

THE LANCET

Global Health

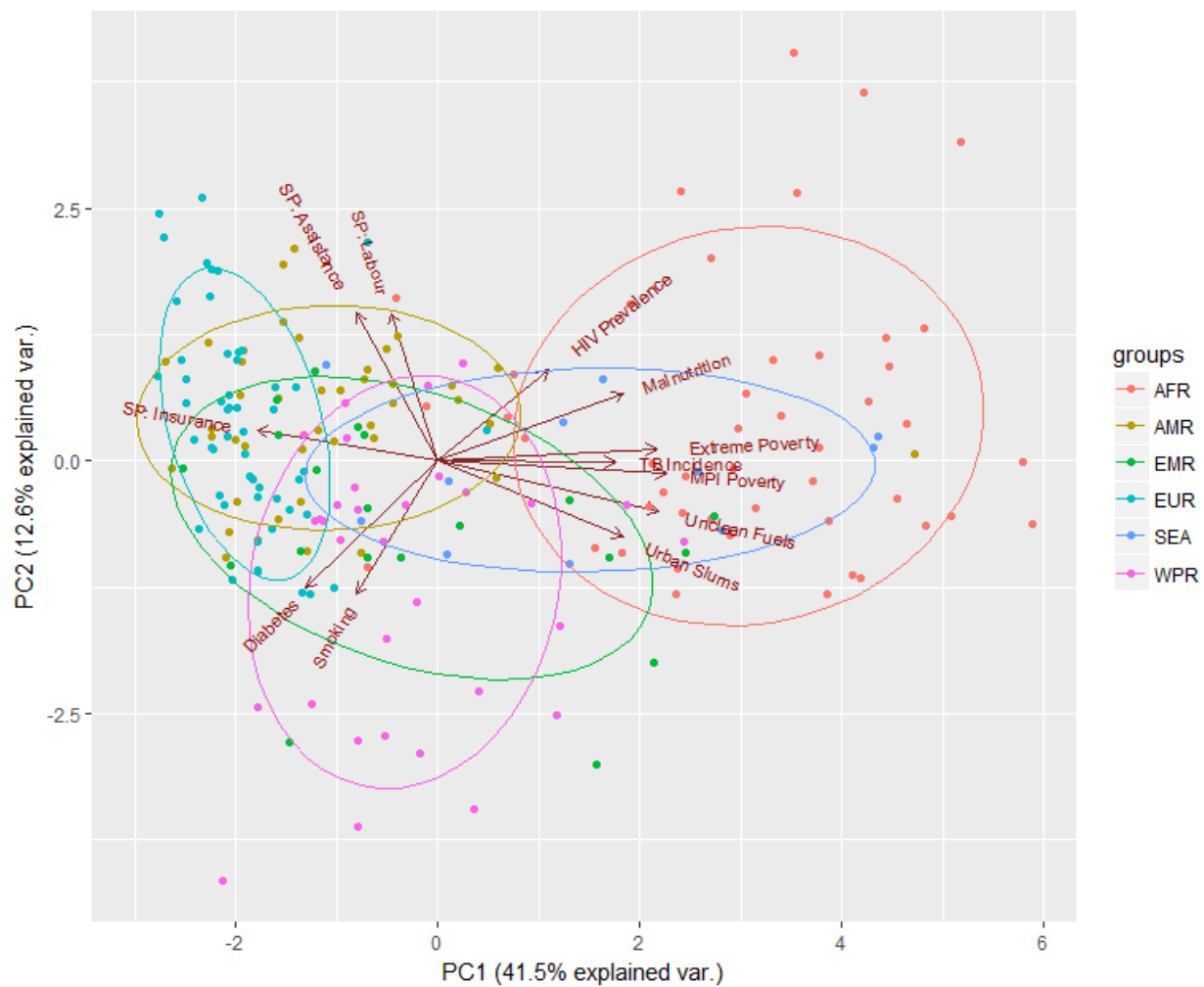
Supplementary appendix

This appendix formed part of the original submission and has been peer reviewed. We post it as supplied by the authors.

Supplement to: Carter DJ, Glaziou P, Lönnroth K, et al. The impact of social protection and poverty elimination on global tuberculosis incidence: a statistical modelling analysis of Sustainable Development Goal 1. *Lancet Glob Health* 2018; published online March 23. [http://dx.doi.org/10.1016/S2214-109X\(18\)30195-5](http://dx.doi.org/10.1016/S2214-109X(18)30195-5).

