

Population Prevalence, Pattern and Associated Factors for Retinal Diseases at High Altitude in Nepal

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Conflict of Interest

No conflict of Interest

Outline of Presentation

- Background and objectives
- Methods
- Result
- Conclusion
- Acknowledgement
- References

Background

- Retinal disorders are the second most common cause of visual impairment and blindness in Nepal ^{1,2}
- Prevalence of retinal disorders in Nepal: 52% \geq age 60 years ²
- Prevalence of retinal disorders at high altitude: 56.2% \geq age 40 years at eye camp.³

Background (contd.)

- High altitude (HA) related health problems: hypoxic and hypobaric conditions, associated lifestyle practices.^{4,6-16}
- Retinal problems: high-altitude sickness (HAS), first described in 1969⁵
- Avoidable ocular and systemic conditions: Awareness, timely precautions and treatment.¹⁴

Background (contd.)

- Mountain: 15% of the land, 7% of the population, Nepal.
- Many hilly districts: altitude greater than 2500 meters ⁴
- Difficult geographical terrain: limited access to transportation and primary eye care services
- Limited information on retinal disorders: high-altitude populations.

Background..

- Study aimed to explore the population prevalence, pattern and associated factors for retinal disorders at high altitude in Nepal.
- Findings helpful in intervention programs: reduce blindness in remote regions.

Objectives

General Objective

- To find the population prevalence, pattern and associated factors of retinal diseases at high altitudes of Nepal

Objectives

Specific Objectives

- To find the prevalence of retinal diseases at high altitudes
- To find out the pattern of retinal diseases at high altitude
- To assess factors associated with retinal diseases at high altitude

Methodology

- Study design: population based, cross sectional study, **Duration:** May 2023 to Aug 2023.
- Study sites: Manang, Mustang, Solukhumbu districts
- HA: usual residence at an altitude of over 2500 meter.⁴
- Existing primary eye care service: for detailed retinal evaluations.

Methodology..

Sample size:

- Retinal problem prevalence (P) 56.8%, 95% confidence and 10% tolerable error, the minimum sample size for the study: 293.
- With 5% non-response rate: sample size (309)

Methodology..

Sampling Techniques

- A list of municipalities and wards of the three study districts.
- Selection of one municipality from each district: using simple random sampling.
- Ward selection: random sampling based on the proportion of the population

Methodology..

Study sites (wards):

- Chame Rural Municipality (Manang)
- Lo-Ghekar Damodarkunda Rural municipality
(Upper Mustang)
- [Khumbu Pasanglhamu](#) rural municipality
(Solukhumbu).

Methodology..

Inclusion Criteria:

Subjects age 40 years and older residing in study sites

Exclusion criteria:

- Retinal evaluation was not possible: hazy media
- Unable to respond the history, examination, and investigation protocol

Methodology (contd.)

Consent: Written informed consent

Detailed history: demographics, ocular, systemic disorder

Examination of eye: presenting and best corrected visual acuity (Snellen chart)

Anterior segment evaluation: slit lamp

Fundus evaluation (mydriasis): Indirect ophthalmoscopy (90 Diopter and 20 Diopter lens)

Methodology (contd.)

- Retina evaluation: fellowship trained retina specialists
- Diabetic retinopathy (DR) and macular edema: Early Treatment Diabetic Retinopathy Study criteria.¹⁸
- AMD: International Age-related Maculopathy Epidemiological Study Group.¹⁹
- Hypertensive retinopathy: Modified Scheie Classification.²⁰

Methodology (contd.)

Fundus photographs: portable fundus camera (Nidek Versacam DS-20) under mydriasis

General examination:

- Blood pressure
- Height and weight

Investigation:

- Pulse oxymetry (Ross-Max)
- Blood sugar: random (venous blood)

Methodology (contd.)

- Diagnostic criteria for diabetes mellitus: use of diabetic medications or a random blood sugar level of 200 mg/dl or greater.^{1,2,17}
- Diagnostic criteria for hypertension: systolic BP \geq 140 mmHg diastolic BP \geq 90 mmHg, or antihypertensive medications.^{1,2}

Methodology..

Ethical approval:

- Ethical Review Board, Nepal Health Research Council (NHRC)
- Protocol Registration number: 199/2023
- Reference number: 3058

Methodology (contd.)

Data analysis

- Data collected: electronic software (Open Data kit) and transferred to MS Excel for cleaning and coding.
- Data analysis: Statistical Package for the Social Science (SPSS) V 20.
- Descriptive results: mean, percentage, tables, and figures, were prepared.

Methodology (contd..)

Data analysis

- Categorical data: Chi Square or Fisher Exact tests
- For numerical data: Independent t test/Mann Whitney U test
- Statistically significant variables: multiple logistic regression to quantify the outcome.
- 95% Confidence interval (CI) of average oxygen level was also calculated.
- A *P*-value <0.05: statistical significant.

