

Ethnic Differences in Presentation and Outcomes for Cardiovascular Disease in North America

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Equity –stocks, options

none

This presentation; No conflicts

The Racial and Ethnic Diversity of the US Population Will Continue to Increase



US Census Bureau. US interim projections by age, sex, race, and Hispanic origin. 2004. Available at: http://www.census.gov/ipc/www/usinterimproj/. Accessed on December 21, 2005.

Ethnicity does not = Race

Race – Dorland's dictionary;

 An ethnic stock, or division of mankind. In genetics, races are considered as populations having different distributions of gene frequency.

Ethnic – Dorland's dictionary;

 Pertaining to a social group who share cultural bonds (religious, national, etc) or physical (racial) characteristics

American Anthropological Association

- "It has become clear that human populations are not unambiguous, clearly demarcated, biologically distinct groups....Any attempt to establish lines of division among biological populations is both arbitrary and subjective."

RISKS OF ETHNICITY RESEARCH

- Defining ethnic origin is not easy and likely to get more difficult in the future with increasing migration and inter-ethnic marriages
 - This in turn can affect the validity and applicability of research conclusions
- Ethnic labels could be used as a source of discrimination by health insurers and employers
 - On the other hand identification of common high risk gene pools (alleles) may in the future allow us to tailor medical therapy to those in whom the costbenefit ratio is most favorable

GEOGRAPHIC CHD DIVERSITY



Geographic Diversity in CHD Deaths By Ethnicity







Asians

Native Indians

Hispanics



← Blacks

Whites -



Geographic/Ethnic Disparity within Georgia



* Average annual age-adjusted rate (deaths per 100,000) for women ages 35 years and older. Data for Hispanics are also included within each of the four categories of race.

INTERHEART: Positive impact of lifestyle factors on acute MI for all ethnic groups



Yusuf S et al. Lancet. 2004;364:937-52.

VBWG

Asian Indian (South Asian) Population

1994 - 38 yr old South Asian malecoronary anatomy at presentation



Who are "South Asians"



AGE-ADJUSTED CV deaths declining in N America but rising in INDIA/CHINA



Visible Minorities in Canada and USA



CHD as Cause of Death in Asians



Don't Blame just the Diet!



Coronary Heart Disease in South Asians

- Two-fold increased risk of developing CHD for SA vs. Caucasians
- SA develop CHD and heart failure at younger ages
- More diffuse CAD at angiography, worse outcome following CABG
- Larger MI size, later presentation to hospital
- Approximately 50% higher CHD mortality
- Similar LDL levels, prevalence of HT, smoking similar or lower
- Lower HDL, higher prevalence of DM
- Conventional risk factors do not explain excess CHD

Gupta M et al. CMAJ. 2002 Deedwania P et al. Indian Heart J. 2005:57;617-631. Vallapuri S et al. Am J Cardiol. 2002;90:1147-50. Hamdoolay Z et al. Circulation. 2004 (AHA) Gupta M et al. Canadian Journal of Cardiology 2001;17(supp C):68C Singh et al. Ethnicity and Disease. 2005:15(4):615-619 Gupta M et al. Circulation. 2006:113;924-929

Prevalence of CVD for Given Levels of Carotid Atherosclerosis (IMT)



Non-Traditional Risk Factors

SHARE, Anand et al. Lancet 2000







South Asians and CAD

- Rapidly growing segment of US population
- Elevated CAD incidence in young adults
- High CAD risk compared to whites, with equivalent risk factors
 - Higher prevalence rates of insulin resistance
 - Metabolic syndrome
 - Diabetes
 - Elevated CRP
 - Lipoprotein (a) levels



Anand SS, et al. *Arterioscler Thromb Vasc Biol.* 2004;24:1509-1515.

McKeigue & Sevak. *Coronary heart disease in South Asian communities.* London: Health Education Authority, 1994. Anand SS, et al. *Lancet* 2000;356:279-284.

Environment, "Thrifty Genotype" and the Metabolic Syndrome



 Genes that convert and store simple sugars to abdominal fat
Selective advantage in times of famine



Low caloric intake High energy expenditure Low BMI and WHR



High caloric intake Low energy expenditure High BMI and WHR









Changes in Risk Factors with Migration



Racial and Ethnic Minority Groups Are More Likely to Have Type 2 Diabetes



McNeely MJ, Boyko EJ. *Diabetes Care.* 2004;27:66-69.

