

## Supplementary Online Content

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This supplementary material has been provided by the authors to give readers additional information about their work.

**eTable 1: General characteristics of all included studies (n=132)**

NMLs = non-melanocytic lesions, nr= not reported, SVM= support vector machine, ANN= artificial neural network, \*histopathology was only used for suspicious lesions

STUDY ID	METHOD	TEST SET SOURCE	TOTAL DATA SET SIZE	# OF MELANOMA	NMLS INCLUDED	CLASSIFIER	INDEPENDENT TEST SET	GOLD STANDARD FOR THE TEST SET	INCLUDED IN QUANTITATIVE ANALYSIS
ABUZAGHLEH 2015 <sup>1</sup>	computer vision	public	200	40	no	SVM	no	histopathology*, dermatoscopy	
AFIFI 2017 <sup>2</sup>	computer vision	public	356	168	nr	SVM	no	nr	
AMOABEDINI 2018 <sup>3</sup>	computer vision	public	76	38	no	threshold	no	nr	Yes
ASCIERTO 2010 <sup>4</sup>	hardware-based	proprietary	54	12	no	nr	yes	histopathology	Yes
BARATA 2015A <sup>5</sup>	computer vision	public	482	241	nr	ensemble classifier	no	expert consensus	
BARATA 2015B <sup>6</sup>	computer vision	public	200	40	no	random forests	no	histopathology*, dermatoscopy	Yes
BARROS MENDES 2018 <sup>7</sup>	deep learning	public	5286	374	yes	ANN	no	nr	
BARZEGARI 2005 <sup>8</sup>	computer vision	proprietary	122	6	yes	ANN	no	histopathology	Yes
BAUR 2018 <sup>9</sup>	deep learning	public	2000	2000	yes	ANN	no	histopathology*, expert consensus	
BHATTACHARYA 2017 <sup>10</sup>	deep learning	public	275	128	no	SVM	no	nr	
BI 2017 <sup>11</sup>	deep learning	public	4350	nr	nr	ANN	yes	histopathology*, expert consensus	Yes
BISSOTO 2018 <sup>12</sup>	deep learning	public	nr	nr	yes	ANN	yes	nr	
BONO 2002 <sup>13</sup>	hardware-based	proprietary	313	66	yes	discriminant analysis	yes	histopathology	
CARCAGNI 2018 <sup>14</sup>	deep learning	public	10015	1113	yes	ANN	yes	histopathology*, follow-up, confocal microscopy, expert consensus	
CELEBI 2007 <sup>15</sup>	computer vision	public	596	88	yes	SVM	no	histopathology*, follow-up	
CELEBI 2008 <sup>16</sup>	computer vision	public	545	186	yes	decision tree	no	histopathology*, follow-up	
CHAKRAVATI 2015 <sup>17</sup>	computer vision	public	456	251	nr	ANN	no	histopathology*, clinical evaluation	

<b>CHANG H 2017<sup>18</sup></b>	deep learning	public	2750	521	yes	ANN	yes	histopathology*, expert consensus	Yes
<b>CHANG WY 2013<sup>19</sup></b>	computer vision	proprietary	769	8	yes	SVM	no	histopathology	
<b>CHANG WY 2015<sup>20</sup></b>	computer vision	proprietary	347	7	yes	SVM	yes	histopathology	
<b>CHOU 2017<sup>21</sup></b>	deep learning	public	200	40	no	SVM	no	histopathology*, dermatoscopy	
<b>CODELLA 2016<sup>22</sup></b>	deep learning	public	1279	248	no	SVM	yes	histopathology*, expert consensus, nr for part of the lesions	Yes
<b>CODELLA 2018<sup>23</sup></b>	deep learning	public	2750	521	yes	kNN	no	histopathology*, expert consensus	
<b>DEL ROSARIO 2018<sup>24</sup></b>	computer vision	proprietary	1076	71	yes	threshold	yes	histopathology	Yes
<b>DING 2015<sup>25</sup></b>	hardware-based	proprietary	46	12	yes	SVM	no	nr	Yes
<b>DO 2014<sup>26</sup></b>	computer vision	proprietary	81	29	nr	SVM	no	nr	
<b>DO 2017<sup>27</sup></b>	computer vision	proprietary	184	67	no	SVM	no	histopathology*, expert consensus	Yes
<b>DREISEITL 2009<sup>28</sup></b>	computer vision	proprietary	3827	31	no	ANN	yes	histopathology*, follow-up	Yes
<b>ESTEVA 2017<sup>29</sup></b>	deep learning	proprietary	129450	nr	yes	ANN	no	test set histopathology, rest unspecified (dermatologist-labelled)	
<b>FERRI 2010<sup>30</sup></b>	computer vision	proprietary	977	50	no	SVM	no	histopathology*, follow-up	Yes
<b>FINK 2016<sup>31</sup></b>	hardware-based	proprietary	360	3	no	threshold	yes	histopathology*, dermoscopy follow-up	Yes
<b>FORNACIALI 2016<sup>32</sup></b>	deep learning	public	1052	267	nr	SVM	no	histopathology*, clinical evaluation	
<b>FORSCHNER 2018<sup>33</sup></b>	hardware-based	proprietary	529	101	yes	ANN	no	histopathology	Yes
<b>FRIEDMAN 2008<sup>34</sup></b>	hardware-based	proprietary	99	49	no	threshold	no	histopathology	
<b>GAREAU 2017<sup>35</sup></b>	computer vision	proprietary	120	60	no	decision tree	no	histopathology	Yes
<b>GARNAVI 2012<sup>36</sup></b>	computer vision	proprietary	289	114	nr	random forests	no	nr	Yes
<b>GAUTAM 2018<sup>37</sup></b>	computer vision	public	294	138	nr	random forests	no	nr	Yes