Photographs



Photographs



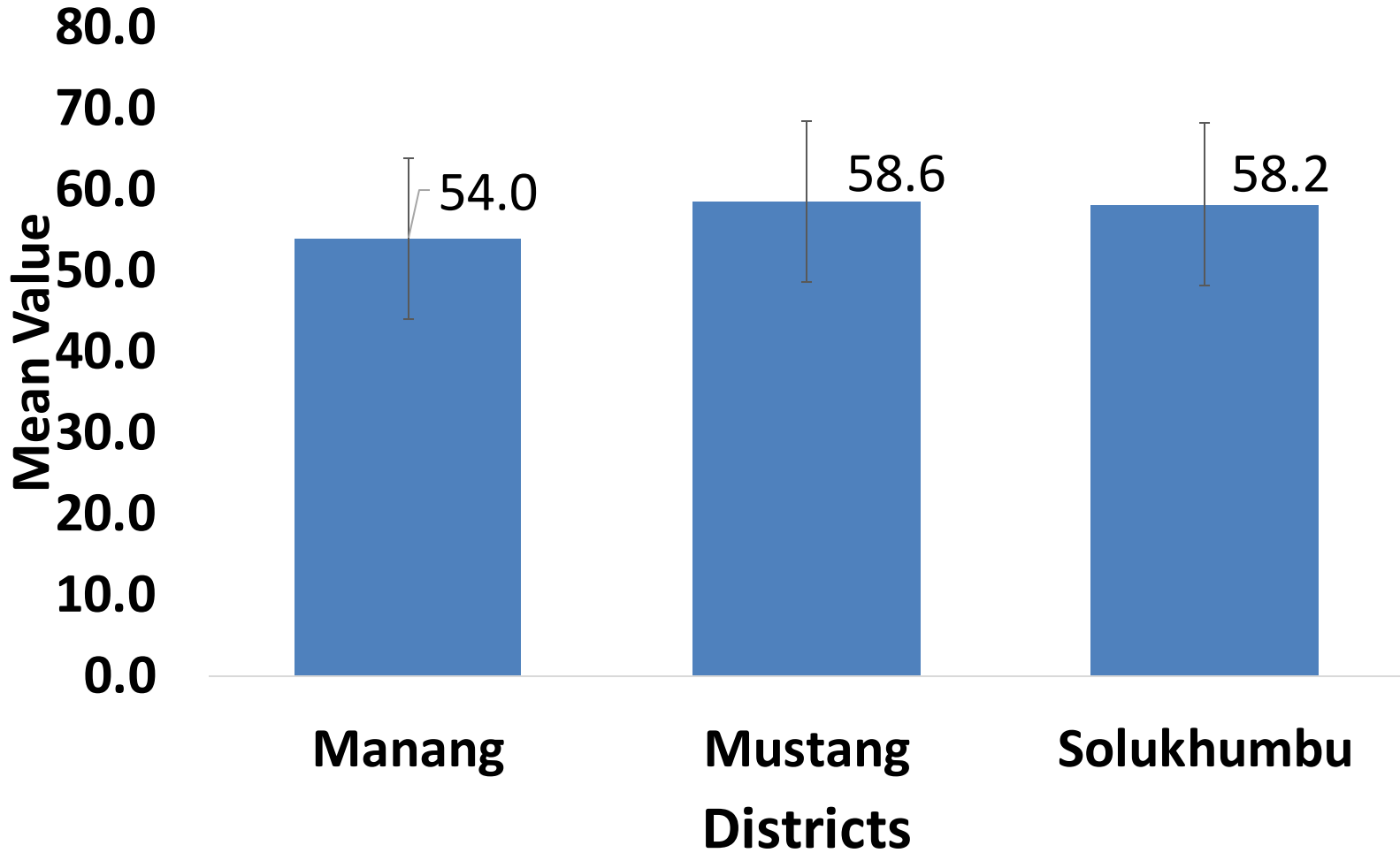
Photographs



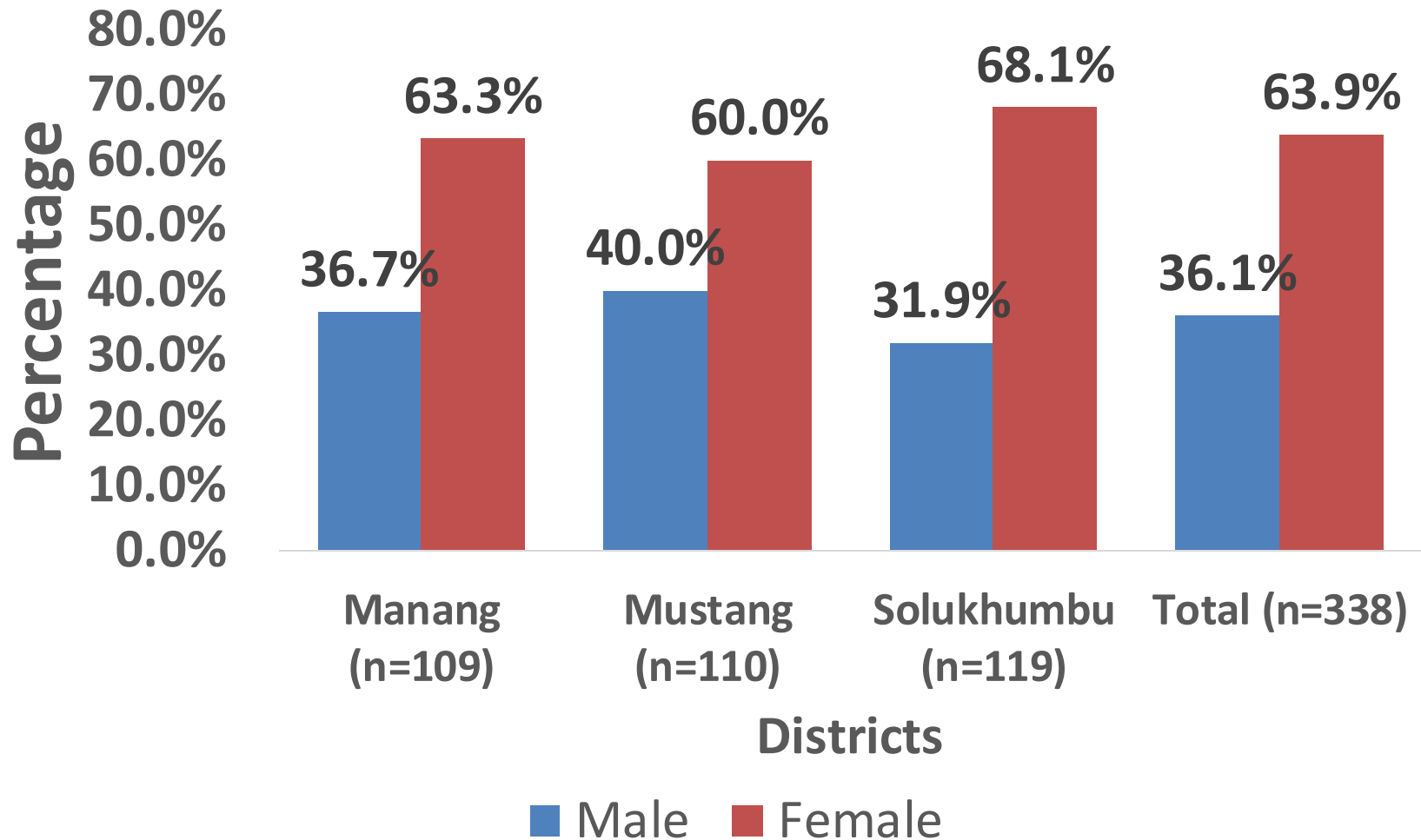
Results

- Total study participants: 338
- Manang: 109 (32.3%)
- Mustang: 110 (32.5%)
- Solukhumbu: 119 (35.2%)
- Mean age of participants: 57.0 years & S.D. 11.1
years

