## Twelve Clusters of U.S. Transfusion-Associated Babesia microti Cases, 1979–2009\*.

7 single-donation clusters§ A B C D E F G S multidonation	Transfusion RI (2004) RI (2006) VA (2009) NY (1997) NY (1999)	Index  Corecipient Corecipient Index Corecipient Index Corecipient Corecipient Corecipient Corecipient Index  Corecipient Corecipient Corecipient Corecipient Corecipient	Preterm infant Age 70 y: Gi bleeding Preterm infant Age 28 y: SCD	Case†  Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear-positive PCR-positive PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive	Case-Patients‡  Another preterm corecipient of RBCs was treated empirically  No additional information  No other corecipients (Lookback: recipient of RBCs donated 3 m earlier tested negative)  Platelet corecipients of RBCs tested negative (Lookback >1 y earlier: RBC recipient test negative; platelet recipient died ≤3 wk aft transfusion)  Platelet corecipients reportedly was asymptomatic and was not tested (Lookback: "on odverse outcomes" report
B C D E F G S multidonation	RI (2006)  VA (2009)  NY (1997)  NY (1999)	Corecipient Corecipient Index Corecipient Corecipient Corecipient Corecipient Index Corecipient Corecipient Corecipient Corecipient Index Corecipient Index	Preterm infant Age 70 y; Gi bleeding Preterm infant	Smear/PCR-positive Smear/PCR-positive PCR-positive PCR-positive PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive	treated empirically  No additional information  No other corecipients (Lookback: recipient of RBCs donated 3 m earlier tested negative)  Platelet corecipients (age 11 y) and 2 preterm corecipients of RBCs tested negative (Lookback >1 y earlier: RBC recipient testr negative; platelet recipient died ≤3 wk aft transfusion)  Platelet corecipient reportedly was asymptomatic and was not tested (Lookback: "no adverse outcomes" report
C  E  F  G  S multidonation	VA (2009)  NY (1997)  NY (1999)	Corecipient Index Corecipient Corecipient Index Corecipient Index Corecipient Index Corecipient Corecipient Index	Preterm infant Age 70 y; Gi bleeding Preterm infant	Smear/PCR-positive Smear-positive PCR-positive PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive	No additional information  No other corecipients (Lookback: recipient of RBCs donated 3 m earlier tested negative)  Platelet corecipient (age 11 y) and 2 preterm corecipients of RBCs tested negative (Lookback > 1 y earlier: RBC recipient tests negative: platelet recipient died ≤3 wk aft transfusion)  Platelet corecipient reportedly was asymptomatic and was not tested (Lookback: "no adverse outcomes" report
C  E  F  G  S multidonation	VA (2009)  NY (1997)  NY (1999)	Index Corecipient Corecipient Index Corecipient Index Corecipient Index Corecipient Index Corecipient	Preterm infant Full-term infant Preterm infant Preterm infant	Smear-positive PCR-positive PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive	No other corecipients (Lookback: recipient of RBCs donated 3 m earlier tested negative)  Platelet corecipient (age 11 y) and 2 preterm corecipients of RBCs tested negative (Lookback >1 y earlier: RBC recipient tests negative; platelet recipient died \$3 wk aft transfusion)  Platelet corecipient reportedly was asymptomatic and was not tested (Lookback: "no adverse outcomes" report
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E F G S multidonation	NY (1997) NY (1999)	Corecipient Index Corecipient Corecipient Index Corecipient Corecipient Corecipient	Preterm infant Preterm infant Preterm infant Preterm infant Full-term infant Preterm infant Age 70 y; GI bleeding Preterm infant	PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive	(Lookback: recipient of RBCs donated 3 m earlier tested negative)  Platelet corecipient (age 11 y) and 2 preterm corecipients of RBCs tested negative (Lookback >1 y earlier: RBC recipient testragative; platelet recipient died \$3 wk aft transfusion)  Platelet corecipient reportedly was asymptomatic and was not tested (Lookback: "no adverse outcomes" report
E F G S multidonation	NY (1997) NY (1999)	Index Corecipient Index  Corecipient Corecipient Corecipient	Preterm infant Preterm infant Preterm infant Full-term infant Preterm infant Age 70 y; GI bleeding Preterm infant	Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive	(Lookback: recipient of RBCs donated 3 m earlier tested negative)  Platelet corecipient (age 11 y) and 2 preterm corecipients of RBCs tested negative (Lookback >1 y earlier: RBC recipient testragative; platelet recipient died \$3 wk aft transfusion)  Platelet corecipient reportedly was asymptomatic and was not tested (Lookback: "no adverse outcomes" report
