

Table S1. P values for associations between TB status and each of the host factors shown in Table 1, by age group *

	All (≥ 18 y/o) p value	YA (18-44 y/o) p value	MAA (45-64 y/o) p value	ELD (≥ 65 y/o) p value
Sociodemographics				
Male sex	<0.001	<0.001	<0.001	<0.001
Education, High school or higher	<0.001	<0.001	0.518	0.562
Current or past smoker	<0.001	<0.001	0.007	0.036
TB-related variables				
Recent TB exposure (ReC vs CoC)				
Past TB	<0.001	0.008	0.162	0.027
BCG	<0.001	0.078	0.295	0.002
Type 2 diabetes and other conditions				
Type 2 diabetes	<0.001	<0.001	<0.001	0.321
Overweight/obese, BMI ≥ 25	<0.001	<0.001	<0.001	<0.001
Central obesity (M ≥ 0.90 M; F ≥ 0.86)	<0.001	<0.001	0.001	0.313
High cholesterol (200 mg/dL)	<0.001	0.069	<0.001	<0.001
High LDL (100 mg/dL)	<0.001	0.001	<0.001	<0.001
Low HDL (40 M, 50 F, mg/dL)	0.607	0.866	0.266	0.748
High Triglycerides (150 mg/dL)	<0.001	<0.001	<0.001	<0.001
Macrovascular diseases	0.234	0.282	0.052	0.014
Microvascular diseases	<0.001	0.008	<0.001	0.356
Anti-inflammatory medications	0.001	0.001	0.131	0.823

* Data from Table 1 was used to identify associations between host characteristics and TB status among all participants or by age group. Analysis was done by chi-square or Fisher's exact; $p \leq 0.099$ shown in bold; Normal range values shown in parenthesis; M, males; F, females.

Table S2. Multivariable logistic regression models for the association between diabetes (exposure) and TB (outcome) by age group *

	All (≥ 18 y/o)	YA (18-44 y/o)	MAA (45-64 y/o)	ELD (≥ 65 y/o)
Models controlling for core variables associated with TB outcomes in all age groups as shown in Table S1				
Diabetes (vs no diabetes)	4.12 (2.86, 5.93)	6.48 (3.35, 12.53)	5.4 (2.95, 9.89)	1.87 (0.75, 10.95)
Male sex (vs female)	3.4 (2.34, 4.94)	2.79 (1.58, 4.91)	4.34 (2.37, 7.92)	4.03 (1.48, 46.78)
BMI, (UW/NOR vs OW/OB)	12.03 (8.31, 17.42)	18.97 (10.28, 35.01)	7.82 (4.38, 13.95)	16.22 (5.63, 4.14)
Current or past smoker (vs never)	1.51 (1.03, 2.22)	1.1 (0.61, 2)	2.05 (1.13, 3.74)	1.46 (0.52, 0)
Sensitivity analysis. Models controlling for variables associated with TB depending on the age group, as shown in Table S1				
Diabetes (vs no diabetes)	3.64 (2.51, 5.29)	5.3 (2.65, 10.62)	5.19 (2.79, 9.67)	1.81 (0.72, 43.09)
Male sex (vs female)	3.47 (2.36, 5.1)	2.73 (1.51, 4.96)	4.65 (2.47, 8.75)	3.78 (1.38, 45.81)
BMI, (UW/NOR vs OW/OB)	12.04 (8.24, 17.59)	19.46 (10.37, 36.53)	7.82 (4.29, 14.26)	14.78 (5.07, 10.34)
Current or past smoker (vs never)	1.51 (1.02, 2.24)	1.07 (0.57, 2.01)	2.16 (1.17, 3.98)	1.46 (0.52, 43.09)
Lower education (vs HS or higher)	1.52 (1.03, 2.23)	1.96 (1.13, 3.4)	1.48 (0.77, 2.85)	NT
BCG at birth (vs none)	NT	NT	NT	0.56 (0.19, 45.81)
Pain/NSAID use (vs none)	1.94 (1.29, 2.9)	3.05 (1.59, 5.84)	1.69 (0.88, 3.27)	NT
* Data expressed as adjusted odds ratio (95% CI) after controlling for the variables analyzed for each model; BMI, body-weight index; UW/NOR, underweight or normal weight; OW/OB, overweight or obese; Pain/NSAID, report use of pain medication, including non-steroidal anti-inflammatory drug (NSAID) use in past month; NT, variable not entered into the model because p value above 0.100 by univariable analysis in the corresponding age group; HS, High school degree				

Table S3. Unique characteristics of diabetes patients without TB for laboratory data, by age groups