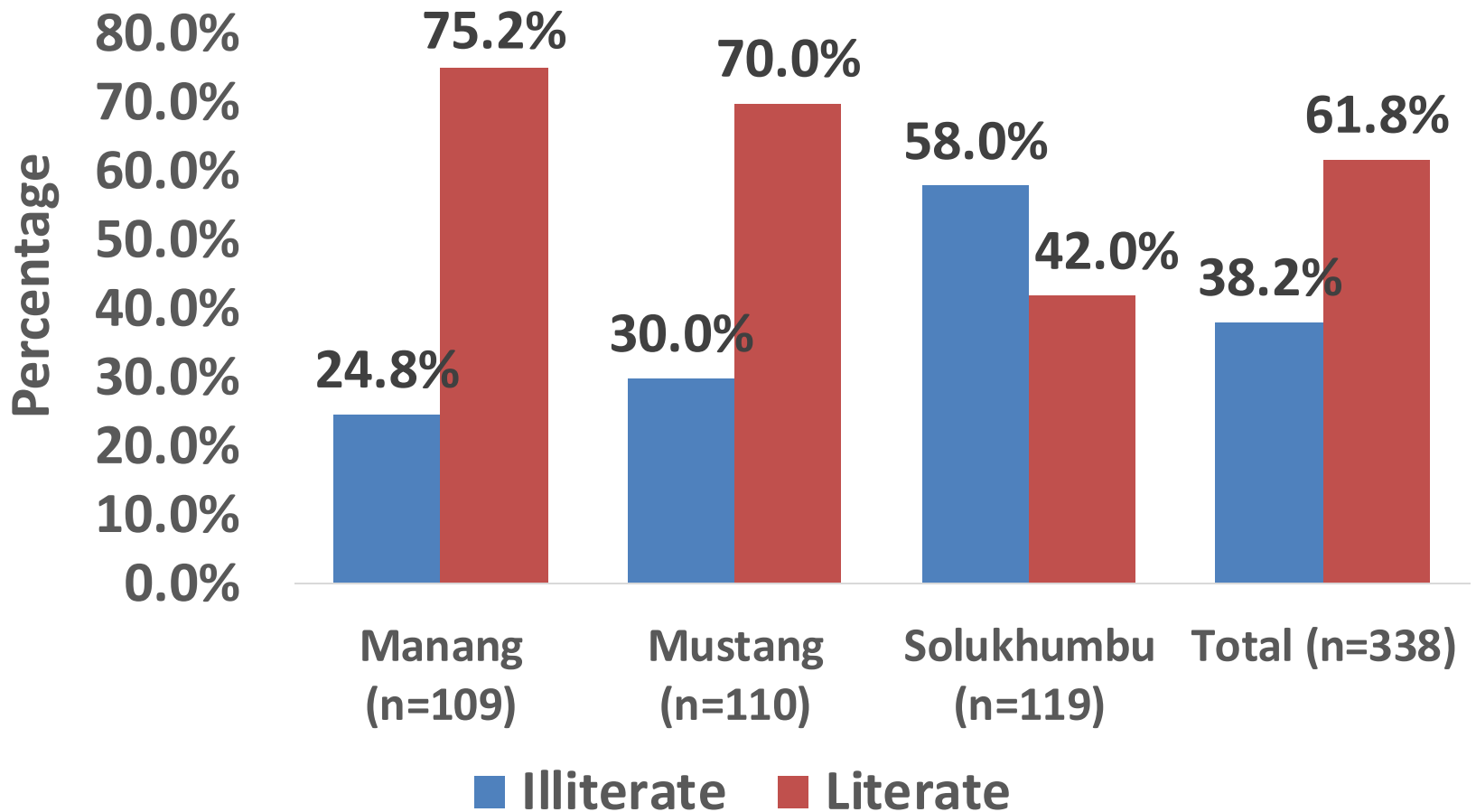
Result: Age (SD) Distribution



Result: Gender Distribution



Results: Literacy Status



Results: Demographic characteristics

Category	Manang, n(%)	Mustang, n(%)	Solukhumbu, n(%)	Total, n(%)
Occupation				
Agriculture	31(28.4)	81(31)	46(38.7)	158(46.7)
Housewife/Household work	24(22.0)	19(24)	18(15.1)	61(18.0)
Business	33(30.3)	2(33)	34(28.6)	69(20.4)
Service	19(17.4)	7(19)	16(13.4)	42(12.4)
Others	2(1.8)	1(2)	5(4.2)	8(2.4)
Religion				
Hinduism	25(22.9)	1(25)	8(6.7)	34(10.1)
Buddhism	81(74.3)	109(81)	106(89.1)	296(87.6)
Others	3(2.8)	0 (0)	5(4.2)	8(2.4)