E F G S multidonation	NY (1997) NY (1999)	Corecipient Index  Corecipient Corecipient Corecipient Index	Preterm infant Preterm infant Full-term infant Preterm infant Age 70 y; Gi bleeding Preterm infant	Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive	(Lookback: recipient of RBCs donated 3 m earlier tested negative)  Platelet corecipient (age 11 y) and 2 preterm corecipients of RBCs tested negative (Lookback >1 y earlier: RBC recipient testragative; platelet recipient died \$3 wk aft transfusion)  Platelet corecipient reportedly was asymptomatic and was not tested (Lookback: "no adverse outcomes" report
F G multidonation	NY (1999)	Corecipient Index  Corecipient  Corecipient Index	Preterm infant Full-term infant Preterm infant Age 70 y; GI bleeding Preterm infant	Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive	earlier tested negative)  Platelet corecipient (age 11 y) and 2 preterm corecipients of RBCs tested negative (Lookback >1 y earlier: RBC recipient test negative: platelet recipient died s3 wk aff transfusion)  Platelet corecipient reportedly was asymptomatic and was not tested (Lookback: "no adverse outcomes" report
F G multidonation	NY (1999)	Index  Corecipient  Corecipient Index	Full-term infant  Preterm infant  Age 70 y; GI bleeding  Preterm infant	Smear/PCR-positive  Smear/PCR-positive  Smear/PCR-positive  Smear/PCR-positive	preterm corecipients of RBCs tested negative (Lookback >1 y earlier: RBC recipient test negative; platelet recipient died ≤3 wk aft transfusion)  Platelet corecipient reportedly was asymptomatic and was not tested (Lookback: "no adverse outcomes" report
F G multidonation	NY (1999)	Corecipient  Corecipient Index	Preterm infant  Age 70 y; GI bleeding  Preterm infant	Smear/PCR-positive Smear/PCR-positive Smear/PCR-positive	preterm corecipients of RBCs tested negative (Lookback >1 y earlier: RBC recipient tests negative; platelet recipient died ≤3 wk aft transfusion)  Platelet corecipient reportedly was asymptomatic and was not tested (Lookback: "no adverse outcomes" report
F G 5 multidonation	7,700	Corecipient Index	Age 70 y; GI bleeding Preterm infant	Smear/PCR-positive Smear/PCR-positive	(Lookback >1 y earlier: RBC recipient test- negative; platelet recipient died ≤3 wk aft transfusion)
F G multidonation	7,700	Index	Preterm infant	Smear/PCR-positive	Platelet corecipient reportedly was asymptomatic and was not tested (Lookback: "no adverse outcomes" report
F G 5 multidonation	7,700	Index	Preterm infant	Smear/PCR-positive	asymptomatic and was not tested (Lookback: "no adverse outcomes" report
F G multidonation	7,700				asymptomatic and was not tested (Lookback: "no adverse outcomes" report
G multidonation		Corecipient	Age 28 y; SCD	PCR-positive	
G multidonation					for recipients associated with 2 previous donations)
multidonation	CT (2006)	Index	Neonate	"Proven infection"	No additional information
multidonation		Corecipient	Age 32 y; SCD	"Proven infection"	-
	MN (2008)	Index	Age 92 y; asplenic	Smear/PCR-positive	Double RBC donation: both recipients became infected and are listed here
		Corecipient	Age 36 y; surgery	PCR-positive	-
clusters   H	MN (1999)	Lookback (July donation)	Age 78 y; GI bleeding	PCR-positive	Platelet corecipient (age 70 y) tested negative about 8 mo after transfusion
	MN (1999)	Lookback (September donation)	Age 80 y	PCR-positive	No corecipients
	MN (1999)	Index (November donation)	Age 68 y; surgery	Smear/PCR-positive	Platelet corecipient (age 81 y) tested
	MN (2000)	Lookforward (January donation)	Age 67 y; surgery	Seropositive	negative about 6 mo after transfusion RBC corecipient (age 73 y) died 2 d after transfusion
I .	NY (2002)	Lookback (March donation)	Age 78 y; surgery	PCR-positive	(Further lookback: recipient associated w December 2001 donation tested negative
	NY (2002)	Index (May donation)	Age 80 y; cirrhosis	Smear-positive	No corecipients
1	NY (2002)	Lookback (October donation)	Age 52 y; surgery	Seropositive	(Further lookback: no information about
					recipient of RBCs donated in August)
	MA (2004)	Index (December 2003 donation)	Age 74 y; carcinoma	Smear-positive	No additional information
К	WI (2007)	Lookback (August donation)	Age 83 y; surgery	Seropositive	(Status of other recipients of RBCs donate in 2007: 2 died; 1 tested negative; 1 lost t follow-up)
	FL (2008)	Index (February 2008 donation)	Age 83 y; GI bleeding	Smear/PCR-positive	No corecipients
L	MN (2008)	Index (August donation)	Age 61 y; leukemia	Smear/PCR-positive	No corecipients (lookback: RBC recipient associated with May donation tested
		Lookforward (October donation)	Age 53 y; surgery	Seropositive	negative) No corecipients

