

Table 1.

## Summary of Included Articles

First Author, Year	Prospective or Retrospective Identification of TRD	Study Region	Study Design	Mean Depression Score	Threshold Depression Score	Total No. of Participants	Reported Risk Factor/Predictor Categories
Anttila 2007 <sup>14</sup>	Retrospective	Finland	Case-control	NA	NA	511	Genetic
Balestri 2016 <sup>a, 15</sup>	Prospective	Europe	Cohort	<b>MADRS, mean (SD):</b> Total = 31.30 (6.18) Resistant = 32.37 (6.37) Non-resistant = 30.89 (6.07)	MADRS > 22	355	Mental
Brenner 2020 <sup>16</sup>	Retrospective	Sweden	Case-control	NA	NA	121,669	Mental
Carvalho 2017 <sup>17</sup>	Prospective	Portugal	Cohort	<b>BDI, mean (SD) [range]:</b> 27.81 (7.40) [20.0–48.0]	BDI > 20	80	Genetic
Cepeda 2018 <sup>18</sup>	Prospective/retrospective	United States	Cohort	NA	NA	230,801	Mental, physical, demographic
Döme 2021 <sup>19</sup>	Retrospective	Hungary	Cohort	NA	NA	99,531	Mental, physical, demographic
de Diego-Adelño 2014 <sup>20</sup>	Retrospective	Spain	Case-control	<b>HDRS, mean (SD):</b> Resistant = 20.8 (4.9) Remitted = 2.1 (0.9) First-episode = 14.6 (7.5)	HDRS > 14	18	Mental, demographic, imaging
Dudek 2010 <sup>21</sup>	Retrospective	Poland	Case-control	NA	HDRS < 18	1,051	Mental, demographic
Fabbri 2015 <sup>a,b, 22</sup>	Retrospective	Other: Europe and United States	Cohort	NA	European = no cutoff Italian sample = ≥ 13 HDRS STAR*D = ≥ 14 HDRS	European MDD = 287, European BP = 86, Italian = 90, STAR*D = 1,892	Genetic
Fabbri 2017 <sup>a,b, 23</sup>	Retrospective	Europe*	Cohort	<b>HDRS, mean (SD):</b> 22.30 (5.29) <b>MADRS, mean (SD):</b> 30.55 (6.14)	MADRS ≥ 22	220	Genetic
Fabbri 2018 <sup>a,b, 24</sup>	Retrospective	Europe*	Cohort	NA	MADRS ≥ 22	NA	Genetic
Fabbri 2019 <sup>a,b, 25</sup>	Retrospective	UK, Europe,* and United States	Cohort	NA	MADRS ≥ 22	3,225	Genetic
Fabbri 2020 <sup>a,b, 26</sup>	Retrospective	United States and Europe*	Cohort	<b>MADRS, mean (SD):</b> 33.85 (7.69)	MADRS ≥ 22	1209	
Fabbri 2021 <sup>27</sup>	Retrospective	UK	Cohort	NA	MADRS ≥ 22	EXCEED = 8,926 of which 1,271 (14.24%) MDD; UKB = 230,096 of which 19,979 (8.68%) MDD	Mental, physical, demographic, personality, genetic
Gronemann 2018 <sup>28</sup>	Prospective/retrospective	Denmark	Cohort	<b>ICD severity codes:</b> Mild (10.2%) Moderate (16.8%) Severe (24.8%)	NA	222,480	Mental
Gronemann 2020 <sup>29</sup>	Prospective/retrospective	Denmark	Cohort	<b>ICD severity codes:</b> Mild (11.2%) Moderate (19.5%) Severe (28.1%)	NA	194,074	Mental, physical, demographic
Guo 2012 <sup>30</sup>	Retrospective	China	Case-control	<b>HDRS, mean (SD):</b> Resistant = 24.52 (4.17) Responsive = 25.89 (6.26)	HDRS ≥ 18	18	Mental, imaging
