**Cohort Profile: Regional Ethnic Cohort Study in Northwest China**

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**Key Features**

* The Regional Ethnic Cohort Study in Northwest China (RECS), a community, population-based, prospective observational study, aims to investigate specific aetiological causes and long-term health hazards of non-communicable diseases (NCDs) among different ethnic groups in the northwest region of China.
* A total of 118,572 participants aged 35–74 years from five provinces of Northwest China, including people from the Han, Uygur, Kazakh, Hui and Tibetan ethnic groups, were recruited between June 2018 and May 2019.
* Approximately 10% of the participants will be actively followed every 3 years via face-to-face interviews, using concise questionnaires to review risk exposures and disease incidence, and with collection of biological specimens, including blood, saliva and stool samples. Passive follow-up will be conducted by periodic linking (every 6 months) of baseline survey data to established electronic disease registries.
* The questionnaire survey, regular medical examination and collection and storage of blood samples were conducted in the RECS baseline survey for all participants. Approximately 40% of the participants had blood and urine samples assessed for several biochemical indices.
* Overweight/obesity and hypertension have become a prominent public health problem in Northwest China, especially in the Uygur ethnic group. Low prevalences of self-reported diabetes and cardiovascular disease among several ethnic groups (other than Han ethnicity) indicate lower levels of access to health services and chronic disease management among the regional ethnic minorities of Northwest China.
* Collaborations from all over the world are welcome. Please contact the corresponding author at tjdshn@xjtu.edu.cn with specific research ideas or questions.

**Why was the cohort set up?**

Non-communicable diseases (NCDs) contribute to more than two-thirds of deaths worldwide. Approximately 80% of NCDs occur in low- and middle-income countries, and 33% of the deaths from NCDs are in people younger than 60 years (1-4). Although great efforts have been made to control NCDs in China, their prevalence continues to increase (5-8). The prevalence of NCDs in China reportedly increased from 15.3% in 2003 to 29.5% in 2013(9). The burden of NCDs and associated healthcare costs is rising for older people with chronic diseases who require long-term medical services and support. Patients with NCDs face the risk of disability and life-threatening conditions that result from complications of NCDs because of a lack of medical service resources and coverage by healthcare systems (5, 9).

The region of Northwest China (31°–50°N, 73°–111°E) is considered to be relatively poorer (in financial terms) than the central and eastern provinces. It includes the provinces of Shaanxi, Gansu and Qinghai, and the autonomous regions of Ningxia Hui and Xinjiang Uygur (10). The region’s topography is highly varied and includes large stretches of arid desert, wastelands, fertile oases, grassy plateaus and high mountain ranges. These provinces face various environmental issues because of decreased water availability and an arid climate (11, 12). Northwest China is also one of the most ethnically diverse areas, with 17 ethnic minorities and two of the five autonomous regions that are designated minority areas. Lifestyle and dietary habits differ between ethnic groups, and high rates of smoking and drinking in many ethnic populations further increase the risk of NCDs (13-17).

China has experienced tremendous changes in the past 30 years, during which time its rapid economic development has substantially influenced environmental exposure (severe air pollution), economic inequalities among regions and ethnicities, and diet and lifestyle changes in the Chinese population. These changes are also thought to have contributed to NCD development and mortality (18-20). To address the urgent need for data on the prevalence of and risk factors for NCDs and associated conditions in resource-constrained settings, the Regional Ethnic Cohort Study in Northwest China (RECS), supported by the National Key R&D Program of China (NKRP), aimed to establish a nationally representative cohort, including people from different ethnicities and areas. The RECS, as one important part of the NKRP, will provide a reliable source of data for studies investigating specific aetiological causes and long-term health hazards of NCDs among different ethnic groups in Northwest China. The findings from studies using the cohort data will serve to promote the health conditions of the local population and to provide references for other countries to formulate prevention and control strategies for NCDs.