CT = Connecticut; FL = Florida; GI = gastrointestinal; MA = Massachusetts; MN = Minnesota; NY = New York; PCR = polymerase chain reaction; RBC = red blood cell; RI = Rhode Island; SCD = sickle cell disease; VA = Virginia; WI = Wisconsin.

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<sup>\*</sup>The 12 identified clusters encompass 30 cases (1 per row) linked to 19 donations by the 12 implicated donors; the 30 cases include 12 index and 18 nonindex cases (11 in corecipients, 5 detected in lookback investigations, and 2 from lookforward investigations). One case was linked to whole blood-derived platelets (cluster H; fourth donation (5); the other 29 were linked to RBC components. Among infants with available data, the smallest transfused volume was approximately 8 mL in 2 multidonation clusters (1 and k1, case-patients were identified in 2 states, in cluster I, both donations were in Maline, by a donor probably exposed in Massachusetts; in cluster K, a Wisconsin resident also donated in Florida. Five of 12 implicated donors had parasitologically confirmed infection, on the basis of testing an original unit segment (8, C, D, and G) or subsequent specimens (H); the donor linked to cluster H still had demonstrable parasitemia, by PCR analyses, 4 mo after the fourth donation, 10 mo after exposure (5). For cluster A's donor, a segment was available but results of PCR analyses were negative.
\*\*Seropositive\*\* is noted only for the 4 nonindex cases that were not parasitologically confirmed: The reciprocal antibody tetres ranged from 256 to 1024 in 8. microti indirect fluorescent antibody testing.

<sup>‡</sup>For recipients other than case-patients, "tested negative" denotes seronegativity, at a minimum.

<sup>§ 18</sup> cases (13 in infants and 5 in adults); 2–3 cases per cluster.

<sup>| 12</sup> cases (all in adults); 1 case per donation; 2-4 donations per cluster.

## Characteristics of U.S. Transfusion-Associated Babesia microti Cases, Stratified by Type and Class (n= 159 Total Cases, Including 141 Index Cases), 1979–2009\*.