635 Appendix 1 –PCA

636 To examine relationships between selected variables, a PCA was performed after conducting a single
 637 imputation by region using the regression substitution method. Five imputations of the dataset were
 638 generated using multiple imputation by chained equations and the data used to generate this biplot
 639 comes from a randomly selected imputed dataset.²⁰ Smoking and diabetes cluster together and not
 640 using clean fuels and living in an urban slum also cluster together, informing a decision to combine
 641 variables. Extreme poverty and MPI poverty are also closely related, suggesting they may be collinear.
 642 Both are quite closely related to TB incidence, as expected. Malnutrition is also quite strongly related to
 643 TB incidence. Social insurance is negatively correlated with TB incidence as expected, and closely
 644 associated with regions with high and middle income countries.
 645



646

647

648 **Figure A.** Biplot of the first two principal components of variables in the conceptual framework. Closeness of the
649 arrows indicates the degree of correlation between the variables. The direction of the arrows points in the
650 direction of increasing values of that variable. Ellipses represent groupings by WHO region.

651

652

653

654 Appendix 2 - Missing Data

655

656 Table A presents the level of missingness in the data prior to the multiple imputation procedure. The
 657 multiple imputation procedure works by running iterated regression analyses, taking one variable as the
 658 outcome variable and the remaining variables in the dataset as the predictor variables. Doing this for all
 659 variables results in a set of predicted values (imputed values) for each missing value ²⁰. Repeating this
 660 procedure multiple times provides multiple datasets with no missing data which are pooled together to
 661 perform analyses.

662

663 The advantage of undertaking this procedure is that the imputed values for a given country closely
 664 resemble values for countries with similar characteristics in the other variables, and are thus more likely
 665 to accurately reflect possible values of the missing data. Doing this multiple times ensures convergence
 666 on a reasonable value. In this dataset, missingness on social insurance and missingness on absolute
 667 poverty was not associated with any of the four proximal TB risk factors, nor was it associated with WHO
 668 region.

669

670

671

Subtarget Indicator Measures	% missing data
SDG 1.1: Absolute Poverty	42.0%
SDG 1.2: Multidimensional Poverty Index (prevalence)	52.2%
SDG 1.3: Labour Market Social Protection	71.7%
SDG 1.3: Social Assistance	48.3%
SDG 1.3: Social Insurance	49.3%
Poverty Subconstruct Measures	% missing data
Malnourished %	0.01%
HIV Prevalence	22.9%
Health Behaviours	39.0%
Housing Quality	54.6%

672 **Table A.** Proportion of missing data for each subtarget indicator and for each measure of poverty. Missingness in
 673 the context of these data is that there was no recorded value after 2010.

674 Appendix 3 – Mediators

675

676 A simple analysis for mediation was performed by adjusting the estimate for the linear relationship
 677 between SDG 1 and TB for each of the risk factors in turn. Evidence for mediation is determined by the
 678 degree of change in the relationship between SDG 1 and TB is adjusted for with the risk factor under
 679 examination. All risk factors will show a degree of mediation for extreme poverty and social insurance.
 680

Indicator		Crude	Adjusted	Mediation?
SDG 1.1: Extreme Poverty	Malnutrition	3.23***	2.29***	Moderate
	HIV	3.23***	1.99***	Moderate
	Health Behaviours	3.23***	4.67***	Moderate
	Housing Quality	3.23***	2.43	Moderate
SDG 1.2: MPI Prevalence	Malnutrition	1.94***	1.17*	Weak
	Health Behaviours	1.94***	2.32**	Weak
	Housing Quality	1.94***	-1.44	Strong
SDG 1.3: Social Assistance	Malnutrition	-0.81	-0.49	Weak
	Housing Quality	-0.81	0.38	Moderate
SDG 1.3: Social Insurance	Malnutrition	-3.73***	-2.79**	Moderate
	HIV	-3.73***	-2.00**	Moderate
	Health Behaviours	-3.73***	-4.98***	Moderate
	Housing Quality	-3.73***	-1.90	Strong

681 Stars indicate significant difference of coefficient from 0: ***p < 0.001, ** p < .01, * p < .05

682 **Table B.** Effect estimates for SDG indicators and TB incidence adjusting for a sole mediator, where included
683 mediators are associated with the SDG indicator in question and TB incidence (Table X; Figure B). Pathways are
684 assumed independent.

685

686

687

688
689
690
691
692
693
694
695
696
697
698
699
700
701
702
703
704
705
706
707

Appendix 4 – Income Analyses

GDP per capita and the Gini coefficient were entered into the multiple imputation model and included as a covariate in the adjusted linear models used for mediation analysis as a sensitivity analysis. Neither GDP nor Gini coefficient had a strong association with TB incidence adjusting for the four TB risk factors. However, adjustment for the Gini coefficient resulted in a loss of conventional statistical significance for the health behaviour pathway, and a gain of conventional statistical significance for the social insurance pathway though the direction and magnitude of effect are still consistent with the prior results without income adjustment. This loss of conventional statistical significance is expected given the higher prevalence of both smoking and diabetes in high and middle-income countries that is likely captured in part by GDP. This result is not unexpected as GDP and Gini coefficient are likely to be distal (upstream) drivers of the four more proximal risk factors and thus some of the association of TB with smoking and diabetes may be accounted for in adjustment for GDP and Gini. As the direction and magnitude of effect still hold, we conclude that even after adjustment for income effects, health behaviour is likely to remain a mediator of the relationship between the SDG subtargets and TB. The results may indicate that the health behaviour pathway is responsible for less of the total effect of the SDG subtargets and TB than malnutrition, HIV, or housing quality.