**Who is in the cohort?**

This cohort consists of people in 13 areas of five western provinces of Northwest China (Xinjiang: Moyu county, Yining city, Urumchi city; Gansu: Wuwei city, Tianshui city, Gannan city, Baiying city; Qinghai: Xining city; Ningxia: Qingtongxia city, Pingluo city; Shaanxi: Zhenba county, Baoji city, Xi’an city). The areas were chosen according to local disease patterns, exposures to certain risk factors, distribution of ethnic groups, population size and stability, quality of death and disease registries, and local commitment and capacity (Figure 1). The cohort was planned to include 110,000 participants and is managed by each provincial project group (Xinjiang: Xinjiang Medical University; Gansu: Gansu University of Chinese Medicine; Qinghai: Qinghai Center of Disease Prevention and Control; Ningxia: Ningxia Medical University; Shaanxi: Xi’an Jiaotong University). Each group is responsible for the baseline survey measurements, including the questionnaire interview, blood sample collection and storage, and further follow-up work. The Shaanxi project team operates as the general manager in charge of the project. The baseline survey was conducted between June 2018 and May 2019.

After selecting the sites for inclusion in the cohort, the local community health centers and township hospitals advertised the study to the residents they served to recruit the participants. Permanent residents who met the inclusion criteria were not formally invited but could volunteer to participate, to help ensure the compliance of participants. After the target sample size for each village/community was reached, the local survey ended. A neighbouring village/community was used if the sample size in the first village/community was not reached. Participants were asked to show their national identity cards before entering the cohort, and only those whose address on the identity card showed them to be a resident of a cohort site were included, as we were unable to follow up people who were not permanent local residents.

The selection of sites was based on the population size and ethnic distribution of each province (21). Therefore, the target sample sizes were 40 000, 30 000, 15 000, 20 000 and 5000 in Shaanxi (the largest proportion of the population), Xinjiang (multi-ethnic gathering area), Ningxia (gathering area of Hui ethnic), Gansu and Qinghai provinces, respectively. As the China Multi-Ethnic Cohort (CMEC), which is also part of the NKRP, investigated Tibetans in Tibet, we reduced the sample size in Qinghai province and focused on Han-ethnic participants living on the plateau, and enlarged the sample size in Shaanxi province to ensure that we reached the total target sample size of 110 000.

A baseline survey was conducted via face-to-face interviews and clinic visits, with assistance from local health staff. The participants in the cohort consisted of permanent residents aged 35–74 years, but any individuals attending the baseline survey who were slightly outside the target age range were not turned away, to encourage participation. We excluded people who had severe mental disease or disabilities that hindered their ability to communicate, so we could obtain accurate data. All participants gave written informed consent, which allowed us to access their medical records and long-term storage supplies of blood samples, solely for medical research purposes.

**How often will participants be followed up?**

The National Death Surveillance System (NDSS), Chronic Disease Registries (CDR), National Central Cancer Registry (NCCR) and National Health Insurance Datasets (NHID) enable us to conduct passive follow-up by periodic linking (every 6 months) of the baseline survey data to each of these datasets. The NDSS and CDR are both based on China’s disease surveillance points system. The causes of death of cohort participants are regularly monitored through the use of official death certificates that are reported to the regional Centers for Disease Control and Prevention (CDC) where the NDSS is based (the NDSS is available in all 13 study areas). Information on chronic diseases (e.g. diabetes, hypertension, heart failure, stroke and cancer) is being collected through linkage with the NCCR, CDR and NHID. The new NHID are now fully established in all 13 study areas, and future information on incidence of chronic disease and hospitalization will be primarily extracted from these datasets (22-26). As the NCCR and CDR are still under construction, their coverage of the sites is limited, so they are not primary data sources but are used as complementary sources for passive follow-up. The unique national identity card and health insurance card numbers were used as key variables to link the baseline data to the NDSS, CDR and NHID.