	All (≥ 18 y/o)		YA (18-44 y/o)		MAA (45-64 y/o)		ELD (≥ 65 y/o)		
	n=241		n=60		n=134		n=47		
Lipids (mg/dL)	n	%	n	%	n	%	n	%	p value
High cholesterol (200)	68	28.2%	16	26.7%	40	29.9%	12	25.5%	0.822
Low HDL (40 M, 50 F)	179	74.3%	49	81.7%	97	72.4%	33	70.2%	0.306
High LDL (100)	126	52.3%	28	46.7%	76	56.7%	22	46.8%	0.459
High Triglycerides (150)	110	45.6%	27	45.0%	65	48.5%	18	38.3%	0.478
Complete Blood Counts (x10e3/μL)	n	median (IQR)	n	median (IQR)	n	median (IQR)	n	median (IQR)	p value
Platelets (146-388)	203	252 (95.00)	49	253 (101.00)	111	264 (85.00)	43	217 (71.00)	<0.001
Eosinophils (0.0-0.4)	213	0.17 (0.17)	50	0.18 (0.21)	119	0.16 (0.14)	44	0.21 (0.24)	0.087
Lymphocytes (0.8-3.3)	213	2.13 (0.86)	50	2.17 (0.92)	119	2.15 (0.87)	44	2.03 (0.72)	0.380
Monocytes (0.2-1.0)	213	0.45 (0.17)	50	0.46 (0.15)	119	0.45 (0.18)	44	0.45 (0.22)	0.799
Neutrophils (2.3-7.7)	213	4.29 (1.90)	50	4.66 (1.76)	119	4.28 (1.76)	44	3.93 (1.73)	0.026
White blood cells (4.8-10.9)	213	7.20 (2.40)	50	7.55 (3.40)	119	7.30 (2.00)	44	6.65 (2.40)	0.050
Hemoglobin (11.6-15.9 mg/dL)	204	13.70 (1.70)	49	13.90 (1.80)	111	13.70 (1.60)	44	13.20 (2.00)	0.005
Oxidation status and Vitamin D	n	median (IQR)	n	median (IQR)	n	median (IQR)	n	median (IQR)	p value
Reduced:oxidized glutathione ratio	97	0.58 (0.10)	21	0.58 (0.09)	45	0.58 (0.14)	31	0.60 (0.09)	0.752
Vitamin D levels (ng/mL)	98	26.09 (11.14)	18	24.59 (8.80)	53	24.59 (10.29)	27	29.09 (13.58)	0.333

Data expressed as column % for categorical variables or median (interquartile range, IQR) for continuous; Normal range values for each parameter shown in parenthesis; M, males; F, Females; p values ≤ 0.099 shown in bold; Bold font in young or middle age adults indicates significant or borderline significant differences when compared to elderly group. YA, young adults; MAA, middle age adults; ELD, elderly.

Table S4. Extended analysis of data from Table 2: Crude and adjusted p values for host characteristics associated with the ELD age group, in T2D-No TB participants

	YA vs ELD	MAA vs ELD	YA vs ELD	MAA vs ELD	YA vs ELD	MAA vs ELD
	Crude p		Adj p (Sex, SFU)		Adj p (Sex, NSAID)	
Sociodemographics, medications and TB-related characteristics						
Male sex	0.047	0.732				
Education, High school or higher	<0.001	0.002	<0.001	0.197	<0.001	0.186
Current smoker	0.161	0.659	0.449	0.386	0.327	0.431
NSAID use	0.018	0.394	0.038	0.477		
BCG vaccination	0.594	0.616	0.549	0.825	0.927	0.481
Latent TB infection	0.671	0.828	0.487	0.489	0.697	0.732
Diabetes history						
Family history of diabetes	0.064	0.115	0.059	0.645	0.253	0.419
Self-reported diabetes	0.261	0.951	0.784	0.516	0.569	0.549
Years with diabetes, yrs	0.001	0.021	0.001	0.002	0.001	0.002
Glucose management						
Hyperglycemia (125 mg/dL)	0.014	0.041	0.114	0.329	0.192	0.426
HbA1c (%)	0.037	0.131	0.002	0.020	0.010	0.058
Glycemic Index (HbA1c*T2Dears)	0.009	0.073	0.041	0.095	0.015	0.076
Insulin Levels (mU/L)	0.158	0.339	0.741	0.187	0.413	0.325
HOMA-IR (Median, IQR)	0.002	0.002	0.001	0.004	0.001	0.005
Diabetes medications in past month						
Any	0.052	0.793	0.702	0.364	0.213	0.369
Insulin	0.276	0.303	0.330	0.634	0.704	0.571
Metformin	0.304	0.383	0.882	0.086	0.424	0.072
Sulfonylureas	0.001	0.127			0.007	0.481
Metformin + Sulfonylureas	0.021	0.572			0.056	0.241
Diabetes-associated conditions						
Body-mass index	0.009	0.305	0.031	0.328	0.023	0.297
Central obesity (M ≥ 0.90 M; F ≥ 0.86)	0.037	0.005	0.255	0.083	0.444	0.090
Macrovascular diseases	<0.001	0.004	<0.001	0.862	<0.001	0.968
Microvascular diseases	0.218	0.231	0.770	0.491	0.208	0.823

* Data from Table 2 was used to expand analysis on associations between host characteristics and age groups in T2D-no TB participants. P values calculated by chi-square or Fisher's exact (crude p), multivariable logistic

regression (Proc LOGISTIC in SAS; categorical variables) or generalized linear models (proc GENMOD in SAS; continuous variables); $p \leq 0.099$ shown in bold; Cut-off values shown in parenthesis; M, males; F, females.