<b>GILMORE 2009</b> <sup>38</sup>	computer vision	proprietary	312	102	no	nr	no	histopathology*, dermatoscopy	
<b>GILMORE 2010</b> <sup>39</sup>	computer vision	proprietary	199	101	no	SVM	no	histopathology	Yes
<b>GILMORE 2018</b> <sup>40</sup>	computer vision	proprietary	250	85	no	SVM	no	histopathology*, dermoscopy evaluation	
<b>GONZALEZ DIAZ 2017</b> <sup>41</sup>	deep learning	public	2750	521	yes	ANN	yes	histopathology*, expert consensus	Yes
<b>GOYAL 2018</b> <sup>42</sup>	deep learning	public	10015	1113	yes	ANN	yes	histopathology*, follow-up, confocal microscopy, expert consensus	
<b>GUO S 2017</b> <sup>43</sup>	deep learning	public	2750	521	yes	ensemble classifier	yes	histopathology*, expert consensus	Yes
<b>GUO Y 2018</b> <sup>44</sup>	deep learning	public	10015	1113	yes	ANN	yes	histopathology*, follow-up, confocal microscopy, expert consensus	
<b>HAENSLE 2018</b> <sup>45</sup>	deep learning	proprietary	>100000	nr	no	ANN	no	histopathology*, dermoscopy follow-up	Yes
<b>HAIDER 2014</b> <sup>46</sup>	computer vision	public	206	119	nr	bayesian	no	histopathology	Yes
<b>HARANGI 2017</b> <sup>47</sup>	deep learning	public	2750	521	yes	ANN	yes	histopathology*, expert consensus	Yes
<b>HARANGI 2018</b> <sup>48</sup>	deep learning	public	2750	521	yes	ensemble on ANNs	no	histopathology*, expert consensus	
<b>HARDIE 2018</b> <sup>49</sup>	computer vision	public	10015	1113	yes	SVM	no	histopathology*, follow-up, confocal microscopy, expert consensus	Yes
<b>HAR-SHAI 2005</b> <sup>50</sup>	hardware-based	proprietary	449	69	yes	threshold	yes	histopathology	Yes
<b>HAUSCHILD 2014</b> <sup>51</sup>	hardware-based	proprietary	130	65	yes	threshold	yes	histopathology	
<b>HUNGER 2012</b> <sup>52</sup>	hardware-based	proprietary	45	12	no	threshold	yes	histopathology*, follow-up	
<b>JAWOREK-KORJAKOWSKA 2016A</b> <sup>53</sup>	computer vision	public	300	60	no	SVM	no	histopathology	Yes
<b>JAWOREK-KORJAKOWSKA 2016B</b> <sup>54</sup>	computer vision	public	200	70	nr	SVM	no	histopathology	
<b>JIJI 2017</b> <sup>55</sup>	computer vision	public	2750	521	yes	SVM	yes	histopathology*, expert consensus	Yes
<b>KAUR 2015</b> <sup>56</sup>	computer vision	proprietary	192	114	no	linear regression	no	histopathology*, follow-up	

<b>KAWAHARA 2018<sup>57</sup></b>	deep learning	proprietary	1011	252	yes	ANN	no	nr	Yes
<b>KHAN 2018<sup>58</sup></b>	computer vision	public	4752	1057	yes	SVM	no	histopathology*, follow-up, expert consensus, unknown for part of the lesions	Yes
<b>KITADA 2018<sup>59</sup></b>	deep learning	public	10015	1113	yes	ANN	yes	histopathology*, follow-up, confocal microscopy, expert consensus	
<b>KOSTOPOULOS 2017<sup>60</sup></b>	computer vision	public	399	69	no	ANN	no	nr	Yes
<b>LAI 2018<sup>61</sup></b>	deep learning	public	2000	374	yes	ANN	no	histopathology*, expert consensus	
<b>LEE 2018<sup>62</sup></b>	deep learning	public	10015	1113	yes	ANN	yes	histopathology*, follow-up, confocal microscopy, expert consensus	
<b>LI K 2018<sup>63</sup></b>	deep learning	public	10015	1113	yes	ANN	no	histopathology*, follow-up, confocal microscopy, expert consensus	Yes
<b>LI L 2014<sup>64</sup></b>	hardware-based	public	187	19	no	bayesian	no	histopathology	
<b>LI Y 2017<sup>65</sup></b>	deep learning	public	2750	521	yes	ANN	yes	histopathology*, expert consensus	Yes
<b>LINGALA 2014<sup>66</sup></b>	computer vision	proprietary	888	195	no	logistic regression model	no	nr	
<b>LIU 2015<sup>67</sup></b>	computer vision	public	258	76	yes	SVM	no	histopathology*, clinical evaluation	Yes
<b>LU 2018<sup>68</sup></b>	deep learning	public	10015	1113	yes	ANN	no	histopathology*, follow-up, confocal microscopy, expert consensus	
<b>MAGLIOGIANNIS 2004<sup>69</sup></b>	computer vision	proprietary	17	7	no	SVM	no	histopathology	
<b>MAGLIOGIANNIS 2015<sup>70</sup></b>	computer vision	proprietary	208	100	no	SVM	no	histopathology	Yes
<b>MAHBOD 2017<sup>71</sup></b>	deep learning	public	2750	521	yes	SVM	yes	histopathology*, expert consensus	Yes
<b>MAITNER 2018<sup>72</sup></b>	deep learning	public	10015	1113	yes	ANN	yes	histopathology*, follow-up, confocal microscopy, expert consensus	
<b>MALVEHY 2014<sup>73</sup></b>	hardware-based	proprietary	2416	265	yes	threshold	yes	histopathology	Yes
<b>MANOUSAKI 2006<sup>74</sup></b>	computer vision	proprietary	132	23	no	logistic regression model	no	histopathology	Yes
<b>MARCHETTI 2018<sup>75</sup></b>	deep learning	public	379	75	no	threshold	no	histopathology*, expert consensus	Yes
<b>MARQUES 2012<sup>76</sup></b>	computer vision	proprietary	163	17	no	nr	no	nr	Yes