Total	109(100)	110(109)	119(100)	338(100)

Result: PaO2(%) (Minimum value) in study sites

Districts	Mean (SD)	95% CI
<i>Manang</i>	88.56 (4.22)	87.76 - 89.36
<i>Mustang</i>	85.25 (3.66)	84.55 - 85.94
<i>Solukhumbu</i>	87.87 (3.73)	87.2 - 88.55
<i>Total</i>	87.24 (4.11)	86.8 - 87.68

Results: Visual Acuity Status

- Presenting visual acuity (better eye): 6/6 to 6/18 : **89.1%**
- Best corrected visual acuity (better eye): 6/6 to 6/18: **96.7%**
- BCVA : **3.9%** had low vision, blindness: **no**
- Presenting VA: **4.2%** low vision; Blind: **0.9%**

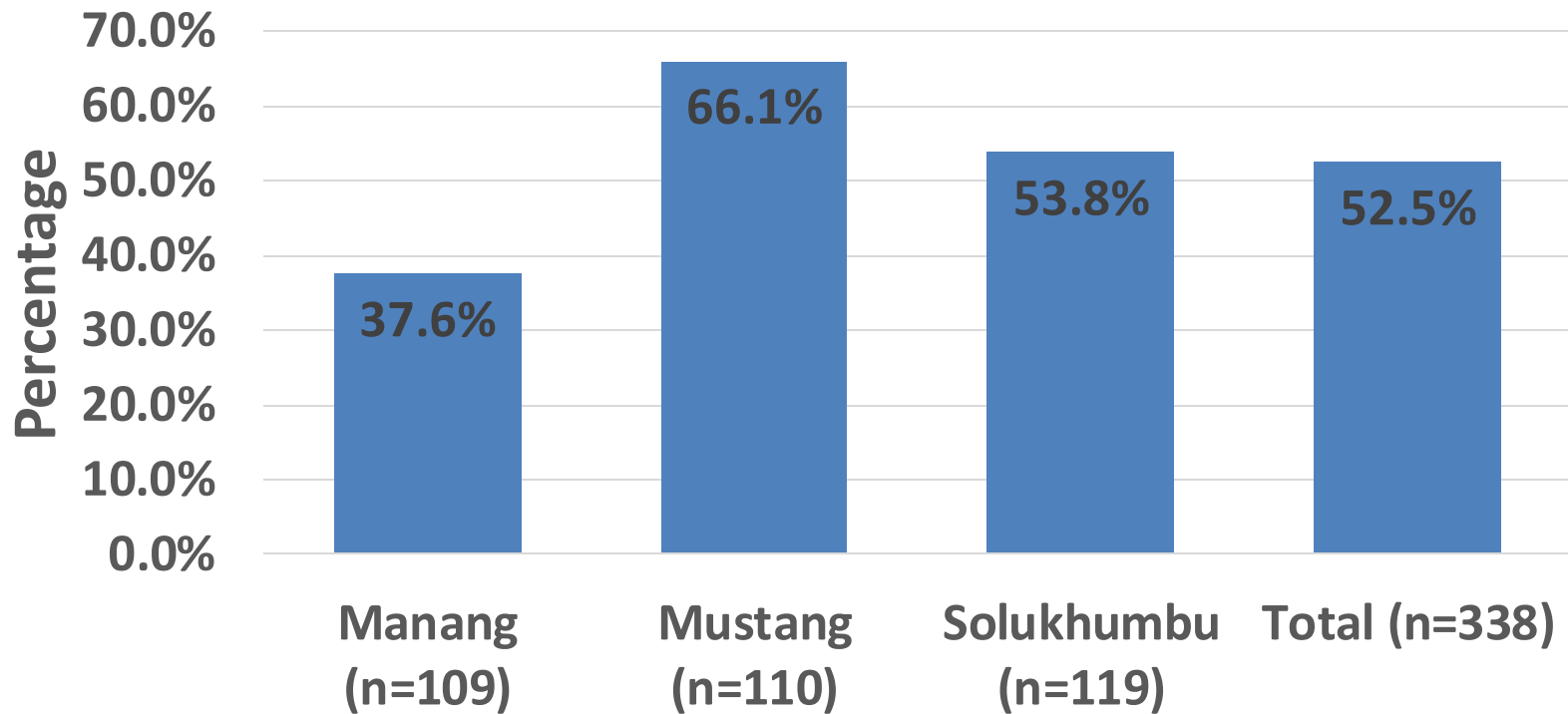
Result: Prevalence of systemic hypertension (HTN)