Table S5. Extended analysis of data from Table S3: Crude and adjusted p values for laboratory characteristics associated with the ELD age group, in T2D-No TB participants

	YA vs ELD	MAA vs ELD	YA vs ELD	MAA vs ELD	YA vs ELD	MAA vs ELD
	Crude p		Adj p (Sex, SFU)		Adj p (Sex, NSAID)	
Lipids (mg/dL)						
High cholesterol (200)	0.947	0.627	0.897	0.623	0.784	0.428
Low HDL (40 M, 50 F)	0.906	0.911	0.193	0.477	0.077	0.353
High LDL (100)	0.226	0.257	0.517	0.133	0.259	0.076
High Triglycerides (150)	0.486	0.243	0.735	0.294	0.784	0.299
Complete Blood Counts (x10e3/μL)						
Platelets (146-388)	0.005	<0.001	0.001	<0.001	0.001	<0.001
Eosinophils (0.0-0.4)	0.249	0.031	0.064	0.001	0.100	0.002
Lymphocytes (0.8-3.3)	0.196	0.236	0.057	0.100	0.045	0.091
Monocytes (0.2-1.0)	0.916	0.652	0.766	0.335	0.982	0.376
Neutrophils (2.3-7.7)	0.010	0.071	0.007	0.044	0.003	0.038
White blood cells (4.8-10.9)	0.022	0.066	0.016	0.086	0.007	0.070
Hemoglobin (11.6-15.9 mg/dL)	0.003	0.012	0.003	0.004	0.002	0.003
Oxidation status and Vitamin D levels						
Reduced to oxidized glutathione ratio	0.830	0.450	NS	NS	0.722	0.221
Vitamin D levels (ng/mL)	0.276	0.156	NS	NS	0.431	0.335

Data from Table S3 was used to expand analysis on associations between host laboratory characteristics and age groups in T2D-no TB participants. P values calculated by chi-square or Fisher's exact (crude p), multivariable logistic regression (Proc LOGISTIC in SAS; categorical variables) or generalized linear models (proc GENMOD in SAS; continuous variables); p \leq 0.099 shown in bold; Cut-off values shown in parenthesis; M, males; F, females.

