**Online-Only Supplemental Material**

**Cardiovascular risk reduction with liraglutide: an exploratory mediation analysis of the LEADER trial**

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# **Supplementary Figure S1. Probability of no MACE occurring in patients treated with liraglutide and placebo, and with adjustment for HbA1c as a candidate mediator, and body weight, UACR, systolic blood pressure, LDL, confirmed hypoglycemic episodes, insulin use and sulfonylurea use as confounders, using the Vansteelandt method.**

HbA1c, glycated hemoglobin; MACE, major adverse cardiovascular event; UACR, urinary albumin-to-creatinine ratio

# **Supplementary Figure S2. On-treatment analysis for the probability of no MACE in patients treated with liraglutide and placebo, and with adjustment for HbA1c as a candidate mediator using the Vansteelandt method.**

HbA1c, glycated hemoglobin; MACE, major adverse cardiovascular event

# **Supplementary Figure S3. Analysis of time to first MACE with reversed adjustment for HbA1c using the Vansteelandt method.**

HbA1c, glycated hemoglobin; MACE, major adverse cardiovascular event

# **Supplementary Figure S4. Probability of no (A) all-cause death, and (B) CV death occurring in patients treated with liraglutide and placebo, and with adjustment for HbA1c as a candidate mediator using the Vansteelandt method.**



CV, cardiovascular; HbA1c, glycated hemoglobin

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