

SUPPLEMENTARY DATA

**Supplementary Table 1. Association analyses between current shift work ( $n = 189,488$ ) and morningness-eveningness preference ( $n = 169,926$ ) with unadjusted/adjusted odds of type 2 diabetes and mean difference in HbA1c (in mmol/mol) across shift work and morningness-eveningness preference categories in employed UK Biobank participants of European descent.**

	Type 2 diabetes			N	HbA1c (mmol/mol)	
	Type 2 diabetes cases /controls	Unadjusted OR [95% CI]	Sleep duration adjusted OR [95% CI]		Unadjusted Beta [95% CI]	Sleep duration adjusted Beta [95% CI]
<b>Shift work</b>						
Day workers	4,047 /154,792	<i>reference</i>	<i>reference</i>	146,993	<i>reference</i>	<i>reference</i>
Shift work without nights	475 /14,863	<b>1.22 [1.11-1.35]</b>	<b>1.26 [1.14-1.39]</b>	14,110	<b>0.29 [0.22-0.35]</b>	<b>0.33 [0.27-0.39]</b>
Sometimes night shift work	284 /8,434	<b>1.29 [1.14-1.46]</b>	<b>1.32 [1.17-1.49]</b>	8,005	<b>0.25 [0.17-0.33]</b>	<b>0.46 [0.38-0.54]</b>
Usual night shift work	80 /2,171	<b>1.41 [1.13-1.77]</b>	<b>1.47 [1.17-1.85]</b>	2,069	0.16 [0-0.32]	<b>0.43 [0.27-0.58]</b>
Always night shift work	156 /4,186	<b>1.43 [1.21-1.68]</b>	<b>1.45 [1.23-1.71]</b>	3,979	<b>0.54 [0.43-0.66]</b>	<b>0.72 [0.60-0.83]</b>
<b>Morningness-eveningness preference</b>						
Definite-morning	1,272 /42,097	<i>reference</i>	<i>reference</i>	39,976	<i>reference</i>	<i>reference</i>
More morning than evening	1,482 /60,064	<b>0.82 [0.76-0.88]</b>	<b>0.86 [0.8-0.93]</b>	57,127	<b>-0.25 [-0.3--0.2]</b>	<b>-0.12 [-0.17--0.08]</b>
More evening than morning	1,268 /48,593	<b>0.86 [0.8-0.93]</b>	0.96 [0.89-1.04]	46,267	<b>-0.28 [-0.32--0.23]</b>	-0.03 [-0.08-0.01]
Definite-evening	497 /14,653	<b>1.12 [1.01-1.25]</b>	<b>1.30 [1.17-1.44]</b>	13,886	<b>-0.17 [-0.24--0.1]</b>	<b>0.11 [0.04-0.18]</b>

**Legend:** Prevalent type 2 diabetes associations are unadjusted and then sex-, age-, and sleep-duration adjusted odds ratios [95% confidence interval]. HbA1c associations are restricted to participants with no prevalent type 2 diabetes. HbA1c associations are unadjusted and then sex-, age-, and sleep-duration adjusted betas [95% confidence interval] in mmol/mol. Bold  $P < 0.05$ .

SUPPLEMENTARY DATA

**Supplementary Table 2. Adjusted odds ratios (OR) or adjusted betas and 95% confidence intervals (CI) of type 2 diabetes and HbA1c (in mmol/mol) with each additional copy of the *MTNR1B* G risk-allele across categories of current work schedule with further adjustment for morningness-eveningness preference ( $n = 169,926$ ).**