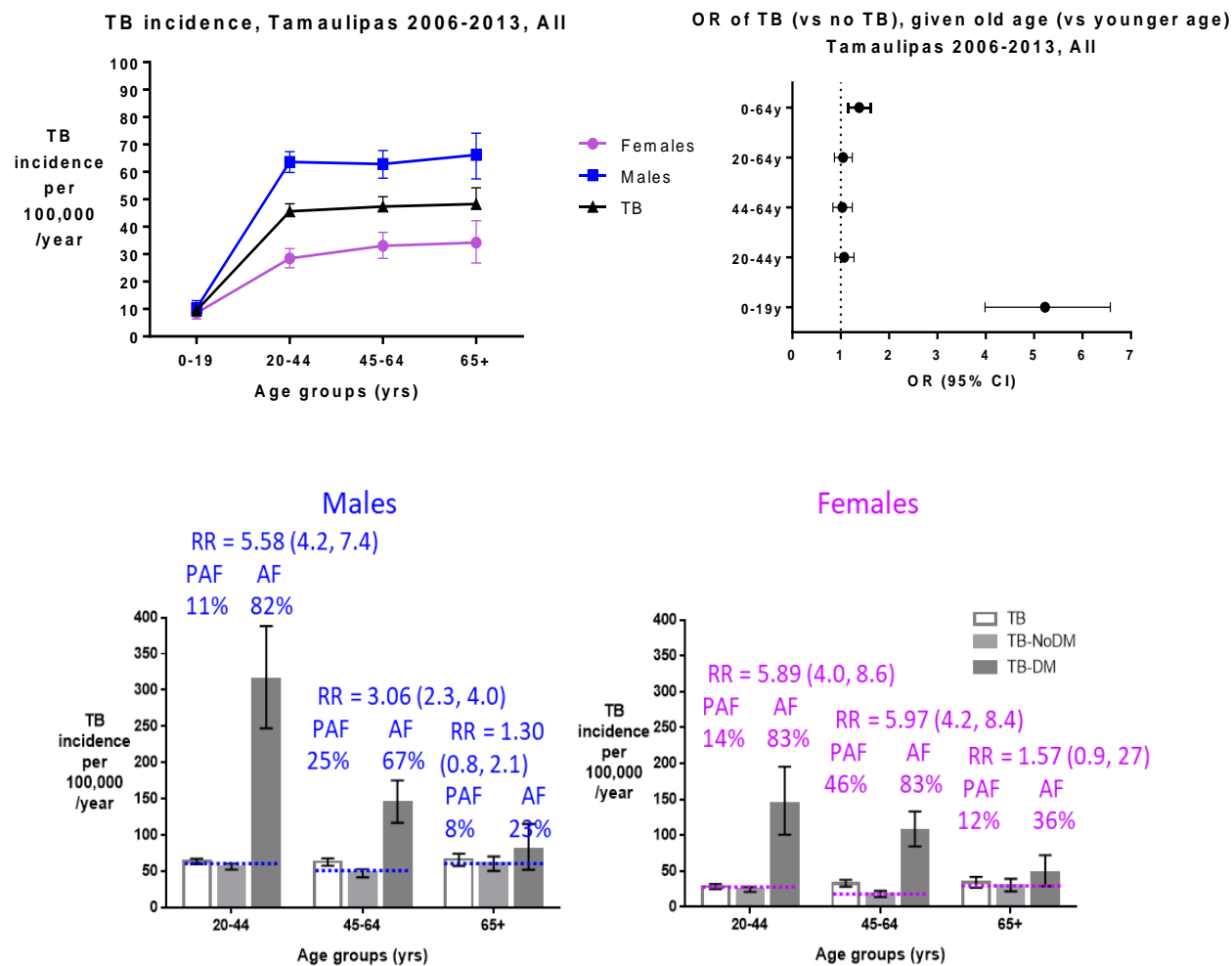


Figure S1. Sensitivity analysis for results from the validation dataset (Tamaulipas 2006-2013) without removal of participants with HIV, intravenous drug use or excess alcohol. A. Prevalence of TB by sex and age. B. OR of TB vs no TB in the ELD vs: YA (OR 1.06, 95%CI 0.88, 1.27), MAA (OR 1.02, 95%CI 0.84, 1.24), all adults (OR 1.04, 95%CI 0.87, 1.24), or all age groups (OR 1.37, 95%CI 1.48, 1.63). C. Relative risk, attributable fraction and population attributable fraction due to T2D, by sex and age group. Symbols and abbreviations: Dots, point estimates; Error bars, 95% confidence intervals; Horizontal dotted line, reference TB prevalence among individuals without T2D for calculation of T2D attributable fraction and population attributable fraction; RR, relative risk; PAF, T2D population attributable risk fraction for TB; AF, T2D attributable risk fraction for TB. CHILD, 0-19 y/o; YA, 20-44y/o; MAA, 45-64 y/o; ELD, ≥65 y/o.

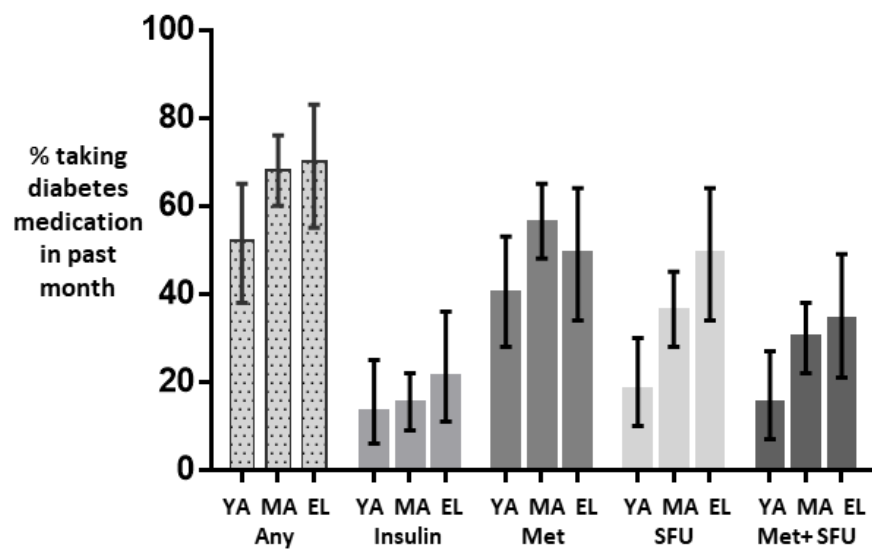


Figure S2. Proportion of diabetes patients using hypoglycemic agents by age group. Few participants also reported use of TZDs, SGLT2I, DPPI or GLP-1 agonists (not shown). Vertical bars, 95% CI; SFU, sulphonylureas; Met, metformin