

**Mortality Rate** was calculated as the proportion of patients who died among all patients diagnosed in a given year, reflecting the overall mortality rate of HIV-infected individuals (1).

### Interrupted Time Series (ITS) Analysis

Annual treatment coverage rate and mortality rate were used as dependent variables (Y) in the models.

The independent variables of the model included:

The time variable (T) representing the research time points, with values 1, 2, 3,..., 17 corresponding to years 2008-2024, respectively. The intervention variable ( $X_t$ ) indicating the implementation of the comprehensive strategy, with a value of 0 before implementation (2008-2017) and 1 after implementation (2018-2024). The time-intervention interaction term  $(T-T_0)X_t$ , which captures the interaction between the time variable and intervention variable with  $T_0$  as the intervention start time, has a value of 0 before implementation and values of 0, 1, 2,... 6 for subsequent post-implementation years, respectively, to assess the intervention's impact on trends.

### The Counterfactual Models

We calculated counterfactual values by extending the pre-implementation regression models (i.e., excluding  $X_t$  and  $(T-T_0)X_t$  terms). The model formula is:

$$Y_t = \beta_0 + \beta_1 T$$

We then calculated differences between the observed and expected (counterfactual) levels.

### Estimated Effect

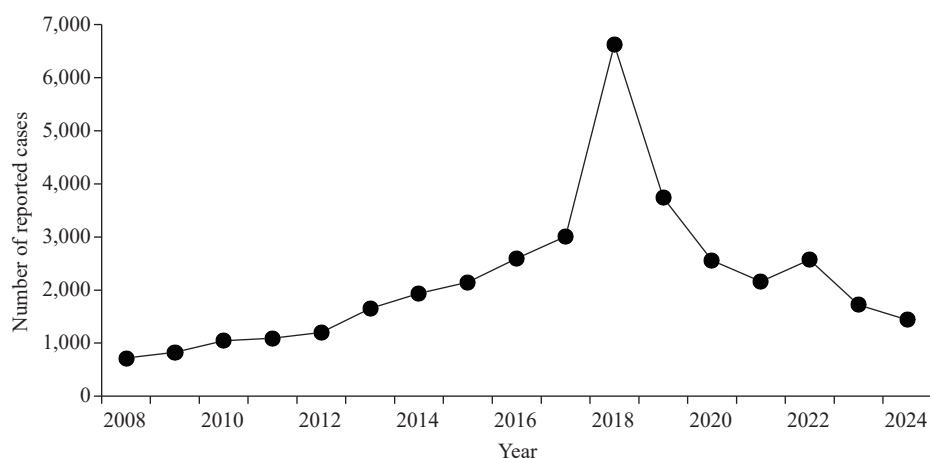
Our analysis evaluated the intervention effects at various timepoints by centralizing time at each timepoint, including both the immediate impact of the comprehensive strategy and its effect by the end of the study period, to estimate reductions in treatment coverage and mortality attributable to the strategy (2).

SUPPLEMENTARY TABLE S1. Characteristics of HIV-infected individuals with heterosexual transmission in Liangshan, 2008 to 2024.

Variables		Total	Before Action Plan (2008–2017)	After Action Plan (2018–2024)	P
Total		37,034	16,326	20,708	
Gender					<0.001
	Male	18,108 (48.9)	7,337 (44.9)	10,771 (52.0)	
	Female	18,926 (51.1)	8,989 (55.1)	9,937 (48.0)	
Age (years)					<0.001
	15–29	12,710 (34.3)	6,776 (41.5)	5,934 (28.7)	
	30–44	15,498 (41.8)	6,726 (41.2)	8,772 (42.4)	
	45–59	6,093 (16.5)	1,741 (10.7)	4,352 (21.0)	
	≥60	2,733 (7.4)	1,083 (6.6)	1,650 (8.0)	
Ethnicity					0.092
	Yi minority	29,839 (80.6)	13,075 (80.1)	16,764 (81.0)	
	Han	6,808 (18.4)	3,082 (18.9)	3,726 (18.0)	
	Others	387 (1.0)	169 (1.0)	218 (1.1)	
Occupation					<0.001
	Farmers	31,803 (85.9)	13,750 (84.2)	18,053 (87.2)	
	Others	5,231 (14.1)	2,576 (15.8)	2,655 (12.8)	
Marital status					<0.001
	Unmarried	6,997 (18.9)	2,788 (17.1)	4,209 (20.3)	
	Married	22,087 (59.6)	10,696 (65.5)	11,391 (55.0)	
	Divorced/Widowed	7,777 (21.0)	2,729 (16.7)	5,048 (24.4)	
	Unknown	173 (0.5)	113 (0.7)	60 (0.3)	
Education					<0.001
	Illiterate	17,614 (47.6)	7,974 (48.8)	9,640 (46.6)	
	Primary School	13,207 (35.7)	5,608 (34.4)	7,599 (36.7)	
	Junior High and Above	6,213 (16.8)	2,744 (16.8)	3,469 (16.8)	
Disease stage					<0.001
	HIV	30,745 (83.0)	13,367 (81.9)	17,378 (83.9)	
	AIDS	6,289 (17.0)	2,959 (18.1)	3,330 (16.1)	
Baseline CD4 count (cell/μL)*					<0.001
	<200	5,802 (15.7)	2,121 (13.0)	3,681 (17.8)	
	201–350	10,336 (27.9)	3,891 (23.8)	6,445 (31.1)	
	351–500	9,050 (24.4)	4,276 (26.2)	4,774 (23.1)	
	>500	9,855 (26.6)	4,838 (29.6)	5,017 (24.2)	
	Missing	1,991 (5.4)	1,200 (7.4)	791 (3.8)	
Region					<0.001
	Non-key counties	18,319 (49.5)	7,597 (46.5)	10,722 (51.8)	
	Key counties	18,715 (50.5)	8,729 (53.5)	9,986 (48.2)	

Note: Data presented as *n* (%).

\* refers to the initial CD4 count at diagnosis. Key counties include Butuo, Zhaojue, Yuexi, and Meigu. Non-key counties comprise all remaining counties in Liangshan Prefecture.



SUPPLEMENTARY FIGURE S1. Annual reported cases of heterosexually transmitted HIV-infected individuals in Liangshan Prefecture from 2008 to 2024.

Abbreviation: HIV=human immunodeficiency virus.

## REFERENCES

1. Zhang HX, Han MJ, Zhou Y, Xiu XF, Xu F, Wang L. Interrupted time series analysis for influence on HIV related fatality of implementation of 'Four Free Services One Care' policy in China. *Chin J Epidemiol* 2020;41(3):406 – 11. <https://doi.org/10.3760/cma.j.issn.0254-6450.2020.03.024>.
2. Wu XS, Wu GH, Ma YM, Huang XJ, Yang YC, Cai YS, et al. The impact of COVID-19 non-pharmaceutical interventions on HIV care continuum in China: an interrupted time series analysis. *Lancet Reg Health West Pac* 2022;29:100569. <https://doi.org/10.1016/j.lanwpc.2022.100569>.