	Type 2 diabetes			HbA1c (mmol/mol)		
	Type 2 diabetes cases /controls	OR [95% CI]	$P_{int}$	$N$	Beta [95% CI]	$P_{int}$
<b>Overall</b>	4,519 /165,407	1.10 [1.05-1.15]	0.10	157,256	0.26 [0.23-0.29]	0.34
<b>Day workers</b>	3,634 /139,090	1.09 [1.03-1.15]		132,249	0.25 [0.22-0.28]	
<b>Shift work without nights</b>	430 /13,389	1.25 [1.07-1.45]		12,720	0.32 [0.22-0.42]	
<b>Sometimes night shift work</b>	247 /7,450	0.94 [0.76-1.16]		7,074	0.35 [0.22-0.48]	
<b>Usual night shift work</b>	70 /1,889	0.84 [0.56-1.27]		1,799	0.21 [-0.05-0.46]	
<b>Always night shift work</b>	138 /3,589	1.24 [0.94-1.63]		3,414	0.18 [-0.01-0.37]	

**Legend:** Association results are adjusted odds ratios [95% confidence interval] of type 2 diabetes per each additional copy of the *MTNR1B* G risk-allele or adjusted betas [95% confidence interval] describing differences in HbA1c in mmol/mol per each additional copy of the *MTNR1B* G risk-allele across categories of current work schedule. Association analyses are adjusted for age, sex, BMI, genotyping array, 10 principal components of ancestry, and morningness-eveningness preference.  $P_{int}$  is log likelihood ratio test comparing models with and without cross-product interaction terms (*MTNR1B* and current work schedule) including main effect terms in logistic or linear regression models adjusted for the aforementioned covariates.

SUPPLEMENTARY DATA

**Supplementary Table 3. Adjusted odds ratios (OR) or adjusted betas and 95% confidence intervals (CI) of type 2 diabetes and HbA1c (in mmol/mol) with each additional copy of the *MTNR1B* G risk-allele across categories of morningness-eveningness preference with further adjustment for current work schedule ( $n = 169,926$ ).**

	Type 2 diabetes			HbA1c (mmol/mol)		
	Type 2 diabetes cases /controls	OR [95% CI]	$P_{int}$	$N$	Beta [95% CI]	$P_{int}$
<b>Overall</b>	4,519/165,407	1.10 [1.04-1.15]	0.044	157,256	0.26 [0.23-0.29]	0.86
<b>Definite morning</b>	1,272/42,097	1.17 [1.07-1.28]		39,976	0.30 [0.25-0.36]	
<b>More morning than evening</b>	1,482/60,064	1.09 [1.00-1.18]		57,127	0.23 [0.19-0.28]	
<b>More evening than morning</b>	1,268/48,593	1.06 [0.97-1.16]		46,267	0.23 [0.18-0.28]	
<b>Definite evening</b>	497/14,653	1.01 [0.87-1.18]		13,886	0.36 [0.27-0.46]	

**Legend:** Association results are adjusted odds ratios [95% confidence interval] of type 2 diabetes per each additional copy of the *MTNR1B* G risk-allele or adjusted betas [95% confidence interval] describing differences in HbA1c in mmol/mol per each additional copy of the *MTNR1B* G risk-allele across categories of morningness-eveningness preference. Association analyses are adjusted for age, sex, BMI, genotyping array, 10 principal components of ancestry, and current work schedule.  $P_{int}$  is log likelihood ratio test comparing models with and without cross-product interaction terms (*MTNR1B* and morningness-eveningness preference) including main effect terms in logistic or linear regression models adjusted for the aforementioned covariates.

SUPPLEMENTARY DATA

**Supplementary Table 4. Sensitivity analyses of morningness-eveningness preference association with adjusted odds of type 2 diabetes ( $n = 298,953$ ) and adjusted mean difference in HbA1c (in mmol/mol;  $n = 272,220$ ) in UK Biobank participants of European descent regardless of employment status.**