Variable	All Cases (n = 159)	Stratification of All Cases, by Type (n = 159)		Stratification of Index Cases, by Class (n = 141)		
		Index Cases Nonindex Cases (n = 141; 89%) (n = 18; 11%)		Definite Cases Probable Cases Possible Cases (n = 61; 43%) (n = 57; 40%) (n = 23; 16%)		
Age at diagnosis, n	157	139	18	60	56	23
Median age (range: IQR), y†	65 (<1-94: 39-78)	66 (<1-94: 44-79)	34 (<1-83: <1-70)	69 (<1-94: 27-81)	65 (<1-92: 45-78)	67 (<1=87: 53=7
Patients age <1 y, n (%)+	18 (11)	11 (8)	7 (39)	9 (15)	1(2)	1(4)
Patients age ≥1 y to <50 y, n (%)	33 (21)	30 (22)	3 (17)	11 (18)	16 (29)	3 (13)
Patients age ≥50 y, n (%)	106 (68)	98 (71)	8 (44)	40 (67)	39 (70)	19 (83)
Male sex, n/n (%)	78/156 (50)	73/138 (53)	5/18 (28)	25/60 (42)	33/55 (60)	15/23 (65)
State of transfusion#						
B. microti-endemic state, subtotal n (%)	138 (87)	122 (87)	16 (89)	44 (72)	57 (100)	21 (91)
Northeast (CT, MA, NJ, NY, or RI), n	118	108	10	-	-	-
Upper Midwest (MN or WI), n	20	14	6	-	-	-
Other state, subtotal n (%)	21 (13)	19 (13)	2 (11)	17 (28)	0	2 (9)
Eastern state, n	17	15	2		-	
Not an eastern state, n	4	4	0	-	-	-
Year of transfusion						
Median (range)	2005 (1979-2009)	2005 (1979-2009)	2004 (1997-2009)	2005 (1980-2009)	2006 (1979-2009)	2005 (1993-2009
By period, n (%)			***************************************			
1979-1984	4(3)	4(3)	0	3 (5)	1(2)	0
1985-1989	3 (2)	3 (2)	0	2 (3)	1(2)	0
1990-1994	6(4)	6(4)	0	1(2)	4(7)	1(4)
1995-1999	24 (15)	19 (14)	5 (28)	9 (15)	5 (9)	5 (22)
2000-2004	31 (20)	26 (18)	5 (28)	13 (21)	9 (16)	4 (17)
2005–2009	91 (57)	83 (59)	8 (44)	33 (54)	37 (65)	13 (57)
Month of symptom onset or diagnosis, n§	-	128	-	56	52	20
Median (range)		Sep (Jan-Dec)	-	Aug (Jan-Dec)	Oct (Jan-Dec)	Sep (Jan-Dec)
Interval from transfusion to diagnosis, n		114			50	
	-	114	-	53		11
Median (range; IQR), d		42 (14-230; 34-53)	-	43 (22-230; 35-52)	42 (14-225; 34-58)	42 (14-54; 21-
Parasitologically confirmed infection, n (%)¶	153 (96)	139 (99)	14 (78)	61 (100)	55 (96)	23 (100)
Surgical splenectomy, subtotal n**	32	32	0	11	12	9
Past history, n	17	17	0	8	8	1
Peritransfusion, n	12	12	0	2	2	
≥1 mo after transfusion, n	3	3	0	1	2	0
Underlying condition or context for transfusion						
(1 per patient), n						
Hematologic disorder, subtotal n	39	37	,	11	20	6
Hematologic cancer	14	14	0	3	7	4
Sickle cell disease	11	9	2	, a		0
Thalassemia major	7	7	0		3	
	7	7	0		3	1
Other hematologic disorder		20	-	1	3	1
Cardiovascular surgery or procedure	22		2	8	7	,
Gastrointestinal disease, bleeding, or surgery	19	17	2	8	6	3
Trauma with posttraumatic splenectomy**	8	8	0	2	2	4
Solid-organ transplantation††	5	5	0	1	4	0
Other surgery, procedure, or trauma	13	9	4	7	2	0
Newborn or complications of prematurity	16	9	7	8	1	0
Carcinoma	13	13	0	5	6	2
Other medical reason or diagnosis	14	14	0	9	3	2
Not specified	10	9	1	2	6	1
All-cause mortality, n (%)##	28 (18)	27 (19)	1(6)	11 (18)	12 (21)	4 (17)
Blood donor, n (%)§§	136 (86)	118 (84)	18 (100)	61 (100)	57 (100)	0
Parasitologically confirmed, subtotal n	24	24	20 (200)	22	2	0
PCR-positive unit segment, n	12	12		12	0	0
CT = Connecticut; IQR = interquartile range; MA = Massacl * Data are number of cases/patients, unless otherwise no * Because a lower proportion of patients with index vs. no adults (P = 0.3).	ted. Diagnosis refers to babesiosis. I onindex cases were younger than 1 y	Transfusion and blood donor refer y of age ( $P = 0.001$ ), the age distrib	to those associated with a case. outions for index vs. nonindex pa	Percentages might not total 100% between tients were significantly different (P	= 0.009), but not if the age com	
adults (P = 0.3).  *See Methods section and Figure 2. The category "easter Washington.  § If both were known and were different, the earlier monify and the second of the	th was specified. Data for the kidney patients ( <b>Table 1</b> ), the interval to d	donor (see text) were not include	ed in analyses of month of diagno s, type of recipient (corecipient v	osis or interval to diagnosis.	investigations. Although most of	