Guo 2012 <sup>31</sup>	Retrospective	China	Case-control	<b>HDRD, mean (SD):</b> Resistant = 23.89 (3.69) Responsive = 25.58 (6.32)	HDRS ≥ 18	23	Mental, imaging
He 2016 <sup>32</sup>	Retrospective	China	Case-control	<b>HDRS, mean (SD):</b> Resistant = 23.94 (3.79) Responsive = 25.59 (6.31)	HDRS ≥ 18	17	Mental, imaging
Huang 2018 <sup>33</sup>	Retrospective	Taiwan	Case-control	<b>HDRS (17-item), mean (SD):</b> TRD = 19.6 (10.3) Non-TRD MDD = 10.86 (7.2)	NA	68	Mental
Huang 2020 <sup>34</sup>	Prospective/retrospective	Taiwan	Cohort	<b>ICD severity codes:</b> Severe 45.9%	NA	31,422	Mental, physical, demographic, other
Huang 2023 <sup>35</sup>	Retrospective	China	Case-control	NA	NA	1,151	Mental, physical, demographic, other
Jeng 2018 <sup>36</sup>	Retrospective	Taiwan	Cohort	NA	NA	127,194	Physical
Kautzky 2017 <sup>a, 37</sup>	Retrospective	Europe*	Cohort	NA	MADRS ≥ 22	480	Mental, physical, demographic
Kautzky 2018 <sup>a, 38</sup>	Prospective	Europe*	Cohort	NA	MADRS ≥ 22	552	Mental, other
Kautzky 2019 <sup>a, 39</sup>	Prospective	Europe*	Cross-sectional	NA	MADRS ≥ 22	916	Mental
Kraus 2023 <sup>40</sup>	Retrospective	Europe*	Cohort	NA	NA	892	Physical,
Lähteenvirta 2022 <sup>41</sup>	Prospective/retrospective	Finland	Cohort	<b>ICD severity codes:</b> Mild (18.8%) Moderate (26.4%) Severe (17.9%) Unknown (37%)	NA	177,144	Mental, demographic
López-Solà 2020 <sup>42</sup>	Prospective	Spain	Case-control	<b>HDRS, mean (SD):</b> Resistant = 17.64 (8.57) Non-resistant = 13.67 (8.65)	NA	229	Mental, other
Lauden 2021 <sup>43</sup>	Retrospective	Israel	Cross-sectional	NA	NA	6,270	Physical
Li 2013 <sup>44</sup>	Prospective	China	Cohort	NA	NA	948	Genetic
Li 2019 <sup>45</sup>	Retrospective	Multinational	Cross-sectional	<b>HDRS, mean (SD):</b> TRD = 23.8 (2.6) Non-TRD MDD = 24.1 (2.6)	NA	21,920	Mental, demographic
Liberman 2020 <sup>46</sup>	Prospective/retrospective	United States	Cohort	NA	NA	35,246	Mental, physical, demographic
Liu 2021 <sup>47</sup>	Retrospective	United States	Cross-sectional	NA	NA	Humana: 2,987,264 Optum: 5,532,986 Total: 8,520,250	Demographic, other
Maes 2009 <sup>48</sup>	Prospective	Belgium	Cohort	NA	NA	57	Biological
Mandelli 2016 <sup>a, 49</sup>	Retrospective	Europe*	Cross-sectional	<b>MADRS, mean (SD):</b> Blue-collar = 23 (11.6) Self-employed = 23.1 (12.2) White-collar = 24.3 (10.6) Manager = 24.9 (8.2)	NA	647	Mental, demographic
Milanesi 2015 <sup>50</sup>	Retrospective	Italy	Non-randomized experimental	NA	NA	627	Genetic
Minelli 2015 <sup>51</sup>	Retrospective	Italy	Cross-sectional	NA	NA	1,088	Genetic
Nie 2018 <sup>b, 52</sup>	Prospective	United States	Cohort	NA	NA	NA	Mental
O'Dushlaine 2014 <sup>b, 53</sup>	Retrospective	United States	Cohort	NA	NA	1,263	Genetic