	Manang	Mustang	Solukhumbu	Total
HTN	n(%)	n(%)	n(%)	n(%)
Known	40(36.7)	18(16.4)	43(36.1)	101(29.9)
New	20(18.3)	32(29.1)	43(36.1)	95(28.1)
No	49(45.0)	60(54.5)	33(27.7)	142(42.0)
Total	109(100)	110(100)	119(100)	338(100)

Result: Prevalence of systemic diabetes mellitus (DM)

	Manang	Mustang	Solukhumbu	Total
DM	n(%)	n(%)	n(%)	n(%)
Known	10(9.2)	7(6.4)	11(9.2)	28(8.3)
New	0	0	9(7.6)	9(2.7)
No	99(90.8)	103(93.6)	99(83.2)	301(89.1)
Total	109(100)	110(100)	119(100)	338(100)

Results: Prevalence of retinal disease



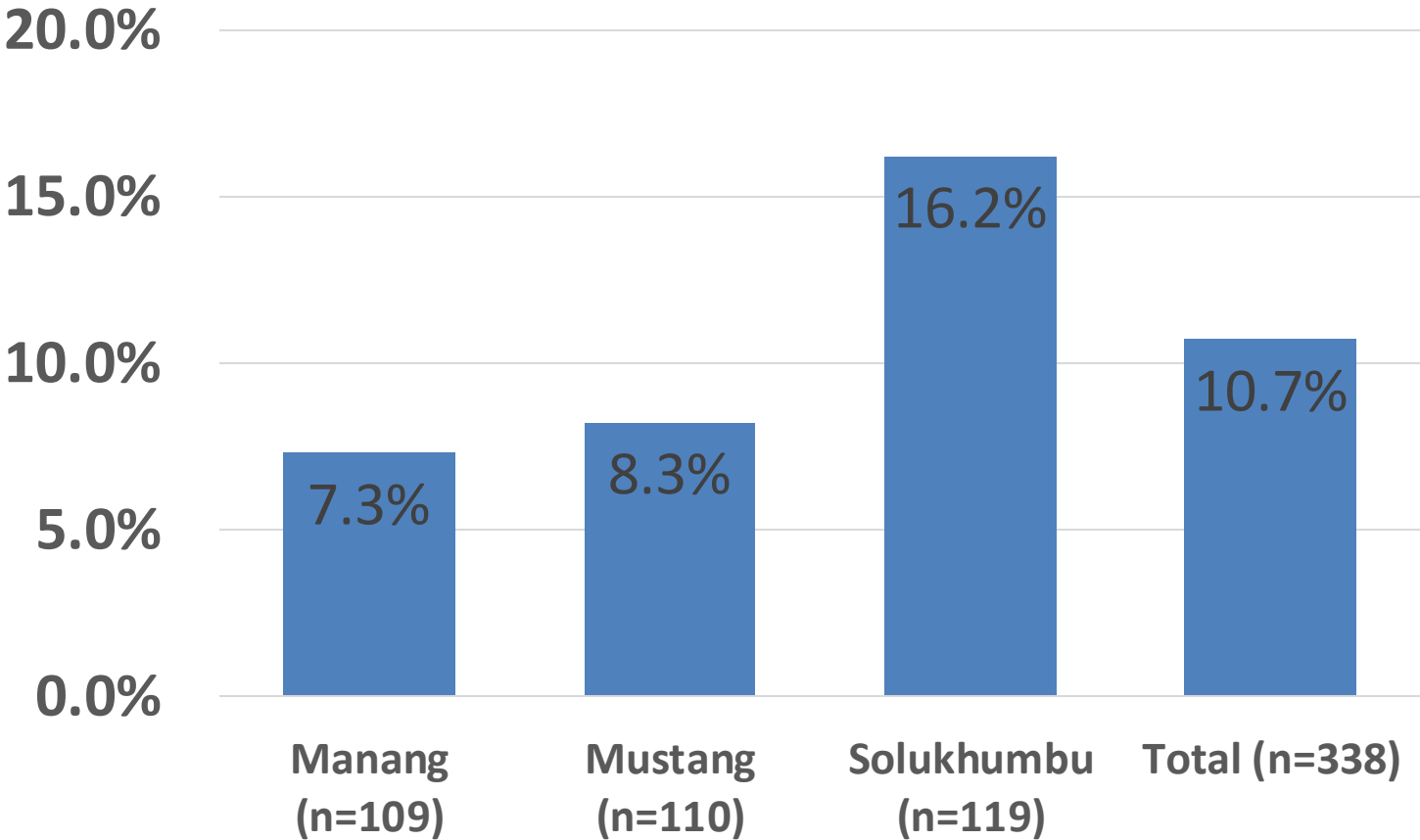
Result: Laterality of retinal disease

Description		Manan	Mustan	Solukhu	Total
		g	g	mbu	