Association with TB Incidence (adjusting additionally for GDP per capita and Gini coefficient)	
Indicator	Adjusted Estimate (more and equally proximal; blue & orange layer)
SDG 1.1: Extreme Poverty	0.87 (-0.89, 2.64)
SDG 1.2: MPI Prevalence	-0.35 (-2.13, 1.44)
SDG 1.3: Labour Protection	-1.09 (-4.58, 2.41)
SDG 1.3: Social Assistance	-0.09 (-0.87, 0.69)
SDG 1.3: Social Insurance	-1.52 (-2.84, -0.20)
Income Covariate	Adjusted Estimate
GDP per capita (in 1000s)	-1.36 (-2.60, 0.12)
Gini Coefficient	0.29 (-2.37, 2.94)

Measure	Adjusted Estimate
Malnourishment	2.75 (0.64, 4.85)
Health Behaviour	2.49 (-0.48, 5.45)
Housing Quality	1.54 (0.55, 2.54)
HIV Prevalence	20.44 (13.25, 27.63)

708 Bold indicates significant difference of coefficient from 0 with $p < .05$

709 **Table C.** The association between each indicator or measure and TB incidence per 100,000 per year. The adjusted
710 estimate represents the association adjusted for all variables more or equally proximal to TB incidence as well as
711 GDP per capita and Gini coefficient for a pooling of 50 datasets from a multiple imputation procedure.

712

713

714

715 Appendix 5 – Sensitivity Analyses to Modelling Assumptions for selected pathways
 716 Complete results from the model at 50% coverage of social protection (an increase of ~30% from baseline) and 90%
 717 elimination of extreme poverty (an increase of ~10% from baseline) are presented in Table D. There remains a
 718 substantial decrease in TB incidence predicted even from partial attainment of one or both of the SDG subtargets.
 719
 720 Results from producing projections for 2035 incidence using only countries with complete information for social
 721 insurance and extreme poverty (n = 94) are presented in Table E. The estimates are broadly similar to the
 722 estimates produced with multiple imputation, but with wider credible intervals – this is expected given the
 723 reduced power from decreased sample size, but also because of the exponential component of the model. The
 724 upper bound of the credible interval, if close to baseline, will decrease much less rapidly than its lower bound
 725 counterpart.

726
727
728

Table D. Projected impact on 2035 TB incidence by mid-level achievement of selected subtargets

Pathway	β (95% Confidence Intervals)	Estimated 2035 Incidence (/100,000)	95% Credible Interval (/100,000)	Reduction from Baseline* (95% CIs)	Expected Annual Proportional Decline
<i>Pathway A:</i> Social Protection for All (50% coverage)	-2.13 (-3.36, -0.90)	69.6 cases	(50.9, 94.8) cases	41.6% (20.5%, 57.3%)	3.3%
<i>Pathway C:</i> Extreme Poverty Elimination (90% Eliminated)	-2.37 (-3.64, -1.10)	97.6 cases	(89.0, 110.1) cases	18.1% (7.6%, 25.3%)	1.3%
<i>Pathway A & Pathway C:</i> Social Protection and Poverty Elimination	---	56.7 cases	(36.9, 86.2) cases	52.5% (27.7%, 69.6%)	4.6%

729
730 **Table D.** The projected TB incidence per 100,000 per year associated with the achievement of selected SDG 1
731 subtargets independent from the other targets – 50% social protection coverage and 90% extreme poverty
732 elimination.

733
734

Table E. Projected impact on 2035 TB incidence: Complete cases

Pathway	β (95% Confidence Intervals)	Estimated 2035 Incidence (/100,000)	95% Credible Interval (/100,000)	Reduction from Baseline* (95% CIs)	Expected Annual Proportional Decline
<i>Pathway A:</i> Social Protection for All	-2.66 (-4.36, -0.95)	76.7 cases	(57.8, 101.6) cases	35.6% (14.8%, 51.6%)	2.7%

(100% coverage)

<i>Pathway C:</i>	-2.01	30.9 cases	(7.8, 112.5)	74.0%	8.2%
Extreme Poverty Elimination (100% Eliminated)	(-3.96, -0.08)		cases	(5.6%, 93.5%)	
<i>Pathway A & Pathway C:</i>	---	19.3 cases	(3.4, 95.8)	83.8%	10.9%
Social Protection and Poverty Elimination			cases	(19.7%, 97.2%)	

735

736

737

738

739

740

741

742

743

744

745

746

Table E. The projected TB incidence per 100,000 per year associated with the achievement of selected SDG 1 subtargets – 100% social protection coverage and 100% extreme poverty elimination. These numbers are from a data subset with complete information on social insurance and extreme poverty.