<b>MATSUNAGA 2017<sup>77</sup></b>	deep learning	public	4194	930	yes	ANN	yes	histopathology*, expert consensus	Yes
<b>MENEGOLA 2016<sup>78</sup></b>	deep learning	public	nr	nr	yes	SVM	no	histopathology	
<b>MENEGOLA 2017A<sup>79</sup></b>	deep learning	public	nr	nr	yes	SVM	no	histopathology*, expert consensus, nr for part of the lesions	Yes
<b>MENEGOLA 2017B<sup>80</sup></b>	deep learning	public	18000	nr	yes	SVM	yes	nr	Yes
<b>MESSADI 2009<sup>81</sup></b>	computer vision	proprietary	180	72	no	ANN	no	clinical evaluation	
<b>MIRUNALINI 2017<sup>82</sup></b>	deep learning	public	2750	521	yes	ANN	yes	histopathology*, expert consensus	Yes
<b>MOLLERSEN 2015<sup>83</sup></b>	computer vision	proprietary	877	45	yes	nr	yes	histopathology	Yes
<b>MONHEIT 2011<sup>84</sup></b>	hardware-based	proprietary	1632	127	yes	threshold	yes	histopathology	
<b>MUNIA 2017<sup>85</sup></b>	computer vision	public	170	70	no	SVM	no	nr	Yes
<b>MURPHREE 2017<sup>86</sup></b>	deep learning	public	2750	521	yes	ANN	yes	histopathology*, expert consensus	Yes
<b>NADER VASCONCELOS 2017<sup>87</sup></b>	deep learning	public	2750	521	yes	ANN	yes	histopathology*, expert consensus	Yes
<b>NANNI 2018<sup>88</sup></b>	deep learning	public	10015	1113	yes	ANN	yes	histopathology*, follow-up, confocal microscopy, expert consensus	
<b>NASIRI 2018<sup>89</sup></b>	deep learning	public	10015	1113	yes	ANN	yes	histopathology*, follow-up, confocal microscopy, expert consensus	
<b>NASR-ESFAHANI 2016<sup>90</sup></b>	deep learning	public	170	70	no	ANN	yes	nr	
<b>NICOLAS 2014<sup>91</sup></b>	computer vision	proprietary	150	nr	yes	kNN	no	histopathology	
<b>PAL 2018<sup>92</sup></b>	deep learning	public	10015	1113	yes	ANN	yes	histopathology*, follow-up, confocal microscopy, expert consensus	
<b>PATWARDHAN 2005<sup>93</sup></b>	hardware-based	proprietary	90	30	no	bell membership function	no	histopathology	Yes
<b>PELLACANI 2004<sup>94</sup></b>	computer vision	proprietary	339	113	no	discriminant analysis	no	histopathology	Yes
<b>PEREZ 2018<sup>95</sup></b>	deep learning	public	2750	512	yes	ANN	no	histopathology*, expert consensus	
<b>PICCOLO 2014<sup>96</sup></b>	computer vision	proprietary	165	33	no	threshold	no	histopathology	Yes