	Type 2 diabetes		HbA1c (mmol/mol)	
	Type 2 diabetes cases /controls	Sex- and age-adjusted OR [95% CI]	N	Sex- and age-adjusted Beta [95% CI]
<b>Morningness-eveningness preference</b>				
Definite-morning	3,490 /75,909	<i>reference</i>	72,013	<i>reference</i>
More morning than evening	4,006 /104,746	<b>0.86 [0.82-0.90]</b>	99,552	<b>-0.12 [-0.15--0.08]</b>
More evening than morning	3,455 /81,559	1.01 [0.96-1.06]	77,517	-0.03 [-0.06-0.01]
Definite-evening	1,373 /24,415	<b>1.40 [1.31-1.49]</b>	23,138	<b>0.17 [0.12-0.22]</b>

**Legend:** Prevalent type 2 diabetes associations are sex-, and age- adjusted odds ratios [95% confidence interval]. HbA1c associations are restricted to participants with no prevalent type 2 diabetes. HbA1c associations are sex- and age-, adjusted betas [95% confidence interval] in mmol/mol. Bold  $P < 0.05$ .

SUPPLEMENTARY DATA

**Supplementary Table 5. Sensitivity analyses of morningness-eveningness preference association with adjusted odds of type 2 diabetes ( $n = 298,953$ ) and adjusted mean difference in HbA1c (in mmol/mol;  $n = 272,220$ ) with each additional copy of the *MTNR1B* G in UK Biobank participants of European descent regardless of employment status.**

	Type 2 diabetes			HbA1c (mmol/mol)		
	Type 2 diabetes cases /controls	OR [95% CI]	$P_{int}$	<i>N</i>	Beta [95% CI]	$P_{int}$
<b>Overall</b>	12,324 /286,629	1.10 [1.07-1.14]	0.17	272,220	0.24 [0.22-0.26]	0.83
<b>Definite morning</b>	3,490 /75,909	1.14 [1.08-1.2]		72,013	0.26 [0.22-0.30]	
<b>More morning than evening</b>	4,006 /104,746	1.11 [1.05-1.16]		99,552	0.22 [0.18-0.25]	
<b>More evening than morning</b>	3,455 /81,559	1.07 [1.02-1.13]		77,517	0.23 [0.19-0.27]	
<b>Definite evening</b>	1,373 /24,415	1.09 [0.99-1.19]		23,138	0.27 [0.20-0.35]	

**Legend:** Association results are adjusted odds ratios [95% confidence interval] of type 2 diabetes per each additional copy of the *MTNR1B* G risk-allele or adjusted betas [95% confidence interval] describing differences in HbA1c in mmol/mol per each additional copy of the *MTNR1B* G risk-allele across categories of morningness-eveningness preference. Association analyses are adjusted for age, sex, BMI, genotyping array and 10 principal components of ancestry.  $P_{int}$  is log likelihood ratio test comparing models with and without cross-product interaction terms (*MTNR1B* and morningness-eveningness preference) including main effect terms in logistic or linear regression models adjusted for the aforementioned covariates.

SUPPLEMENTARY DATA

**Supplementary Table 6. Characteristics of subset of UK Biobank participants of European descent with up to 7-day actigraphy information.**

	Employed Participants	All Participants
<i>N</i>	38,701	82,923
Age, years	56.4	62.5
Sex, % male	41.90	43.9
BMI, kg/m <sup>2</sup>	26.5	26.7
Type 2 diabetes, <i>n</i> cases (%)	555 (1.4)	2,165 (2.6)
Sleep midpoint, clock time	26.91	26.99
Townsend Index*	-1.64	-1.80
People in Household, <i>n</i> (%)		
Husband, Wife or Partner	29,166 (75.4)	63,354 (76.4)
Son and/or Daughter	21,163 (54.7)	29,413 (35.5)
Mother and/or Father	149 (0.4)	287 (0.3)
Grandchild	809 (2.1)	1,285 (1.5)
Other	7 (0.02)	10 (0.01)

Data are mean (SD), median (interquartile range), or percentages.

\*Positive values of the index will indicate areas with high material deprivation, whereas those with negative values will indicate relative affluence.

