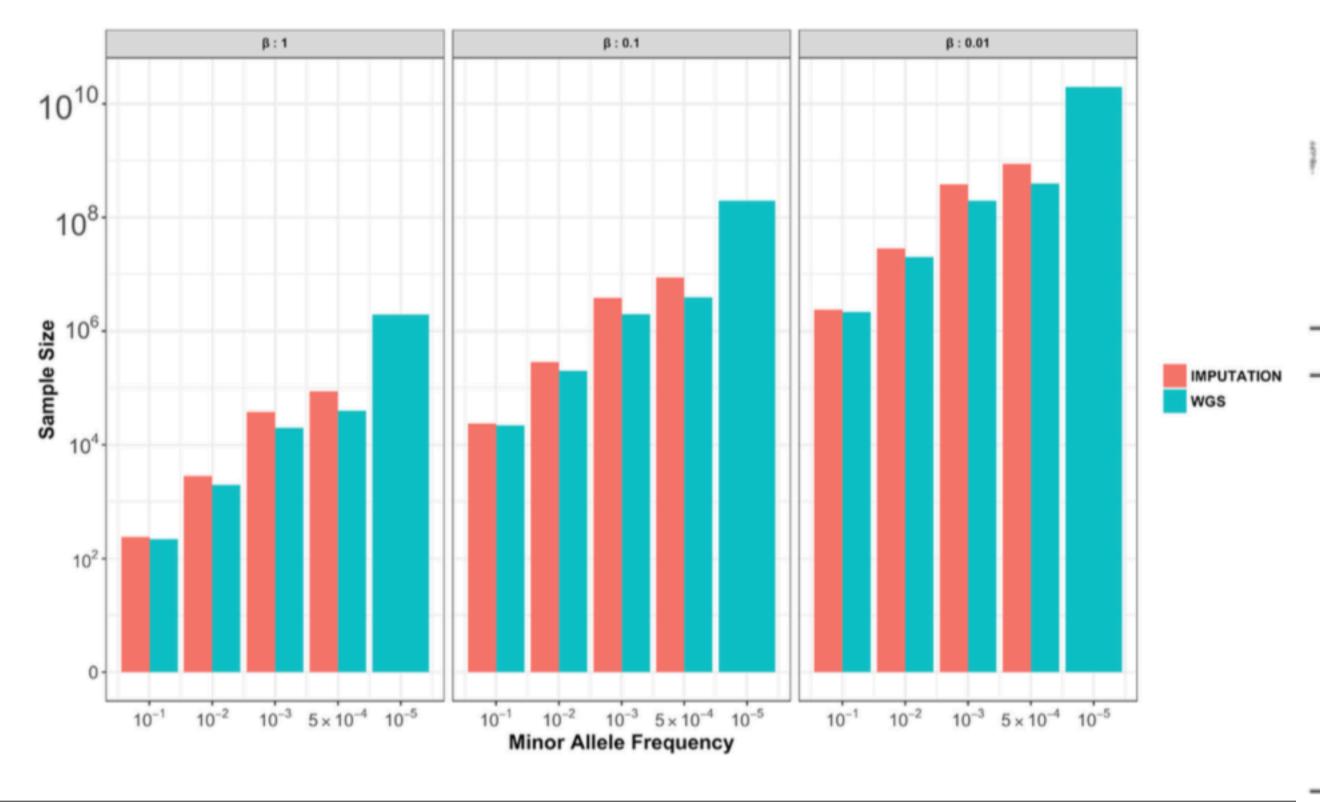
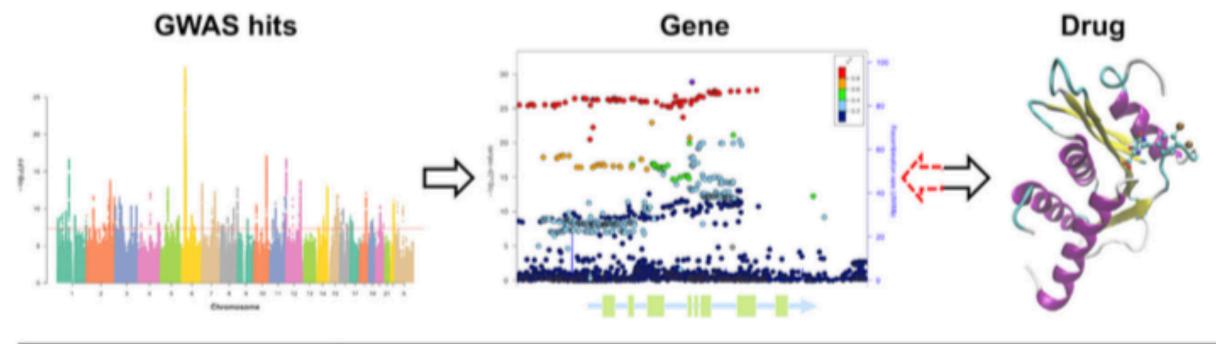
GWAS