<b>POLAP 2018</b> <sup>97</sup>	computer vision	public	200	40	no	ANN	no	histopathology*, dermoscopy evaluation	Yes
<b>QUINZAN 2013</b> <sup>98</sup>	hardware-based	proprietary	32	7	no	qualified majority voting with 5 SVMs	no	histopathology	Yes
<b>RADHAKRISHNAN 2017</b> <sup>99</sup>	deep learning	public	1279	248	no	ANN	no	histopathology*, expert consensus, nr for part of the lesions	Yes
<b>RADHAKRISHNAN 2018</b> <sup>100</sup>	deep learning	public	2750	512	yes	ANN	no	histopathology*, expert consensus	
<b>RAMEZANI 2014</b> <sup>101</sup>	computer vision	public	282	133	nr	SVM	no	nr	Yes
<b>RAY 2018</b> <sup>102</sup>	deep learning	public	10015	1113	yes	deep forest	no	histopathology*, follow-up, confocal microscopy, expert consensus	
<b>REZVANTALAB 2018</b> <sup>103</sup>	deep learning	public	10135	1153	yes	ANN	no	histopathology*, follow-up, confocal microscopy, expert consensus, dermoscopy evaluation	Yes
<b>ROCHA 2017</b> <sup>104</sup>	hardware-based	proprietary	160	6	no	threshold	yes	histopathology*, follow-up	Yes
<b>RUBEGNI 2015</b> <sup>105</sup>	computer vision	proprietary	856	272	no	logistic regression model	no	histopathology	Yes
<b>RUIZ 2008</b> <sup>106</sup>	computer vision	public	110	nr	nr	voting system based on ANNs and a bayesian classifier	no	nr	
<b>SABBAGHI 2018</b> <sup>107</sup>	computer vision	proprietary	825	181	no	SVM	no	histopathology*, dermoscopy follow-up	
<b>SATHEESHA 2017</b> <sup>108</sup>	computer vision	public	263	nr	no	SVM	no	nr	
<b>SBONER 2003</b> <sup>109</sup>	computer vision	proprietary	152	42	no	ensemble classifier	no	histopathology	Yes
<b>SHAIFIEE 2017</b> <sup>110</sup>	deep learning	public	9152	567	yes	ANN	no	histopathology	
<b>SITU 2008</b> <sup>111</sup>	computer vision	proprietary	100	30	no	SVM	no	nr	
<b>SITU 2010</b> <sup>112</sup>	computer vision	proprietary	100	30	no	SVM	no	nr	Yes
<b>SITU 2011</b> <sup>113</sup>	computer vision	public	360	90	nr	SVM	no	histopathology	
<b>STANLEY 2007</b> <sup>114</sup>	computer vision	public	226	113	no	threshold	no	histopathology	Yes
<b>STOECKER 2005</b> <sup>115</sup>	computer vision	proprietary	512	165	no	ANN	no	histopathology	

<b>SWANSON 2010</b> <sup>116</sup>	computer vision	proprietary	118	11	yes	threshold	no	histopathology	Yes
<b>TEIXEIRA SOUSA 2017</b> <sup>117</sup>	deep learning	public	2750	521	yes	ANN	yes	histopathology*, expert consensus	Yes
<b>TENENHAUS 2010</b> <sup>118</sup>	computer vision	proprietary	227	32	no	partial least squares model	no	histopathology*, clinical evaluation	Yes
<b>TOMATIS 2003</b> <sup>119</sup>	hardware-based	proprietary	573	132	yes	ANN	no	histopathology	
<b>TOMATIS 2005</b> <sup>120</sup>	hardware-based	proprietary	2507	184	yes	ANN	no	histopathology*, clinical evaluation	
<b>TSCHANDL 2018</b> <sup>121</sup>	deep learning	public	20329	3044	yes	ANN	no	histopathology*, expert consensus, nr for part of the lesions	Yes
<b>VALLE 2017</b> <sup>122</sup>	deep learning	public	19931	2401	yes	ANN	no	nr	
<b>WADHAWAN 2011</b> <sup>123</sup>	computer vision	public	347	110	nr	SVM	no	dermatoscopy	Yes
<b>WOLF 2013</b> <sup>124</sup>	computer vision	proprietary	188	60	yes	nr	yes	histopathology	Yes
<b>WU 2018</b> <sup>125</sup>	deep learning	public	10015	1113	yes	ANN	no	histopathology*, follow-up, confocal microscopy, expert consensus	
<b>YANG 2017</b> <sup>126</sup>	deep learning	public	2750	521	yes	ANN	yes	histopathology*, expert consensus	Yes
<b>YAP 2018</b> <sup>127</sup>	deep learning	proprietary	2917	727	yes	ANN	no	histopathology	Yes
<b>YI 2018</b> <sup>128</sup>	deep learning	public	1479	288	no	ANN	no	histopathology*, dermoscopy evaluation, expert consensus	
<b>YU C 2018</b> <sup>129</sup>	deep learning	proprietary	724	350	no	ANN	no	histopathology	Yes
<b>YU L 2017</b> <sup>130</sup>	deep learning	public	1279	248	no	ANN	no	histopathology*, expert consensus, nr for part of the lesions	Yes
<b>ZEKOVIC 2014</b> <sup>131</sup>	hardware-based	proprietary	48	11	yes	partial least squares model	no	histopathology	
<b>ZHOU 2011</b> <sup>132</sup>	hardware-based	proprietary	76	23	nr	ensemble classifier	no	nr	



**eTable 2: Assessment of bias risk (br) and applicability concerns (ap) of the included studies (n=132) using the QUADAS-2 tool.**