		n(%)	n(%)	n(%)	n (%)
Retinal diseases (Person)	Yes	41(37.6)	72(66.1)	63(53.8)	176(52.5)
	Unilateral	8(7.3)	2(1.8)	9(7.7)	19(5.7)
Laterality	Bilateral	33(30.3)	70(64.2)	54(46.2)	157(46.9)
	Total	109(100)	109(100)	117(100)	335(100)

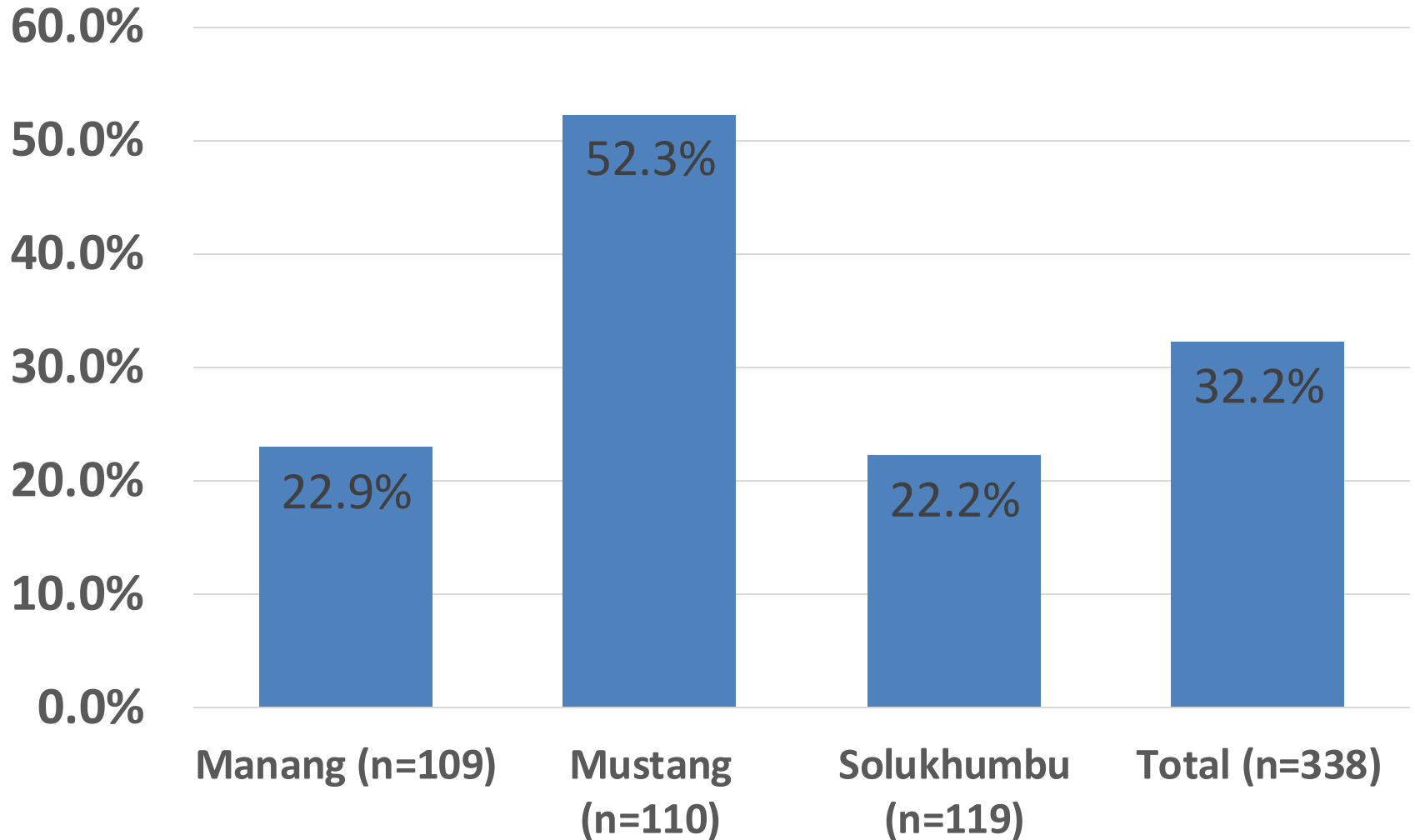
Result: Pattern of retinal diseases

		Manang	Mustang	Solukhum bu	Total
Hypertensive Retinopathy	Yes	25(22.9)	57(52.3)	26(22.2)	108(32.2)
High Altitude Retinopathy(Person)	Yes	8(7.3)	9(8.3)	19(16.2)	36(10.7)
AMD(Person)	Yes	6(5.5)	4(3.7)	17(14.5)	27(8.1)
Diabetic Retinopathy(Person)	Yes	5(4.6)	0	1(0.9)	6(1.8)
Others Retinal Disease(Person)	Yes	10(9.2)	9(8.3)	10(8.5)	29(8.7)