Racial and Ethnic Minorities Tend to Have Worse Glycemic Control



 * Includes individuals who reported all remaining single-race responses, individuals who reported more than one race but did not identify a major race, individuals with multi-races, and people with missing values on race.
Based on data from 757 patients with type 2 diabetes who participated in the National Health and Nutrition Examination Survey (NHANES) 1999-2002.

Fan T, et al. Diabetes Care. 2006. In press.

The Mortality Rate due to Diabetes Is Higher in Racial and Ethnic Minority Groups



Summary of Our Research

- South Asians have an increased risk of cardiovascular disease and a presentation profile similar to diabetics
- Aggressive risk factor modification can reduce morbidity and possibly mortality
- Insulin resistance / diabetes is a key factor in the development of diffuse atherosclerosis
- Early identification and intervention with evidence-based therapy can reduce morbidity and possibly mortality
- MY OPINION; Consider SA ethnicity + 1 risk factor as a CAD equivalent

ETHRISK CALCULATOR: www.epi.bris.ac.uk/CVDethrisk

10-year risk of CVD event (%) in 60-year-olds

Ethnic group	Male nonsmoker	Male smoker	Female nonsmoker	Female smoker
Indian	22.4	36.7	8.7	15.1
Pakistani	24.7	40.0	13.4	22.8
Bangladeshi	24.9	40.4	7.4	13.0
All South Asians	23.6	38.5	9.7	16.9
Chinese	11.1	19.1	2.5	4.4
Caribbean	13.3	22.7	9.7	16.9
Black African	20.1	33.3	10.8	18.7
Irish	16.0	27.0	9.1	15.8
Framingham score	14.1	22.3	8.7	14.7

Brindle P et al. *Heart* 2006; available at: http://heart.bmjjournals.com.



ASIAN BMI (Body Mass Index) Targets

- South Asians have a higher percentage of body fat and increased intra-abdominal fat compared to white Caucasians
- Insulin resistance is common in South Asians at BMI lower than 25
- South Asians develop type 2 diabetes at a younger age and at lower BMI compared to white Caucasians

Classification	Current BMI Target	Asian BMI Target	Risk of Co- Morbidity
Underweight	<18.5	<18.5	Low
Ideal Range	18.5-24.9	18.5-22.9	Average
Overweight	25 or more	23 or more	Increased
Obese I	30-34.9	25-29.9	Moderate
Obese II	35 or more	30 or more	Severe

IDF Diagnostic Criteria for Metabolic Syndrome

Central adiposity (waist circumference in cm)

Men Women

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- South Asian <90 <80
- Chinese <90 <80
- Japanese <85 <90

IDF 2004	
Central Adiposity (waist)	Plus 2 or more of:
Triglycerides	≥ 1.7 mmol/L
HDL cholesterol	
Men	<0.9 mmol/L
Women	<1.1 mmol/L
Blood pressure	≥130 or ≥ 85 mmHg
Fasting glucose	<u>></u> 5.6 mmol/L or DM or IGT

Challenges of Rx of risk factors in Ethnic populations

- Language barriers for information especially with 1st generation immigrants
- Lack of adequate insurance many selfemployed individuals
- Inappropriate dietary and lifestyle advice
- Reluctance to accept "Western medicine"
- Fatalistic attitude to life "reincarnation"

South Asian Dietary Advice - RESOURCES



Stanford University School of Medicine General Clinical Research Center



Various Recipes Available Latha Palaniappan, MD, MS Internal Medicine, Clinical Epidemiology Stanford Center for Research in Disease Prevention Phone: (650) 498-4427 E-mail: saiwwls@yahoo.com Indian Foods: AAPI's Guide To Health, Nutrition, and Diabetes









© American Association of Physicians of Indian Origin Photographs by Amish Thakkar; Cover page designed by Ranjita Misra

Resources- British Heart Foundation



NHS

Heart Disease and South Asians

Delivering the National Service Framework for Coronary Heart Disease



Improving access to treatment and services for South Asians

The National Service Framework for Coronary Heart Disease aimed to secure fair access to high quality services for all. As the NSF is implemented, facilities and services are being deliberately targeted at the areas which need them most to reduce inequalities in access to treatment and services.