STUDY ID	SAMPLE SELECTION (BR)	INDEX TEST (BR)	REFERENCE STANDARD (BR)	FLOW AND TIMING (BR)	SAMPLE SELECTION (AP)	INDEX TEST (AP)	REFERENCE STANDARD (AP)
ABUZAGHLEH 2015 <sup>1</sup>	high	high	low	low	low	low	low
AFIFI 2017 <sup>2</sup>	high	high	moderate	high	low	low	low
AMOABEDINI 2018 <sup>3</sup>	high	high	moderate	high	low	low	low
ASCIERTO 2010 <sup>4</sup>	low	low	low	low	low	low	low
BARATA 2015A <sup>5</sup>	high	high	low	low	low	low	low
BARATA 2015B <sup>6</sup>	high	high	low	low	low	low	low
BARROS MENDES 2018 <sup>7</sup>	high	high	moderate	high	low	low	low
BARZEGARI 2005 <sup>8</sup>	low	high	low	low	low	low	low
BAUR 2018 <sup>9</sup>	high	high	low	low	low	low	low
BHATTACHARYA 2017 <sup>10</sup>	high	high	moderate	moderate	low	low	low
BI 2017 <sup>11</sup>	high	low	moderate	low	low	low	low
BISSOTO 2018 <sup>12</sup>	high	low	moderate	low	moderate	moderate	low
BONO 2002 <sup>13</sup>	high	low	low	low	moderate	moderate	low
CARCAGNI 2018 <sup>14</sup>	high	low	low	low	moderate	moderate	low
CELEBI 2007 <sup>15</sup>	high	high	low	high	low	low	low
CELEBI 2008 <sup>16</sup>	high	high	low	low	moderate	moderate	low
CHAKRAVATI 2015 <sup>17</sup>	high	high	moderate	high	low	low	low
CHANG H 2017 <sup>18</sup>	high	low	moderate	low	low	low	moderate
CHANG WY 2013 <sup>19</sup>	high	high	low	low	moderate	moderate	low
CHANG WY 2015 <sup>20</sup>	high	high	low	low	moderate	moderate	low
CHOU 2017 <sup>21</sup>	high	high	low	low	low	low	low
CODELLA 2016 <sup>22</sup>	high	high	high	low	low	low	low
CODELLA 2018 <sup>23</sup>	high	high	low	low	low	low	low
DEL ROSARIO 2018 <sup>24</sup>	high	high	low	high	low	low	low
DING 2015 <sup>25</sup>	high	high	moderate	moderate	low	low	moderate
DO 2014 <sup>26</sup>	high	high	moderate	moderate	moderate	low	moderate

<b>DO 2017<sup>27</sup></b>	high	high	low	low	low	low	low
<b>DREISEITL 2009<sup>28</sup></b>	low	low	low	high	low	low	low
<b>ESTEVA 2017<sup>29</sup></b>	high	low	low	low	low	low	low
<b>FERRI 2010<sup>30</sup></b>	high	low	low	low	low	low	low
<b>FINK 2016<sup>31</sup></b>	high	low	high	high	low	low	low
<b>FORNACIALI 2016<sup>32</sup></b>	high	high	low	low	low	low	low
<b>FORSCHNER 2018<sup>33</sup></b>	high	low	low	low	moderate	moderate	low
<b>FRIEDMAN 2008<sup>34</sup></b>	high	low	low	low	moderate	moderate	low
<b>GAREAU 2017<sup>35</sup></b>	high	high	low	high	low	low	low
<b>GARNAVI 2012<sup>36</sup></b>	high	high	high	moderate	low	low	moderate
<b>GAUTAM 2018<sup>37</sup></b>	high	high	moderate	moderate	low	low	low
<b>GILMORE 2009<sup>38</sup></b>	high	high	high	low	low	low	low
<b>GILMORE 2010<sup>39</sup></b>	high	high	low	low	low	low	low
<b>GILMORE 2018<sup>40</sup></b>	high	high	low	low	low	low	low
<b>GONZALEZ DIAZ 2017<sup>41</sup></b>	high	low	low	low	low	low	low
<b>GOYAL 2018<sup>42</sup></b>	high	low	low	low	moderate	moderate	low
<b>GUO S 2017<sup>43</sup></b>	high	low	low	low	low	low	low
<b>GUO Y 2018<sup>44</sup></b>	high	low	low	low	moderate	moderate	low
<b>HAENSSLE 2018<sup>45</sup></b>	high	low	low	low	low	low	low
<b>HAIDER 2014<sup>46</sup></b>	high	high	low	low	low	low	low
<b>HARANGI 2017<sup>47</sup></b>	high	low	low	low	low	low	low
<b>HARANGI 2018<sup>48</sup></b>	high	high	low	low	low	low	low
<b>HARDIE 2018<sup>49</sup></b>	high	high	low	low	moderate	moderate	low
<b>HAR-SHAI 2005<sup>50</sup></b>	low	low	low	high	low	low	low
<b>HAUSCHILD 2014<sup>51</sup></b>	high	low	low	low	moderate	moderate	low
<b>HUNGER 2012<sup>52</sup></b>	low	low	low	low	low	low	low
<b>JAWOREK-KORJAKOWSKA 2016A<sup>53</sup></b>	high	high	low	low	low	low	low
<b>JAWOREK-KORJAKOWSKA 2016B<sup>54</sup></b>	high	high	low	low	low	low	low
<b>JIJI 2017<sup>55</sup></b>	high	low	low	low	low	low	low
<b>KAUR 2015<sup>56</sup></b>	high	high	low	low	low	low	low