Index cases were known or presumed to be parasitologically confirmed, with the exception of 2 cases classified as probable transfusion cases: the case in the kidney donor (see text) and a case diagnosed in retrospect, after recovery (30).

\*\* The data constitute minimum numbers of case-patients. Among the 12 known to have undergone spienectomy during the peritransfusion period, the contexts were trauma (n = 8) or abdominal surgery for other reasons (n = 4). The cases in the 3

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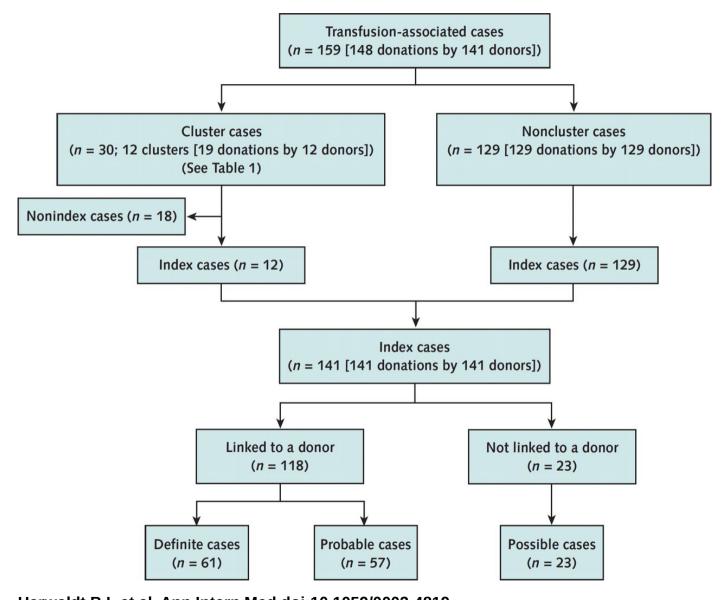
<sup>\*\*</sup> The data constitute minimum numbers of case-patients. Among the 12 known to have undergone splenectomy during the peritransfusion period, the contexts were trauma (n = 8) or abdominal surgery for other reasons (n = 4). The cases in the patients known to have undergone posttransfusion splenectomy include 1 definite case (the index case of cluster L [25]: Table 1) and 2 probable cases, including the first described transfusion case (14).

\*\*Three received a kidney (living related [31]. Inving unrelated, or cadeveric), 1 received a heart [29], and 1 underwent bilasteral lung transplantation.

<sup>##</sup> Although outcome data were unavailable for some patients, we assumed that no other case-patients died in the short term. The patients known to have died include 2 cluster-associated infants whose gestational ages were 23 and 24 wk, 2 (of patients 290 y of age, and 6 (of 32) patients known to have undergone surgical splenectomy.

patients Zery or age, and o (is 2) patients known to make undergone surgical specification. Explicitly of receipt of more than 1 contaminated is could not be excluded.

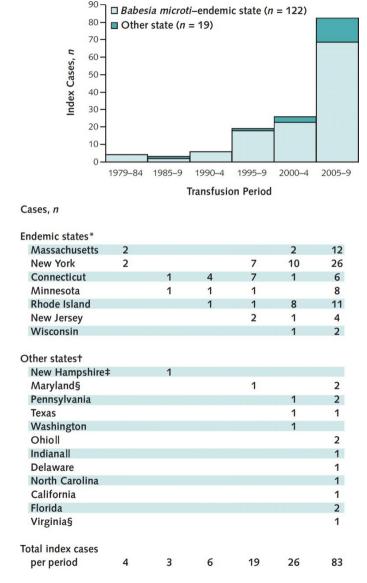
Stratification of 159 U.S. transfusion-associatedBabesia microti cases, 1979–2009.By type of case (cluster vs. not; index vs. not) and by class of index case (definite, probable, or possible).