Potential barriers to access

Some barriers are common to all communities: poor health, lack of time and absence of support may all influence people's ability and motivation to access services and lead a healthy lifestyle. Or practical problems, for example transport issues, may make it difficult for people to get to hospital. However, South Asian communities potentially face a number of additional barriers that service providers need to be awere of and address.



CASE STUDY

CASE STUDY Khush Dil "Happy Heart" Project

The team offers the following services:

The Coriander Club, Spitalfields City Farm



women attending have diabetes and high blood pressure. The class looks at ways of cooking more healthily for example ideas on how to limit fat and sait, and ways to eat five portions of fuilt and vegetables a day. The women cook traditional Bangladeshi dishes and have also theid dishes from other cuisines, for example Mediterranean dishes, as well as dishes to appeal to children and grandchildren. The dass ta bo used to helo

RESOURCES – Clinician Update

 Gupta M, Singh N, Verma S.
South Asians and Cardiovascular Risk: What Clinicians Should Know
Circulation 2006;113 924-929

South Asians and Cardiovascular Risk What Clinicians Should Know

CLINICIAN UPDATE

Milan Oupia, MD; Narendra Singh, MD; Suboth Vernat, MD, PhD

Sprawpoint A 36-marchi meanolisy, sommanive South Asian case greented to the emergency department of a commuaite Impliti with rebosienzi thest pain cf will minister damains. No 12-bad SITS descentrated 10 mm of 5Tseptent elevatos in fields V, through V, and he received libring tic theory within 90 minutes of symptom cased His cale marined, but his ST seemesh cole partially contrained he had a cask creates house of \$565 U.L. and he show of signs of early haat follows 1.00. cholesteral was 135 angld1, HDL 32 agit1, tiglyonide 21 agit1, ad thi chicketeral 206 aug/d.; his body man inder (SMI) wur 24 ingint. Cardan collectronics depundented serves and diffuse triple-wavel disease, including occlusion of the proximal left minut? decreasing sales, in well in moderate left vestricular distriction. While in the loaded, he was disasted with anycost ince I datates galling and salseptently and event tarm opticated outcomy by put wagers. Seath Arlens and

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thei includer India, Pakielan, Sci-Lasta Nepil and Regulateth Collec-Undy, South Asiana represent sub 15th of the global population. In North America, more than 2 million South Aniant reside in the United String and simplei 1 cultion in Change 3 is importial to recognize that the local "Sinch Anias" other to a believag secon population, with important offferences in det, mitten, and Medvie nance different South Anian providtings and reliations. Multiple studies of migrati Sculi Asian populatinas hava, however, coeffemed a 3- to 5-frid increase in the drit for asymptotic isterios and continuacelar deals or compand with other ethnic groups."* là manipa d' que ministrat concern beef disease (CSD) cardatite to Cataly over a 15-year second South Asian had the highlet CRD carcality concept with influiture of Chinese and Environm descent.» In addition, Smith Anima are prove to developing CRD at a promper age, obta before the age of all years in man.² Case-control sitis have three that compared with while, South Asian is Canada present to the banging interior the connect scale meteorial infection and an more

the time of codice constant rules that while yet are more likely to have decodenative main, endineered, and datal commeny strary disease." In minimo, South Arians for disalimently younger is the time of the "implicit rules for heart induces

Traditional Cardiac Risk Factors

Risk Factors The INTERNEART study demosstated the validout codimacular Genne risk factors day an imported rate is the predictor of any contral induction in populations around the world, including forth Asians, Rowway comercial case-coaird singles decompating pression CSD is South Asian decreation similar or lower prevalence of institutal cirk factors that with other populations.""" A re-view of cross-sectional data from the United Eingdom, iarbufag the 1999 Realth Survive of Realand, owneds that the coordinate of Incontention in the the in South Ashier and the white population was Tobacco and it gener-Ely low manag South Asian cash and strictst antipued of second South Asian wannen.» Tebarro coommptan is rapidly iscressing is South Asian consider in conjunction with ecocontric expensions.

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African American and Hispanic Populations

Risk of Cardiovascular Disease

Death Rates per 100,000 Persons Among US Ethnicities



American Heart Association. 2006. Heart Disease and Stroke Statistical Update.

