

The role of galectins in immunity and infection

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Supplemental Table 1. Extracellular binding partners identified for galectins*

Galectin	Binding partner
Galectin-1	CD2 ¹ , CD3 ¹ , CD4 ² , CD7 ²⁻⁴ , CD11b, CD11a, CD43 ^{2,3} , CD44 ⁵ , CD45 ^{2,3,6,7} , CD69 ⁸ , CD107, CD146 ⁹ , CD326 ⁵ , TCR ^{10,11} , CR3 ¹² , CEA ¹³ , laminin ^{14,15} , fibronectin ^{14,16} , GM1 ¹⁷ , glycoprotein 90/MAC-2BP ¹⁸ , CA125 ¹⁹ , **pre-B cell receptor ^{20,21} haptoglobin ²² , Vascular Endothelial Growth Factor R2 ²³ , Neuropilin 1 ²⁴ , $\alpha 7\beta 1$ integrin ²⁵ , $\alpha 4\beta 7$ integrin ²⁵ , $\beta 1$ integrin ²⁶ , Osteospondin ²⁷ , Thrombospondin ²⁷ , Thy-1 ⁷ , Tissue plasminogen activator ²⁸ , Lamp1 ^{13,29} Lamp 2 ¹³
Galectin-2	CD14 ³⁰ , $\beta 1$ integrin ³¹ , MUC5AC ³²
Galectin-3	CD7 ³³ , CD66a ³⁴ , CD66b ³⁴ , CD29/B1 integrin ³³ , CD45 ³⁵ , Mac2 ³⁶ , IgE, TRAIL ³⁷ , EGFR ³⁸ , Mac2 BP ³⁹ , laminin ⁴⁰ , Tenascin-C ⁴¹ , Tenascin-R ⁴¹ , Collagen IV ⁴² , Elastin ⁴³ , Hensin ⁴⁴ , Fibronectin ⁴⁵ , NG2 ⁴⁶ , CEA ^{47,48} , $\alpha 2\beta 1$ integrin ⁴² , Muc1 ⁴⁹ , Lamp 1 ⁵⁰ , Lamp 2 ⁵⁰
Galectin-4	Transferrin receptor ⁵¹ , ***CD14 ⁵² , SB1a ⁵³ , SM3 ⁵³ , SM4s ⁵³ , SB2 ⁵³ , SM2a ⁵³ , GM1 ⁵³ , CEA ⁵³
Galectin-7	E-cadherin ^{54,55}
Galectin-8	CD11a/CD18 ⁵⁶ , CD44 ⁵⁷ , CD45 ⁵⁸ , BCR ⁵⁹ , glycoprotein 1b ⁶⁰
Galectin-9	CD5 ⁶¹ , CD22 ⁶² , CD44 ⁶³ , CD45 ^{62,64} , CD180 ⁶¹ , Tim 3 ^{65,66} , PD1 ⁶⁷ , V-domain Ig-containing suppressor of T cell activation (VISTA) ⁶⁸ , B cell receptor ^{61,62} , TLR4 ⁶¹ , Lamp 2 ⁶⁹
Galectin-12	

*Not all binding partners have been tested for every galectin. As a result, a binding partner for one galectin could certainly be engaged by a different galectin family member. However, it should be noted that positive interactions in vitro does not demonstrate that these interactions likewise occur in vivo (see Box 2)

**Extracellular binding partner that interacts with galectins through a carbohydrate-independent processes.

Supplemental Table 2. Impact of common glycan modifications on galectin binding

Galectin	Enhances glycan recognition	Inhibits glycan recognition
Galectin-1	Polylactosamine extension, 3' galactose sulfation ⁷⁰⁻⁷⁶	α 2-6 sialylation, ABO(H) blood group modifications, Lewis ^a or Lewis ^x modification ^{70,74}
Galectin-2	Polylactosamine extension, ABO(H) blood group modifications, ABO(H) blood group modifications ⁷⁴	α 2-3 sialylation, α 2-6 sialylation, Lewis ^a or Lewis ^x modification ^{70,74}
Galectin-3	Polylactosamine extension, ABO(H) blood group modifications ^{70-72,74}	α 2-6 sialylation, Lewis ^a or Lewis ^x modification ^{70,74}
Galectin-4	Polylactosamine extension, ABO(H) blood group modifications, 3' galactose sulfation ^{53,70,77,78}	α 2-3 sialylation, α 2-6 sialylation, Lewis ^a or Lewis ^x modification ⁷⁰
Galectin-7	Polylactosamine extension, H blood group modification ^{70,71,79}	α 2-3 sialylation, α 2-6 sialylation, Lewis ^a or Lewis ^x modification ^{70,79}
Galectin-8**	Polylactosamine extension, 3' galactose sulfation, α 2-3 sialylation, ABO(H) blood group modifications ^{70,72,77,78,80-82}	α 2-6 sialylation, Lewis ^a or Lewis ^x modification ^{70,80,81}
Galectin-9	Polylactosamine extension, 3' galactose sulfation, ABO(H) blood group modifications ^{70,77,78,83}	α 2-6 sialylation, Lewis ^a or Lewis ^x modification ⁷⁰