Results: Prevalence of HAR



Results: Prevalence of HTN Retinopathy



Result: Pattern of retinal disease (others) (n=19)

Retinal Disease (Bilateral)	n (%)
Optic Atrophy with Macular Scar	1 (0.3%)
HTN Retinopathy, Lamellar Macular Hole	1 (0.3%)
Lattice Degeneration	1 (0.3%)
Central serous retinopathy	1 (0.3%)
Congenital retinal disorders (macular dystrophy)	2 (0.6%)
Retinitis Pigmentosa	2 (0.6%)
Chorioretinal Scarring	1 (0.3%)
Epiretinal membrane	1 (0.3%)

Result: Pattern of retinal disease (Others)

Retinal Disease (Unilateral)	n (%)
Central serous retinopathy	1 (0.3%)
HTN Retinopathy, foveal Atrophy	2 (0.6%)
Retinal Tear	3 (0.9%)
HTN Retinopathy, Moderate NPDR, Macular Scar	1 (0.3%)
Post Traumatic Maculopathy	1 (0.3%)
Branch retinal vein occlusion	5 (1.5%)
Macular Scar	1 (0.3%)

Result: Pattern of retinal disease (others)

Retinal Disease (Unilateral)	n (%)
Macular Scar	1 (0.3%)
HTN Retinopathy, Macular Hole	1 (0.3%)
RPE Atrophy	1 (0.3%)
HTN Retinopathy, Lattice Degeneration	1 (0.3%)
HTN Retinopathy, Intermediate Dry AMD, Old branch retinal vein occlusion	1 (0.3%)
HTN Retinopathy, Dry Mild AMD, Old branch retinal vein occlusion	1 (0.3%)

Results: Pattern of other ocular morbidities (anterior segment)

No ocular problems: 76 (11.2%)

Anterior Segment eye diseases:

- **Pingecula: 420 eye (62.1%)**
- **Cataract: 197 (29.1%)**
- **Refractive error: 175 (25.9%),**
- **Pterygium: 63(9.3%),**
- **Pseudophakia with capsular opacification: 57 (8.4%)**

Result: Factors with Retinal Diseases (bivariate)

Variable	Category	Total
<i>Age Group</i>	<i><60, 60 and Above</i>	<i><0.001</i>
<i>Gender</i>	<i>Male, Female</i>	<i>0.055</i>
<i>Religion</i>	<i>Buddhism, Others (Hindu, Kirat)</i>	<i>0.332</i>

Result: Factors with Retinal Diseases (bivariate)

Variable	Category	Total
<i>Education</i>	<i>Illiterate, Informal Education, Literate</i>	0.084
<i>Smoking History</i>	<i>Non-smoker, Present smoker, Past smoker</i>	0.047
<i>Alcohol consumption</i>	<i>Yes, No</i>	<0.001

Result: Factors with Retinal Diseases (bivariate)

Variable	Category	Total
<i>Systemic hypertension</i>	<i>Known HTN, New HTN, No HTN</i>	<0.001
<i>Diabetes mellitus</i>	<i>Known DM, New DM, No DM</i>	0.383
<i>Best Corrected Visual Acuity Category</i>	<i>6/6 to 6/18, <6/18 to 6/60, <6/60 to 3/60</i>	0.293
<i>PaO2:(in%) (Lowest value) category</i>	<i><90, 90 and above</i>	0.004
<i>Trekking</i>	<i>Yes, No</i>	0.151
<i>Blood Sugar</i>	<i>Mean (SD)</i>	0.997

Result: Multiple logistic regressions

	p value	OR (95% CI)
Age Group (>60 compared with <60)	0.003	2.19(1.3 to 3.1)
Occupation: (Others)	Reference	
Occupation: Agriculture	0.212	1.43(0.82 to 2.51)
Occupation: Business	0.517	0.79(0.4 to 1.6)
Smoking History (Past Smoker)	Reference	
Smoking History(Non Smoker)	0.334	0.72(0.38 to 1.39)
Smoking History(Present Smoker)	0.938	1.04(0.4 to 2.69)

Result: Multiple logistic regressions

	p value	OR (95% CI)
Alcohol consumption(Yes compared with No)	0.382	1.26(0.75 to 2.12)
HTN (No)	Reference	
HTN(Known)	0.000	5.31(2.86 to 9.86)
HTN(New)	0.045	1.8(1.01 to 3.18)
Constant	0.034	0.42(0 to 0)
SPO2 <90>90%	0.113	1.54 (0.9 to 2.64)

Results: Awareness of major retinal diseases

- Diabetic retinopathy awareness: 15 (4.4%),.
- AMD awareness: 19 (5.6%)
- Hypertensive retinopathy awareness: 10 (23.8%)
- Awareness of high-altitude retinopathy: 10.7%

Fundus photographs



Conclusion

- Over half of the study participants had some form of retinal diseases in one or both eyes.
- Most common retinal disorder were hypertensive retinopathy, high altitude retinopathy and AMD.
- Retinal diseases had significant association with age, and systemic hypertension.