Park 2021 <sup>54</sup>	Prospective/retrospective	Korea	Cohort	NA	NA	15,887	Mental, demographic
Perlis 2013 <sup>b, 55</sup>	Prospective	United States	Cohort	NA	NA	2,555	Mental, demographic, other
Petersen 2002 <sup>56</sup>	Retrospective	United States	Cross-sectional	NA	HDRS ≥ 18	158	Mental
Qiao 2018 <sup>82</sup>	Retrospective	China	Cohort	<b>HDRS, mean (SD):</b> TRD = 28.67 (5.71) Non-TRD MDD = 29.95 (5.33)	HDRS ≥ 17	206	Mental, biological
Sagud 2013 <sup>57</sup>	Retrospective	Croatia	Cross-sectional	<b>HDRS, mean (SD):</b> TRD = 27.7 (6.3) Non-TRD MDD = 26.1 (5.9)	NA	203	Mental, biological
Santos 2016 <sup>58</sup>	Prospective	Portugal	Non-randomized experimental	<b>BDI, mean (SD):</b> 27.81 (7.40)	BDI > 20 threshold	80	Genetic
Santos 2015 <sup>59</sup>	Retrospective	Portugal	Cohort	<b>BDI, mean (SD):</b> 27.81 (7.40)	BDI > 20 threshold	80	Genetic
Scherrer 2016 <sup>60</sup>	Prospective/retrospective	United States	Cohort	NA	NA	6,169	Physical, other
Sherbourne 2004 <sup>61</sup>	Prospective	United States	Cohort	NA	NA	542	Mental, demographic
Souery 2007 <sup>a, 62</sup>	Retrospective	Europe*	Cross-sectional	NA	HDRS ≤ 17	702	Mental
Sternat 2018 <sup>63</sup>	Retrospective	Canada	Cross-sectional	NA	NA	160	Mental, demographic
Takahashi 2013 <sup>64</sup>	Retrospective	Japan	Non-randomized experimental	<b>HDRS, mean (SD):</b> 18.31 (4.04)	HDRS ≥ 14	128	Personality
Takahashi 2013 <sup>65</sup>	Retrospective	Japan	Non-randomized experimental	<b>HDRS, mean (SD):</b> 18.31 (4.04)	HDRS ≥ 14	240	Personality
Wang 2022 <sup>66</sup>	Retrospective	China	Non-randomized experimental	<b>HDRS, mean (SD):</b> TRD = 29.40 (5.02) Non-TRD MDD = 22.41 (3.6)	NA	261	Genetic
Wu 2022 <sup>67</sup>	Retrospective	China	Case-control	<b>HDRS, mean (SD):</b> TRD = 32.87 (0.55)	NA	90	Biological
Yamasaki 2020 <sup>68</sup>	Retrospective	Japan	Non-randomized experimental	<b>HDRS, mean (SEM):</b> TRD = 18.5 (18.1–18.9) Non-TRD MDD = 16.7 (16.2–17.2)	NA	42	Biological
Zhang 2014 <sup>69</sup>	Retrospective	China	Non-randomized experimental	<b>HDRS, mean (SD):</b> TRD = 24.0 (2.7) Non-TRD MDD = 24.1 (2.6)	HDRS ≥ 17	1,569	Genetic

<sup>a</sup>Article from the European Group for the Study of Resistant Depression (GSRD).<sup>b</sup>Article from STAR\*D.

\*Patient population is from the GSRD European cohort, which includes individuals from Belgium, France, Italy, Israel, Austria, and Greece.

Abbreviations: BDI = Beck Depression Inventory, HDRS = Hamilton Depression Rating Scale, ICD = International Classification of Diseases, MADRS = Montgomery-Asberg Depression Rating Scale, NA = not applicable, SEM = standard error of the mean, STAR\*D = Sequenced Treatment Alternatives to Relieve Depression, TRD = treatment-resistant depression, UKB = UK Biobank.