SUPPLEMENTARY DATA

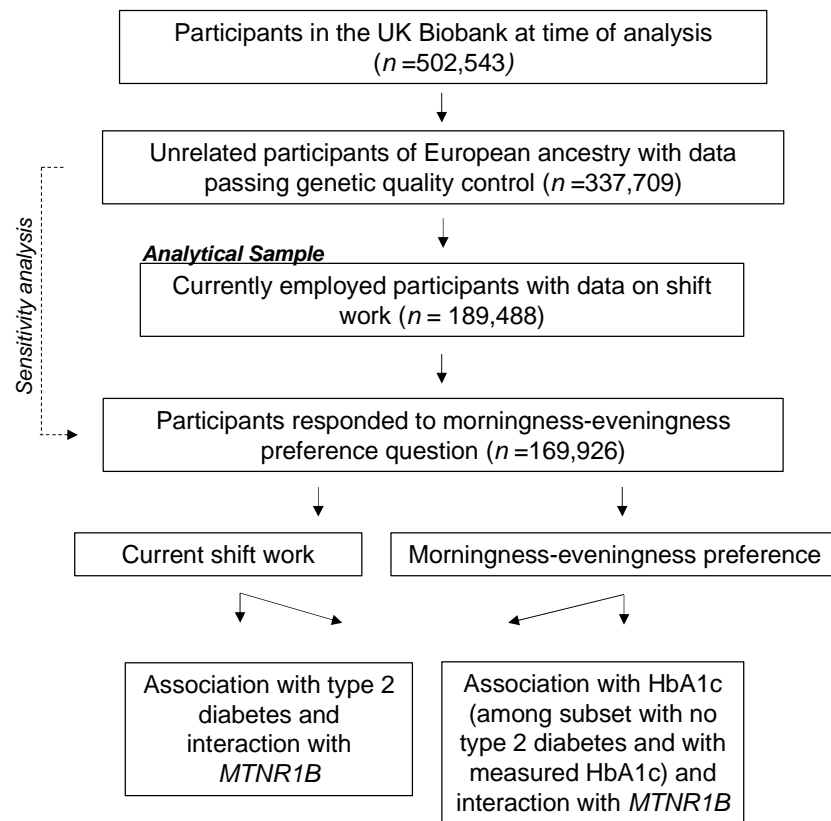
**Supplementary Table 7. Associations between quartiles of accelerometer-derived sleep midpoint ( $n = 82,923$ ) with adjusted odds of type 2 diabetes in UK Biobank participants of European descent regardless of employment status.**

	Type 2 diabetes cases /controls	Sex- and age-adjusted OR [95% CI]	Sex- and age- and household status- adjusted OR [95% CI]
<b>Sleep midpoint (Q1)</b>	590 / 20,141	<i>reference</i>	<i>reference</i>
<b>Sleep midpoint (Q2)</b>	474 / 20,258	<b>0.79 [0.72-0.88]</b>	<b>0.80 [0.73-0.89]</b>
<b>Sleep midpoint (Q3)</b>	481 / 20,249	<b>0.79 [0.71-0.87]</b>	<b>0.80 [0.72-0.88]</b>
<b>Sleep midpoint (Q4)</b>	620 / 20,110	0.99 [0.88-1.11]	1.00 [0.89-1.12]

**Legend:** Prevalent type 2 diabetes associations are sex- and age- adjusted odds ratios [95% confidence interval] and bold  $P < 0.05$ . In sensitivity analysis, associations were further adjusted for people residing in the household with the participant (household status).

SUPPLEMENTARY DATA

Supplementary Figure 1. Analysis workflow.





SUPPLEMENTARY DATA

**Supplementary Figure 2. Adjusted odds ratios (OR) of type 2 diabetes with each additional copy of the *MTNR1B* G risk-allele in the overall sample and across categories of morningness-eveningness preference in employed UK Biobank participants of European descent ( $n = 169,926$ ).**