Conclusion..

- Awareness of major retinal disease was very low, despite of high prevalence of blinding disease among the population.

Takeaway message

- There is high prevalence of blinding retinal disease among study population.
- Majority of retinal diseases are avoidable on timely precaution.
- Enhancement of access to eye care services and awareness on major blinding retinal diseases need to be emphasized among population at high altitude in Nepal.

Acknowledgement

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References

1. Thapa SS, Thapa R, Paudyal I, Khanal S, Aujla J, Paudyal G, Rens G. Prevalence and pattern of Vitreo-retinal disorders in Nepal: the Bhaktapur glaucoma study. *BMC Ophthalmology* 2013, 13, 9. doi:10.1186/1471-2415-13-9.
2. Thapa R, Khanal S, Tan HS, Thapa SS, van Rens GHMB. Prevalence, pattern and risk factors of retinal diseases among an elderly population in Nepal: The Bhaktapur Retina Study. *Clinical Ophthalmology* 2020;14:2109-2118. PMID:32801619. PMCID:PMC: 7399464.
3. Thapa R, Paudyal G, Crandall A, Tabin G. Vitreo-retinal disorders at High altitude in Nepal. *Nepalese Journal of Ophthalmology*. 2013; 5(8):57-62. PMID: 23584648

References

4. Aryal N, Weatherall M, Bhatta YKD, Mann S. Lipid Profiles, Glycated Hemoglobin, and Diabetes in People Living at High Altitude in Nepal. *International Journal of Environmental Research and Public Health*. 2017; 14(9):1041. <https://doi.org/10.3390/ijerph14091041>)
5. [Daniel S Morris](#), [John Somner](#), [Michael J Donald](#), [Ian J C McCormick](#), [Rupert R A Bourne](#), [Suber S Huang](#), [Peter Aspinall](#) & [Baljean Dhillon](#). The Eye at Altitude. [Hypoxia and Exercise](#). 2007:249-270.
6. Rennie D, Morrossey J. Retinal changes in Himalayan climbers. *Arch Ophthalmol* 1975;93:395-400.
7. Karki P, Sijapati MJ, Pandey M, Khadka N. Ocular presentations in high altitude sickness: experience of our centre. *Nep Med J* 2019;2(1):141- 4.

8. Mashkovskiy E, Szawarski P, Ryzhkov P, Goslar T, Mrak I. Fulminant high altitude blindness. *Journal of Travel Medicine*, 2016, 1–4.
9. Izadi M, Pourazizi M, Alemzadeh-Ansari Mh. Ocular problems in high-altitude traveling: a review with focus on management. *Int J Travel Med Glob Health*. 2017;5(2):41-45.
10. Wiedman M. High altitude retinal hemorrhage. *Arch Ophthalmol*. 1975;93:401-3.
11. Karakucuk S, Mirza GE. Ophthalmological effects of high altitude. *Ophthalmic Res*. 2000;32:30-40.
12. Houston CS. High altitude illness. Disease with protean manifestations. *JAMA* 1976;236:2193–5.
13. [Montgomery AB, Mills J, Luce JM.](#) Incidence of Acute Mountain Sickness at Intermediate Altitude. *JAMA*. 1989;261(5):732-734.

14. Wieldman M, Tabin GC. High altitude retinopathy and altitude illness. *Ophthalmology*. 1999;106:1924-7.
15. Izadi M, Pourazizi M, Alemzadeh-Ansari MH. Ocular problems in high-altitude traveling: a review with focus on management. *International Journal of Travel Medicine and Global Health*. 2017 Jun 1;5(2):41-5.
16. Bosch MM, Barthelmes D, Merz TM, et al. High incidence of optic disc swelling at very high altitudes. *Arch Ophthalmol*. 2008;26:644- 50.
17. Report of World Health Organization/International Diabetes Federation Consultation: definition and diagnosis of diabetes mellitus and intermediate hyperglycemia. World Health Organization 2006; 1–50

18. Early Treatment Diabetic Retinopathy Study Research Group. Early photocoagulation for diabetic retinopathy: ETDRS report 9. *Ophthalmology*. 1981;98:766–785.
19. Bird AC, Bressler NM, Bressler SB; International ARM epidemiological study group. An international classification and grading system for age-related maculopathy and age-related macular degeneration. *Surv Ophthalmol*. 1995;39:367–374. doi:10.1016/S0039-6257(05) 80092-X
20. Schubert HD. Ocular manifestations of systemic hypertension. *Curr Opin Ophthalmol*. 1998;9:69–72. doi:10.1097/00055735-199812000-00012

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Thank you