

Study	Study type	Population (n)	Age (mean; range)	Baked-milk challenge	Incidence	Risk factors univariate not associated/ reported	Estimates of univariate prognostic effects						
							Risk factor	Cut-off	AUC (95% CI)	Sens (%)	Spec (%)	PPV (%)	NPV (%)
Bartnikas, 2012	Retrospective cohort	N= 35 Children with IgE-mediated CMA	<i>Median [range]</i> 8.1[3.1-18.1] <u>years</u>	Open OFC with baked muffin. Incremental portions. Max 2.6 g of CM protein	6 (17.1%)	- Age - Sex - other atopic conditions - sIgE α -lactalbumin and β -lactoglobulin. - food challenge results	Casein sIgE CM sIgE Casein SPT (wheat size) Milk SPT (wheat size)	- - ≥ 9 mm ≥ 13 mm	0.65* 0.60* 0.64* 0.70*	- - 66.7 66.7	- - 85.7 72.4	- - NR 33.3	- - 92.3 91.3
Karaman, 2022	prospective cohort	<u>N</u> =35 Children >6 months with IgE-mediated CMA without a history of anaphylaxis during OFC and with low responsiveness to unheated milk	17 (7-48) <u>months</u>	Open OFC with baked muffin. Incremental portions. Max 4.5 g of CM protein	23 (66%)	- Sex; age of allergy onset; accompanying food allergy; reaction to breast milk. - SPT for: unheated milk, β -lactoglobulin, casein, baked milk, - Total IgE - Total eosinophil count	unheated CM (during OFC) BM SPT (wheat size) SPT α -lactoalb. sIgE β -lactogl. sIgE α -laktalb. sIgE casein	620 mg 3 mm 3mm 1.7 kU/l 2.4 kU/l 0.94 kU/l	0.88 (0.77-0.99) NR NR NR NR NR	83.3 74 80 68 62 81	82.6 67 70 70 70 70	NR NR NR NR NR NR	NR NR NR NR NR NR
Kilic, 2021	Prospective cohort	N= 80 Children with IgE-mediated CMA without chronic systemic diseases	<u>unclear</u> article states: "OFC at two years in most cases". In contrast age is stated to be 7.25 (2.45) <u>months</u>	Open OFC with baked muffin. Incremental portions. Max 2.6 g of CM protein	30 (37.5%)	- SPT for CM - Prick-to-prick SPT wheal size for CM - sIgE α -lactalbumin and β -lactoglobulin. - Patch test with CM - Vitamin D levels - Classification of vitamin D status - Total IgE levels -Peripheral eosinophil count	Muffin SPT Casein sIgE Unheated CM (during OFC) CM sIgE	3 mm 3.01 1458 1.79	0.711* 0.755* 0.771* 0.633*	86.6 80.0 88.2 73.3	56.0 74.0 51.2 60.0	54.2 75.0 42.9 52.4	87.5 87.5 91.3 78.9
Kiykim, 2019	Prospective cohort	N=15 Children with IgE-mediated CMA aged 6-48 months without wheat allergy, uncontrolled asthma, rhinitis, dermatitis, or eosinophilic gastroenteropathy	28.4 \pm 12.1 <u>months</u> (range 13-48)	DBPCFC with bread. Incremental portions. Max 3.2g CM protein	6 (40%)	- Total IgE - Easinophils - CM SPT wheal diameter	CM sIgE sIgE β -lactogl. sIgE α -laktalb. casein sIgE	25 7.1 17.6 32 kU/L	0.981* 0.796* 0.981* 0.983*	100 83.3 100 100	89 66.7 89 89	NR NR NR NR	NR NR NR NR
Vilar,2021	Cohort	N=54 Children with IgE mediated CMA	Mean age (range): 3.14(0.5-15) years	open OFC with muffin. no incremental sizes max 1.3 g CM protein	15 (27.8%)	- Age - Sex - Other food allergies - sIgE for: casein, α -lactalb, β -lactoglobulin - SPT to: raw CM, boiled CM, baked CM, α -lactalb, β -lactoglobulin	CM sIgE SPT casein SPT boiled CM SPT BM	5.13 3.2 5.7mm 3.2mm	0.70 (0.53 -0.87) NR NR NR	54 64 76 78	80 77 69 46	NR NR NR NR	NR NR NR NR
Mehr, 2014	Prospective cohort	N=70 Children with IgE mediated CMA following strict avoidance of CM.	5.3 <u>years</u> (SD NR) Neg. responders Median (IQR) 4.5 (2.5-8) <u>years</u> Pos. responders: median (IQR) 7.3 (4.9-9.6) <u>years</u>	open OFC with muffin. Incremental portions. Max 0.5 gram CM protein	19 (27%)	- Age - Sex - CM SPT - baked milk SPT - Eczema - history of anaphylaxis to food other than CM				N neg	N pos	RR	95% CI
										23	18	12.7	1.8-90.1
										12	13	2.8	1.3-6.0
										20	13	2.4	1.0-5.7
										4	6	2.8	1.4-4.5
										14	10	2.1	1.0-4.5