<b>KAWAHARA 2018<sup>57</sup></b>	high	high	moderate	low	moderate	moderate	low
<b>KHAN 2018<sup>58</sup></b>	high	high	moderate	moderate	low	low	low
<b>KITADA 2018<sup>59</sup></b>	high	low	low	low	moderate	moderate	low
<b>KOSTOPOULOS 2017<sup>60</sup></b>	high	low	moderate	low	low	low	low
<b>LAI 2018<sup>61</sup></b>	high	high	low	low	low	low	low
<b>LEE 2018<sup>62</sup></b>	high	low	low	low	moderate	moderate	low
<b>LI K 2018<sup>63</sup></b>	high	high	low	low	moderate	moderate	low
<b>LI L 2014<sup>64</sup></b>	high	high	low	low	low	low	low
<b>LI Y 2017<sup>65</sup></b>	high	low	low	low	low	low	low
<b>LINGALA 2014<sup>66</sup></b>	high	high	low	low	low	low	low
<b>LIU 2015<sup>67</sup></b>	high	low	high	low	low	low	low
<b>LU 2018<sup>68</sup></b>	high	high	low	low	moderate	moderate	low
<b>MAGLIOGIANNIS 2004<sup>69</sup></b>	high	high	low	low	low	low	low
<b>MAGLIOGIANNIS 2015<sup>70</sup></b>	high	high	low	low	low	low	low
<b>MAHBOD 2017<sup>71</sup></b>	high	low	low	low	low	low	low
<b>MAITNER 2018<sup>72</sup></b>	high	low	low	low	moderate	moderate	low
<b>MALVEHY 2014<sup>73</sup></b>	low	low	low	low	moderate	moderate	low
<b>MANOUSAKI 2006<sup>74</sup></b>	high	high	low	low	low	low	low
<b>MARCHETTI 2018<sup>75</sup></b>	high	high	low	low	low	low	low
<b>MARQUES 2012<sup>76</sup></b>	high	low	low	low	low	low	low
<b>MATSUNAGA 2017<sup>77</sup></b>	high	low	low	low	low	low	low
<b>MENEGOLA 2016<sup>78</sup></b>	high	high	low	low	low	low	low
<b>MENEGOLA 2017A<sup>79</sup></b>	high	high	low	low	low	low	low
<b>MENEGOLA 2017B<sup>80</sup></b>	high	low	low	low	low	low	low
<b>MESSADI 2009<sup>81</sup></b>	high	high	high	low	low	low	low
<b>MIRUNALINI 2017<sup>82</sup></b>	high	low	low	low	low	low	low
<b>MOLLERSEN 2015<sup>83</sup></b>	low	low	low	low	moderate	moderate	low
<b>MONHEIT 2011<sup>84</sup></b>	high	low	low	high	moderate	moderate	low
<b>MUNIA 2017<sup>85</sup></b>	high	high	moderate	low	low	low	low
<b>MURPHREE 2017<sup>86</sup></b>	high	low	low	low	low	low	low
<b>NADER VASCONCELOS 2017<sup>87</sup></b>	high	low	low	low	low	low	low

<b>NANNI 2018<sup>88</sup></b>	high	low	low	low	moderate	moderate	low
<b>NASIRI 2018<sup>89</sup></b>	high	low	low	low	moderate	moderate	low
<b>NASR-ESFAHANI 2016<sup>90</sup></b>	high	high	moderate	low	low	low	low
<b>NICOLAS 2014<sup>91</sup></b>	high	high	low	low	low	low	low
<b>PAL 2018<sup>92</sup></b>	high	low	low	low	moderate	moderate	low
<b>PATWARDHAN 2005<sup>93</sup></b>	high	high	low	low	low	low	low
<b>PELLACANI 2004<sup>94</sup></b>	high	high	low	low	low	low	low
<b>PEREZ 2018<sup>95</sup></b>	high	high	low	low	low	low	low
<b>PICCOLO 2014<sup>96</sup></b>	high	low	low	low	low	low	low
<b>POLAP 2018<sup>97</sup></b>	high	high	low	low	low	low	low
<b>QUINZAN 2013<sup>98</sup></b>	low	high	low	low	low	low	low
<b>RADHAKRISHNAN 2017<sup>99</sup></b>	high	high	high	low	low	low	low
<b>RADHAKRISHNAN 2018<sup>100</sup></b>	high	high	low	low	low	low	low
<b>RAMEZANI 2014<sup>101</sup></b>	high	high	moderate	moderate	low	low	low
<b>RAY 2018<sup>102</sup></b>	high	high	low	low	moderate	moderate	low
<b>REZVANTALAB 2018<sup>103</sup></b>	high	high	low	low	moderate	moderate	low
<b>ROCHA 2017<sup>104</sup></b>	low	low	low	low	low	low	low
<b>RUBEGNI 2015<sup>105</sup></b>	high	high	low	low	low	low	low
<b>RUIZ 2008<sup>106</sup></b>	high	high	moderate	moderate	low	low	low
<b>SABBAGHI 2018<sup>107</sup></b>	high	high	low	low	low	low	low
<b>SATHEESHA 2017<sup>108</sup></b>	high	high	moderate	low	low	low	low
<b>SBONER 2003<sup>109</sup></b>	high	high	low	low	low	low	low
<b>SHAIFIEE 2017<sup>110</sup></b>	high	low	low	low	moderate	moderate	low
<b>SITU 2008<sup>111</sup></b>	high	high	moderate	low	low	low	moderate
<b>SITU 2010<sup>112</sup></b>	high	high	moderate	low	low	low	moderate
<b>SITU 2011<sup>113</sup></b>	high	low	low	low	low	low	low
<b>STANLEY 2007<sup>114</sup></b>	high	high	low	low	low	low	low
<b>STOECKER 2005<sup>115</sup></b>	high	low	low	low	low	low	low
<b>SWANSON 2010<sup>116</sup></b>	low	high	low	low	low	low	low
<b>TEIXEIRA SOUSA 2017<sup>117</sup></b>	high	low	low	low	low	low	low
<b>TENENHAUS 2010<sup>118</sup></b>	high	low	moderate	high	low	low	low