Yrs of potential life lost

Circulation 111: 2005

- Figure 4. Years of potential life lost (YPLL) before 75 years of
- age resulting from diseases of the heart, ischemic heart disease,
- and stroke, United States, 2001. API indicates Asian or Pacific
- Islander; AIAN, American Indian or Alaska Native. Source: CDC,
- Health United States, 2003.


Clinical Study Examples - LIPIDS

Study	Study Aim	Overall n	% Minority Subjects	Major Results	Subgroup Results
CARE, 1996	Effect of pravastatin		7-8%	Fatal coronary event + nonfatal MI RRR=24%	
Cholesterol and Recurrent Events Trial	on fatal/nonfatal coronary events; MI and 115 <ldl- C<174mg/dL</ldl- 	4159	non-white participants	Need for CABG RRR=26% Stroke BRR=31%	
LIPID, 1998 Long-term Intervention with Pravastatin in Ischemic Disease	Effect of pravastatin on coronary/all cause mortality in CHD patients	9014		Coronary death RRR=24% Overall mortality RRR=22%	Women: Coronary mortality + non-fatal MI RRR=11% Age ≥ 70 yrs: Coronary mortality + non-fatal MI RRR=15%



Clinical Study Examples - HYPERTENSION

Study	Study Aim	Overall n	% Minority Subjects	Major Results	Subgroup Results
ALLHAT, 2002 Antihypertensive and Lipid Lowering Treatment to Prevent Heart Attack Trial	Effect of CCB or ACEi vs. diuretic CHD	33,357	W-47 B-32 Hispanic W-12.5 Hispanic B-3.3 Other-5	No difference in outcomes Thiazide type diuretics are superior in preventing CVD	Blacks: Heart Failure RR=1.47 (favors Cholthlidone)
MRFIT, 1985 Multiple Risk Factor Intervention Trial	Special intervention vs. Usual care	11,342	B-7.2	No favorable association between lowering BP and CHD rate	

Clinical Study Examples – CONGESTIVE HEART FAILURE

Study	Study Aim	Overall n	% Minority Subjects	Major Results	Subgroup Results
EPHEHSUS, 2005 Eplerenone Post Acute MI Heart Failure Efficacy and Survival Study	Impact of eplerenone on mortality (30 days post MI LVEF clinical signs of heart failure)	6632	1%	All cause mortality RRR= 31% CV mortality RRR=32% Sudden cardiac death RRR=37%	No sub-analysis of minority subjects Women: all cause mortality RR~ 0.5
HOPE, 2000 Heart Outcomes Prevention Evaluation	Role of ramipril (In patients at high risk for cardiovascular events, without LV Dysfunction or heart failure)	9297		CV death RR=0.74 MI RR=0.80 Stroke RR=0.68 All cause mortality RR=0.84 Heart failure RR=0.77	Women: MI + stroke + cardiovascular death RR ~ 0.8



The Jackson Heart Study (2000 – Present)

- Single site, prospective epidemiologic study of cardiovascular disease among African-Americans in Jackson, Mississippi
- 6500 African American subjects
- Largest investigation of CVD that has ever been undertaken in African Americans
- Questionnaires: lifestyle habits, medical history, medications, social/cultural factors
- Physical Assessment: weight, height, body size, BP, EKG echocardiogram
- Labs: lipids, glucose, hemostatic factors



The Strong Heart Study (1988-1992)

- Population based, epidemiologic, longitudinal study
- Studies CVD and pulmonary disease and their risk factors
- 4,549 Native Americans in Arizona, Oklahoma and North and South Dakota
- Included personal interview, physical exam, BP measurements and labs (OGTT)



The San Antonio Family Heart Study (1991- Present)

- Studies cardiovascular and diabetes and their risk factors
- 1,400 members of > 40 large Mexican American families
- Risk factor data collected by questionnaire plus physical exam and phlebotomy for genetic studies
- Two follow-up examinations through 2005



The Atherosclerosis Risk In Communities (ARIC) (1987-1998)