- Quanto menor a associação, mais amostras necessárias.
- Quanto menor a MAF (Minor Alelle Frequency), mais amostras necessárias.
- Permite identificar genes alvo.
- Desenvolvimento de drogas.





Trait	Gene with GWAS hits	Known or candidate drug		
Type 2 Diabetes	SLC30A8/KCNJ11	ZnT-8 antagonists/Glyburide		
Rheumatoid Arthritis	PADI4/IL6R	BB-CI-amidine/Tocilizumab		
Ankylosing Spondylitis(AS)	TNFR1/PTGER4/TYK2	TNF- inhibitors/NSAIDs/fostamatinib		
Psoriasis(Ps)	IL23A	Risankizumab		
Osteoporosis	RANKL/ESR1	Denosumab/Raloxifene and HRT		
Schizophrenia	DRD2	Anti-psychotics		
LDL cholesterol	HMGCR	Pravastatin		
AS, Ps, Psoriatic Arthritis	IL12B	Ustekinumab		

Diabetes tipo II

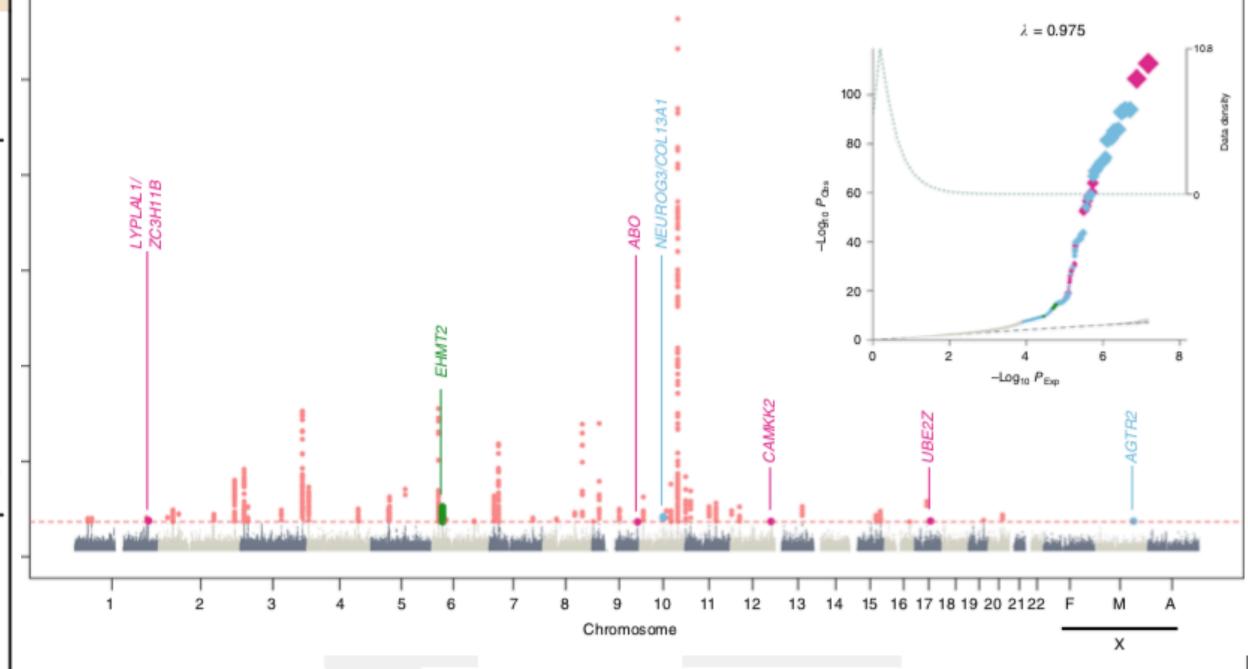
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- Background genético
- Influência ambiental e comportamental.