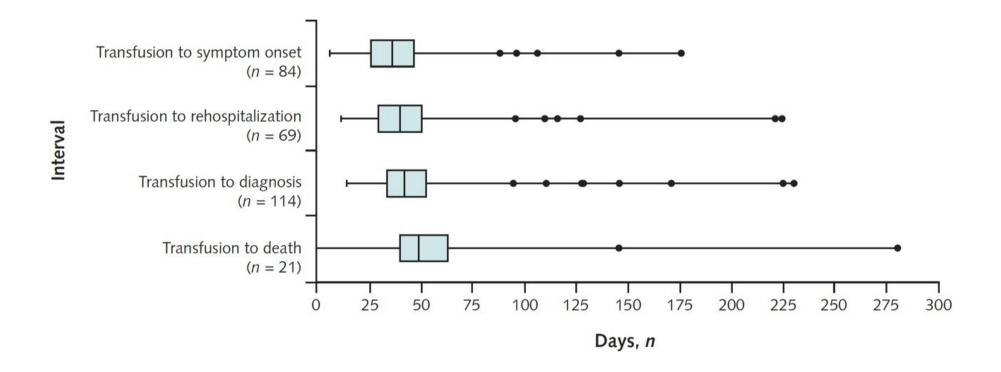
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## Distribution of U.S. transfusion-associatedBabesia microti index cases, 1979–2009.By period and state of transfusion (n = 141 cases).



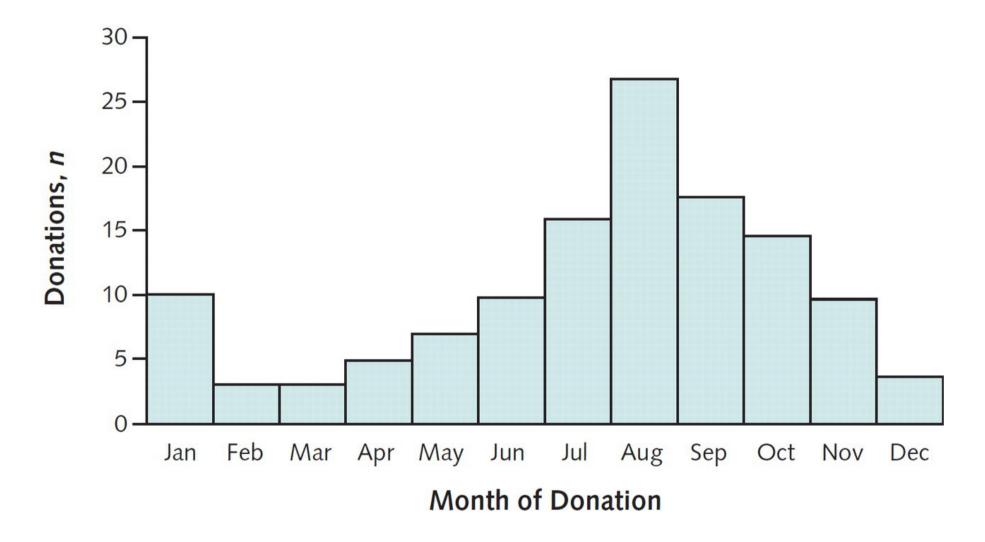
Herwaldt B L et al. Ann Intern Med doi:10.1059/0003-4819-155-8-201110180-00362 Annals of Internal Medicine Box-and-whisker plots of the distributions of time from transfusion to various events for U.S. transfusion-associated Babesia microti index cases, 1979–2009. The data are limited to the subsets of the 141 index patients for whom particular intervals were rel...



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Distribution by month of the blood donations associated with U.S. Babesia microti transfusion cases (n = 128 of 148 total donations), 1979–2009. The month of donation was known or estimable for 128 of 148 donations (by 141 donors) associated with transmissio...



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