<b>TOMATIS 2003</b> <sup>119</sup>	low	high	low	low	low	low	low
<b>TOMATIS 2005</b> <sup>120</sup>	high	low	low	low	low	low	low
<b>TSCHANDL 2018</b> <sup>121</sup>	high	high	low	low	low	low	low
<b>VALLE 2017</b> <sup>122</sup>	high	low	high	low	low	low	low
<b>WADHAWAN 2011</b> <sup>123</sup>	high	low	high	low	low	low	low
<b>WOLF 2013</b> <sup>124</sup>	high	low	low	low	low	low	low
<b>WU 2018</b> <sup>125</sup>	high	high	low	low	moderate	moderate	low
<b>YANG 2017</b> <sup>126</sup>	high	low	low	low	low	low	low
<b>YAP 2018</b> <sup>127</sup>	high	high	low	low	moderate	moderate	low
<b>YI 2018</b> <sup>128</sup>	high	high	low	low	low	low	low
<b>YU C 2018</b> <sup>129</sup>	high	high	low	low	low	low	low
<b>YU L 2017</b> <sup>130</sup>	high	low	high	low	low	low	low
<b>ZEKOVIC 2014</b> <sup>131</sup>	high	low	low	low	low	low	low
<b>ZHOU 2011</b> <sup>132</sup>	high	high	moderate	moderate	low	low	low

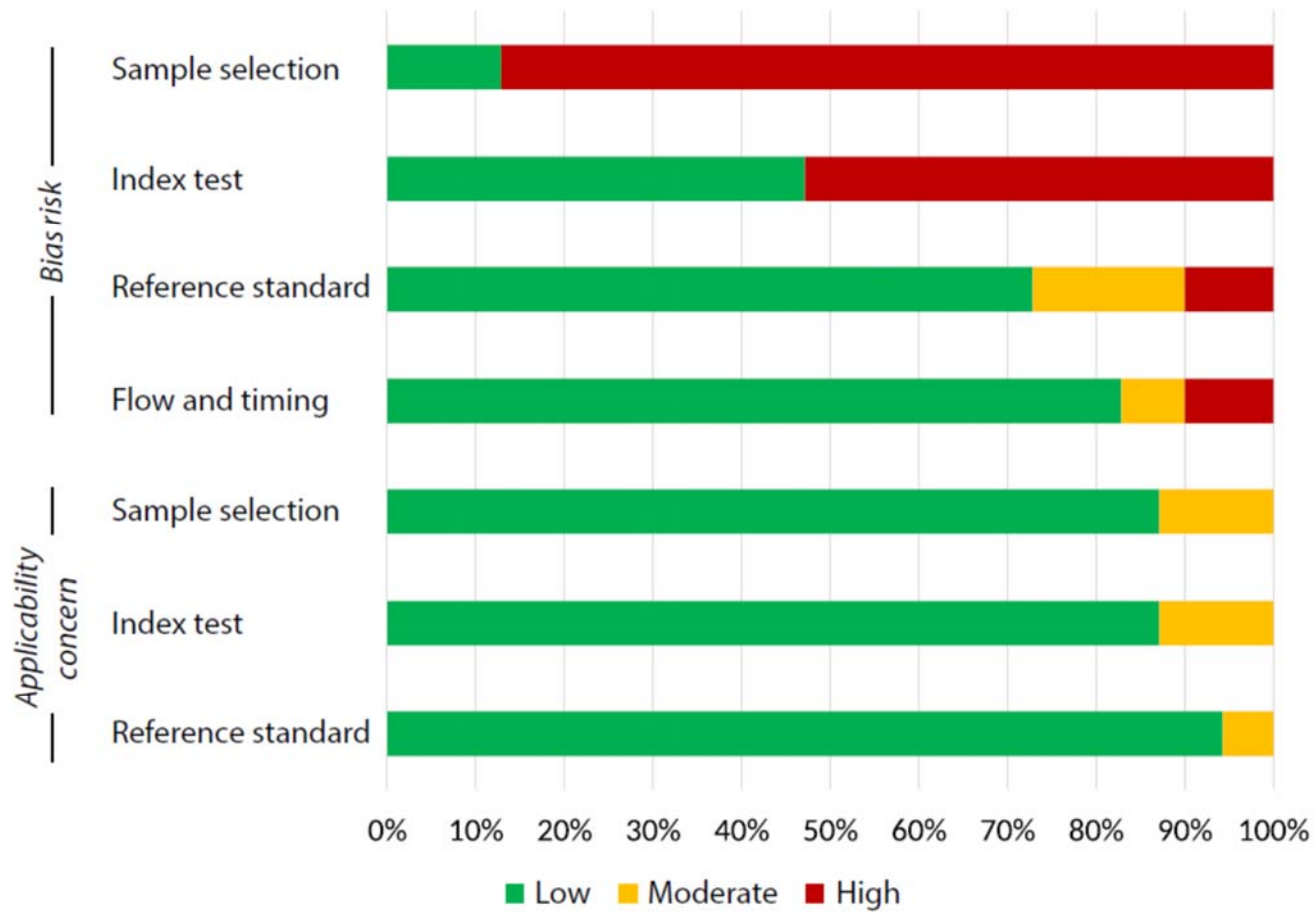
**eTable3: Sensitivity, specificity and covariable effects calculated with two identified outliers (Wolf 2013<sup>124</sup> and Jiji 2017<sup>55</sup>) excluded.**