- Prospective population based, epidemiologic, study
- Studies CHD, atherosclerosis, stroke and risk factors
- 15,792 participants from 4 U.S. communities, including 12% Blacks in the Forsyth County, NC site
- Included personal interview, risk factor determination, physical exam, BP, BMI and waist circumference measurements, labs (HDL-C, TG, glucose levels)
- Echocardiographic studies were preformed in the Jackson site between 1993-1996



Multi-Ethnic Study of Atherosclerosis (MESA) (2000-2002)

- Studies prevalence, treatment and control of dyslipidemia
 - A multi-center cohort
 - 6814 participants, free of CVD at baseline, 28% black, 12% Chinese, 22% Hispanic
 - Participants were evaluated for CVD risk, CT (CAC) and fasting lipid profiles



The Dallas Heart Study (2000-2002)

- Single-site population based probability sample of Dallas residents
- Aim: to evaluate ethnic differences in cardiovascular health
- Included survey questionnaires, labs and imaging studies
- 6101 subjects, 50% Black

Ethnic differences in risk factors in women 25-64 years old, NHANES 1988-1994

Benjamin, E. J. et al. Circulation 2005;111:e124-133e Circulation

Ethnic differences in risk factors in men 25-64 years old, NHANES 1988-1994



JNC 7- Minority Populations

- In general, treatment similar for all demographic groups.
- Socioeconomic factors and lifestyle important barriers to BP control.
- Prevalence, severity of HTN increased in African Americans.
- African Americans demonstrate somewhat reduced BP responses to monotherapy with BBs, ACEIs, or ARBs compared to diuretics or CCBs.
- These differences usually eliminated by adding adequate doses of a diuretic.

BEST Trial ; Effect of Bucindolol on Mortality by Race



Eichhorn EJ et al. Circulation. 2000;102(Suppl):II-778. Abstract #3759. HF NOT AN APPROVED INDICATION

US Carvedilol Trials: Effect of Race on Death or Hospitalization for Any Cause



Yancy CW et al. N Engl J Med. 2001;344:1358–1365.

SOLVD Studies: Racial Differences in Mortality Rates



*Mortality rates include treatment and placebo arms. Adapted from Dries DL et al. N Engl J Med.1999;340:609–616.

V-HeFT II Retrospective Analysis: Effect on All-Cause Mortality in Black Patients

Retrospective analyses are hypothesis generating only



V-HeFT I Retrospective Analysis: Survival Benefit in Black Patients

Retrospective analyses are hypothesis generating only



A-HeFT: Study Design

- African American Heart Failure Trial (A-HeFT)¹
 - Hypothesis: Fixed-dose of ISDN/HYD will improve outcomes in black patients with moderate-to-severe symptomatic HF
- 169 sites²
- 1050 randomized patients (518 BiDil[®], 532 placebo)^{1,2}
- Up to 18 months of follow-up^{1,2}
- No patient lost to follow-up for vital status¹
- Study initiated 6/12/01, and terminated early due to significant survival benefit in the BiDil[®] group^{1,2}

ISDN/HYD=isosorbide dinitrate/hydralazine.

- 1. Taylor AL et al. N Engl J Med. 2004;351:2049-2057.
- 2. BiDil[®] Prescribing Information.

A-HEFT Trial - Kaplan-Meier Estimates of Overall Survival



The NEW ENGLAND JOURNAL of MEDICINE

Disparities in Cardiovascular Care

Ethnic differences in Medicine

today in **CARDIOLOGY**

Focus on Minorities and Heart Disease

Increased awareness crucial to CVD prevention in minority patients

Minority patients are less likely to be screened for CVD risk factors and less likely to receive risk-reducing treatments.

Evidence of Racial/Ethnic Differences in Cardiac Care, 1984-2001



11 studies find no racial/ethnic difference in care (14%)

2 studies find racial/ethnic minority group more likely than whites to receive appropriate care (2%)

Total = 81 studies



CRUSADE/NSTE ACS: In-Hospital Procedures



Sonel EF, et al. *Circulation*. 2005;111:1225-1232.

CRUSADE/NSTE ACS Recommendations at Discharge



* For current smokers.

[†] For patients with positive markers.

Sonel EF, et al. Circulation. 2005;111:1225-1232.