*This list is far from comprehensive. Instead, the intention of this list is to provide a general sense of the impact of common glycan modifications on glycan binding by distinct members of the galectin family

**Each domain of galectin-8 has been shown to have distinct glycan binding preferences.

Supplemental Table 3. Intracellular binding partners identified for galectins*

Galectin	Binding partner
Galectin-1	H-ras, actin ⁸⁴ , Gemin4 ⁸⁵ , cytochrome oxidase subunit III ⁸⁵ , HBGP82 ⁸⁶ , SUMO-3/SMT3B ⁸⁵ , Protocadherin 24 ⁸⁷ , Toll-interacting protein (Tollip) ⁸⁸
Galectin-2	
Galectin-3	Bcl2 ⁸⁹ , K-Ras ⁹⁰ , Akt ^{91,92} , CD95 (cytoplasmic domain) ⁹³ , Gemin4 ⁸⁵ , Chrp55 ⁹⁴ , Axin, CBP70 ⁹⁵ , β -catenin ⁹⁶ , TTF-1 ⁹⁷ , Synexin ⁹⁸ , Protocadherin 24 ⁸⁷ , Cytokeratins ⁹⁹ , Nucling ¹⁰⁰ , Alix ^{101,102} , NLRP3 ¹⁰³ , TRIM16 ¹⁰⁴
Galectin-4	APC ¹⁰⁵ , Axin ¹⁰⁵ , β -catenin ¹⁰⁵ , Sp1 ¹⁰⁶ , p27 ¹⁰⁶
Galectin-7	Bcl2 ¹⁰⁷ , Toll-interacting protein (Tollip) ⁸⁸
Galectin-8	NDP52 ¹⁰⁸ , E3 ligase ¹⁰⁹ , Regulator-Rag/SLC38A9 ¹¹⁰ , TAK1 ¹¹⁰ , AMPK ¹¹⁰ , SLC38A9 ¹¹⁰
Galectin-9	TAK1 ¹¹¹ , AMPK ^{110,111} , NLRP3 ¹¹² , USP9X ¹¹¹
Galectin-12	VPS13C ¹¹³

*Not all binding partners have been tested for every galectin. As a result, a binding partner for one galectin could certainly be engaged by a different galectin family member. However, it should be noted that positive interactions *in vitro* does not demonstrate that these interactions likewise occur *in vivo* (see Box 2). Glycan-dependent interactions with binding partners exposed to the cytosol can likewise occur and are not outlined here.

Supplemental Table 4. Galectin interactions with microbes

Galectin	Bacteria*	Viruses	Fungus	Parasites
Galectin-1	<i>Streptococcus pneumoniae</i> ¹¹⁴	HIV ¹¹⁵ , human T-cell leukemia virus type 1 (HTLV-1) ¹¹⁶ , Nipah virus ¹¹⁷ , influenza ¹¹⁸		<i>Trichomonas vaginalis</i> ^{119,120} , <i>Trypanosoma cruzi</i> ¹²¹
Galectin-2				
Galectin-3	<i>Klebsiella pneumoniae</i> ¹²² , <i>Pseudomonas aeruginosa</i> ¹²³ , <i>Neisseria gonorrhoeae</i> ¹²⁴ , <i>Neisseria meningitidis</i> ¹²⁵ , <i>Helicobacter pylori</i> ^{126,127} , <i>Salmonella enterica</i> subsp. <i>S. enterica</i> serovar <i>S. minnesota</i> ¹²² , <i>Streptococcus pneumoniae</i> ^{114,128} , <i>Providencia alcalifaciens</i> ¹²⁸	HSV ¹²⁹	<i>Candida albicans</i> ^{130,131} , <i>Cryptococcus neoformans</i> ¹³² , <i>Aspergillus fumigatus</i> ¹³³	<i>Trypanosoma cruzi</i> ^{121,134} , <i>Trichomonas vaginalis</i> ¹²⁰ , <i>Leishmania major</i> ¹³⁵ , <i>Schistosoma mansoni</i> ¹³⁶ , <i>Toxoplasma gondii</i> ¹³⁷
Galectin-4	<i>Escherichia coli</i> ¹³⁸ , <i>Providencia alcalifaciens</i> ¹³⁹ , <i>Klebsiella pneumoniae</i> ¹³⁹			<i>Trypanosoma cruzi</i> ¹²¹
Galectin-7	<i>Escherichia coli</i> ⁷⁹ , <i>Providencia alcalifaciens</i> ⁷⁹ , <i>Klebsiella pneumoniae</i> ⁷⁹ , <i>Streptococcus pneumoniae</i> ⁷⁹			<i>Trypanosoma cruzi</i> ¹²¹
Galectin-8	<i>Escherichia coli</i> ¹³⁸ , <i>Providencia alcalifaciens</i> ¹³⁹ , <i>Klebsiella pneumoniae</i> ¹³⁹			<i>Trypanosoma cruzi</i> ¹²¹
Galectin-9	<i>Escherichia coli</i> ⁸³ , <i>Providencia alcalifaciens</i> ⁸³ , <i>Klebsiella pneumoniae</i> ⁸³ , <i>Streptococcus pneumoniae</i> ⁸³			<i>Leishmania major</i> ¹⁴⁰ , <i>Trypanosoma cruzi</i> ¹²¹

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