Table 1 Novel T2D-associated loci							
			OR (95% CI) <i>P</i> -value				
Novel Locus	Chr	rsIDRisk Allele	Stage1 Discovery Meta- analysis	Stage2 Replication Meta-analysis	Stage1 + Stage2 Combined Meta- analysis	MAF	
LYPLAL1/ZC3H11B (1q41)	1	rs2820443-T	1.08 (1.04-1.13) 2.94 × 10 ⁻⁴ a	1.06 (1.03-1.09) 2.10 × 10 ^{-5 b}	1.07 (1.04-1.09) 2.56 × 10 ^{-8 c}	0.28	
EHMT2 (6p21.33-p21.32)	6	rs115884658-A	1.34 (1.18-1.53) 1.00 × 10 ⁻⁵ a	1.17 (1.09-1.26) 2.90 × 10 ^{-6 c, d}	1.21 (1.14-1.29) 3.00 × 10 ^{-10 c}	0.02	
ABO (9q34.2)	9	rs505922-C	1.07 (1.03-1.11) 6.93 × 10 ⁻⁴ a	1.06 (1.03-1.09) 1.90 × 10 ^{-5 b}	1.06 (1.04-1.09) 4.94 × 10 ^{-8 c}	0.34	
NEUROG3 (10q22.1)	10	rs2642587-G	1.12 (1.08-1.16) 8.45 × 10 ⁻⁹ e	-	-	0.22	
CAMKK2 (12q24.31)	12	rs3794205-G	1.09 (1.05-1.14) 4.18 × 10 ⁻⁵ a	1.06 (1.03-1.09) 1.60 × 10 ^{-4 b}	1.07 (1.04-1.10) 4.11 × 10 ^{-8 c}	0.32	
CALCOCO2/ATP5G1/ UBE2Z/SNF8/GIP (17q21.32)	17	rs12453394-A	1.08 (1.04-1.12) 7.86 × 10 ⁻⁵ a	1.07 (1.03-1.11) 9.60 × 10 ^{-5 b}	1.07 (1.05-1.10) 3.23 × 10 ^{-8 c}	0.47	
AGTR2 (Xq23)	Х	rs146662075-T	3.09 (2.06-4.60) 3.24 × 10 ^{-8 f}	1.57 (1.19-2.07) 1.42 × 10 ^{-3 g}	1.95 (1.56-2.45) 7.85 × 10 ⁻⁹	0.008	

Chr chromosome, OR odds ratio, MAF minor allele frequency

⁸Replication Men Cohort SIGMA UK10K imputation + InterAct + Danish Cohort (case control and follow-up) + Partners Biobank + UK Biobank (18,370 cases and 88,283 controls older than 55 years and OGTT > 7.8 mmol I⁻¹, when available)



almputed based public GWAS discovery meta-analysis (NuGENE + GERA cohort, 7,522 cases and 50,446 controls)

^bTransancestry DIA GRAM Consortium (26,488 cases and 83,964 controls)^cMeta P-value estimated using a weighted Z-score method due to unavailable SE information from Stage 2 replication cohorts Diabetes Genetic Portal (Exome-Chip + Exome Sequencing, 35,789 cases and 56,738 controls)^cFull imputed based public GWAS meta-analysis (NuGENE + GERA cohort + GENEVA + FUSION + WTCCC, 12,931 cases and 57,196 controls)

^f70KforT2D Men Cohort (GERA cohort + GENEVA + FUSION, 5,277 cases and 15,702 controls older than 55 years)