TACTICS-TIMI 18: Influence of Race in Non-ST Elevation ACS

Nonwhite patients vs. white patients

- Less likely to be
 - Taking aspirin
 - Taking beta-blockers
 - Taking statins
 - Referred for coronary angiography in the absence of clear-cut ischemia



TACTICS-TIMI 18: Influence of Race in Non-ST Elevation ACS

Nonwhite patients vs. white patients

- Less likely to receive a stent when undergoing percutaneous coronary revascularization
- Less successful procedural outcomes
- Regardless, after adjustment for differences in rates and types of revascularization, nonwhite patients still had a worse prognosis



The Effect of Race and Sex on Physicians' Recommendations for Cardiac Catheterization "Patients" Experiencing Symptoms of Heart Disease



Results: Referral for cardiac catheterization according to race

	Mean Referral Rate %	Odds Ratio (95% CI)	P value
White	90.6	1.0	
Black	84.7	0.6 (0.4-0.9)	0.02

Rate of Use of Cardiac Catheterization within 60 Days after Acute Myocardial Infarction among Black Patients and White Patients, According to the Race of Their Physicians

 TABLE 2. RATE OF USE OF CARDIAC CATHETERIZATION WITHIN 60 DAYS

 AFTER ACUTE MYOCARDIAL INFARCTION AMONG BLACK PATIENTS AND WHITE PATIENTS,

 ACCORDING TO THE RACE OF THEIR PHYSICIANS.*

Rate	Wнге	Physicians	BLACK PHYSICIANS		
	WHITE PATIENTS $(N=35,176)$	BLACK PATIENTS $(N=3476)$	WHITE PATIENTS $(N=500)$	BLACK PATIENTS (N=563)	
		percent (95 percer	nt con fiden ce in terval)		
Unadjusted	45.7	38.4†	49.6	38.2†	
Adjusted	45.7‡	32.9 (30.1 36.1)§	53.4 (43.4-65.8)	36.5 (29.2 45.2)¶	

*In the adjusted analyses, adjustments were made for the characteristics of the patients, the physicians, and the hospitals. There were no significant differences in the rates of cardiac catheterization among either the white patients or the black patients according to the race of their physicians.

[†]P<0.001 for the comparison between black patients and white patients, regardless of the race of their physician.

‡These patients served as the reference group.

§P<0.001 for the comparison between black patients treated by white physicians and white patients treated by white physicians.

P=0.04 for the comparison between black patients treated by black physicians and white patients treated by black physicians.



LESS REVASCULARIZATION, MORE ANGINA



Figure 2. Revascularization (PCI and/or CABG surgery) status among blacks (n=295) and whites (n=1367) during the initial catheterization hospitalization and at 6 months (overall and by presence of single-vessel/multivessel disease). *P* values correspond to χ^2 tests of revascularization rates among black and white patients.

Circulation 111: 2005



Figure 3. Angina symptoms at baseline and 6 months among black (n=295) and white (n=1367) patients (overall and by presence of single-vessel/multivessel disease). *P* values correspond to χ^2 tests of revascularization rates among black and white patients.

Weight of the Evidence

- African Americans less likely than whites to receive catheterization, angioplasty, bypass surgery and thrombolytic therapy
- Racial/ethnic differences in care remain after adjustment for clinical and socioeconomic factors
- Evidence of disparities in other procedures and treatments mixed
- Limited data on Latinos, Asians and Native Americans

Proportion responding that they believe that within the healthcare system in general, patients receive different care on the basis of the factors shown (n=344)

Proportion responding that they believe that clinically similar patients receive different care on the basis of race/ethnicity by proximity to practice (n=344)



Percentage reporting that specific patient characteristics contribute a great deal to racial/ethnic disparities in cardiovascular care in the United States



Lurie, N. et al. Circulation 2005;111:1264-1269

Percentage reporting that specific nonclinical factors contribute a great deal to racial/ethnic disparities in cardiovascular care in the United States



Percentage achieving goals at 1-year follow-up by intervention group (community based vs standard clinic)



Becker, D. M. et al. Circulation 2005;111:1298-1304

Potential Primary Prevention Implantable Cardioverter-Defibrillator (ICD) Patients Lack Understanding, Want More Information and are Willing to Reconsider Implant Refusal

N Singh, E Hall, E Lucas, S Chandra, D Song, L Yan, D Suh, A Dorsey, S Ramamurthy



Summary


Back up Slides