			<b>Summary Estimate with 95% CI</b>	<b>Univariate p-value</b>
<b>Overall</b>		Sensitivity	0.74 (0.66-0.81)	
		Specificity	0.85 (0.80-0.89)	
<b>Dataset Independence</b>	Independent	Sensitivity	0.51 (0.32-0.70)	
		Specificity	0.85 (0.74-0.92)	
	Non-Independent	Sensitivity	0.82 (0.77-0.86)	p<0.001
		Specificity	0.85 (0.80-0.88)	p=0.967
<b>Dataset Public</b>	Public	Sensitivity	0.57 (0.44-0.69)	
		Specificity	0.92 (0.88-0.94)	
	Proprietary	Sensitivity	0.87 (0.83-0.91)	p<0.001
		Specificity	0.73 (0.64-0.80)	p<0.001
<b>CAD Method</b>	Computer Vision	Sensitivity	0.86 (0.82-0.89)	
		Specificity	0.79 (0.71-0.85)	
	Deep Learning	Sensitivity	0.44 (0.30-0.59)	p<0.001*
		Specificity	0.92 (0.89-0.95)	p<0.001*
	Hardware Based	Sensitivity	0.86 (0.77-0.92)	p=0.899*
		Specificity	0.70 (0.54-0.82)	p=0.352*

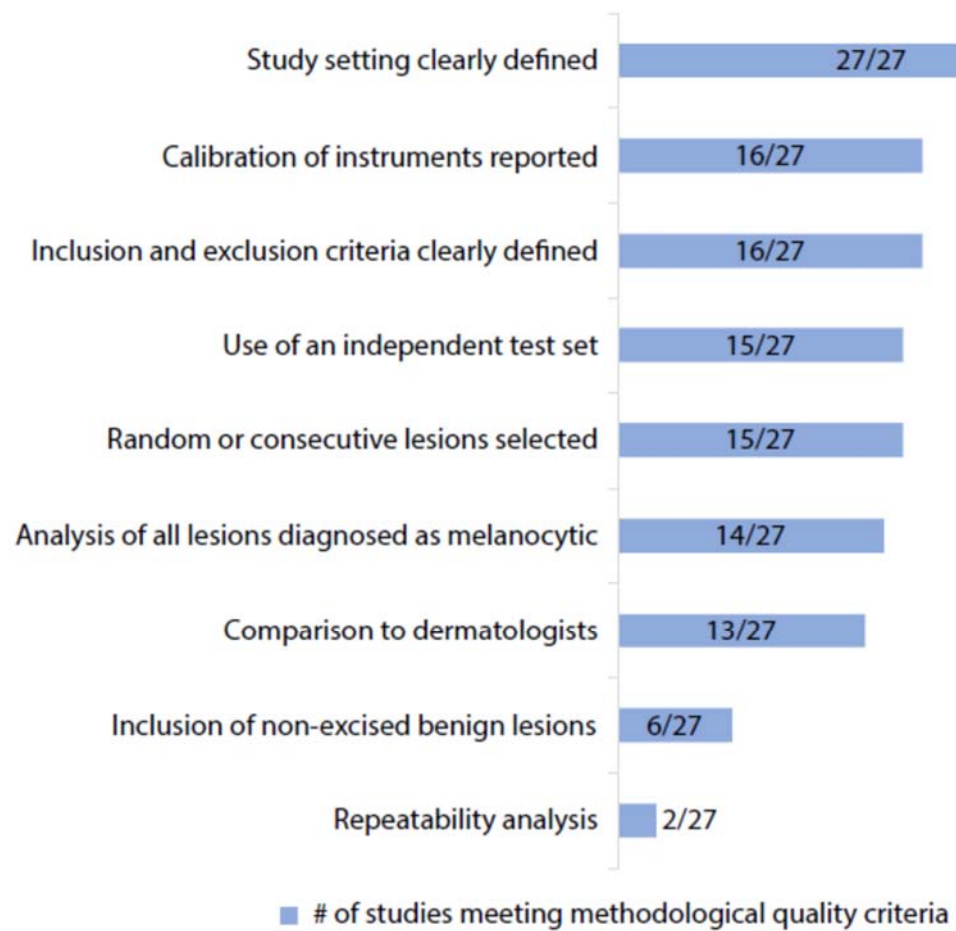
\* Compared to Computer Vision

**eFigure.** Risk of Bias, Applicability Concerns, and Methodologic Quality

eFigure A, Summary of bias risk and applicability concern evaluations for studies included in the quantitative analysis (n = 70) using the QUADAS-2 tool.



eFigure B, Assessment of the methodologic quality of studies from the medical field (n = 27) using predefined criteria.





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