The diagnoses of chronic diseases are based on well-accepted international standards. Causes of death are coded according to the 10th version of the International Statistical Classification of Diseases (ICD-10) (27). In addition, we finished one to two rounds of passive follow-up (varying between sites), and 534 deaths and 3083 new cases of cardiovascular disease (CVD) during 2019–2020 were identified.

Approximately 10% of the participants will be actively followed up every 3 years via face-to-face interviews, to estimate changes in lifestyle and exposure factors and to identify participants who have permanently moved out of the study areas, as well as to minimize the under-reporting of vital status and death data. Biological specimens, including blood, saliva and stool samples, are also planned to be collected. We aim for the rate of loss of individuals to follow-up to be less than 10%.

**What has been measured?**

The types of measures are summarized in Table 1. In brief, a questionnaire survey, physical examination, biological sample collection and laboratory testing were conducted. The entire visit procedure, including the electronic questionnaire interview, medical examination and sample collection, typically took 60–75 min for each participant to complete.

**Questionnaire survey**

The face-to-face questionnaire interview collected baseline information about the participants, including demographic and socio-economic characteristics, lifestyle factors, environmental exposures, medical history, mental health status and, for women, reproductive history (Table 1). The questionnaire referred to the China Kadoorie Biobank as a reference (28), and validated instruments for food frequency (Food Frequency Questionnaire [FFQ]) and quality of life (12-item short form survey [SF-12]) were used (28-30). A tablet computer with a self-developed application (the RECS App) was used to collect the questionnaire information at face-to-face interviews performed by well-trained technicians. To ensure the quality of the data, interviewers were trained in the standardized process and mode of query to be used.

**Medical examination and sample collection**

After the questionnaire survey, 10 mL venous blood samples and anthropometric measurements were taken for all participants by professional physicians or nurses at the local clinical centre, with participants asked to fast overnight for at least 8 hours beforehand. The doctors, nurses and technicians received standardized training before conducting the examinations. For all participants, the medical examination included measurement of height, weight, waist circumference (WC), heart rate, body fat percentage, visceral fat index, basal metabolic rate, bone mass, muscle mass, total body water and resting blood pressure. Each project group (the provincial CDC or a local medical college) had primary responsibility of quality control, as guided by the overall protocol of the study. The measurements obtained in the medical examinations were estimated to have optimal consistency among the devices that were used in the five provinces. In addition, blood tests (e.g, liver and kidney function, blood lipids, fasting blood glucose and glycosylated haemoglobin, high-sensitivity C-reactive protein), urine tests, lung function tests and B-ultrasonic examinations were selectively used in approximately 40% of the participants.

**What has it found?**

The provinces and autonomous regions in Northwest China largely comprise China’s inland poverty belt, with a total land area accounting for 33.3% of the national land area (13). The population density in the areas suitable for human habitation exceeds the critical level of population density in arid and semi-arid areas (as designated by the United Nations), and the distribution of Gross Domestic Product (GDP), life expectancy and forest coverage is unbalanced (15) (Table 2).

Table 2 shows the socio-economic characteristics of the participants (in total and in each of the five provinces). Among the total 118,572 participants, the proportions of Han, Uygur, Kazakh, Hui and Tibetan ethnicities were 75.3%, 13.0%, 1.7%, 8.2% and 1.3%, respectively. Participants in the provinces of Gansu, Shaanxi and Qinghai were primarily of Han ethnicity (over 90%), with a small proportion of people of Tibetan ethnicity in the Gansu (7.2%) and Qinghai (1.3%) provinces. In Xinjiang province, 67.2% of participants were of Uygur, Kazakh or Hui ethnicity, and in Ningxia province, 37.3% of participants were of Hui ethnicity.

Table 3 shows lifestyle and self-reported medical history factors across the different ethnic groups. Kazakh (30.0%) and Tibetan participants (19.9%) had higher proportions of current smokers. There were also higher proportions of participants who usually drank alcohol in the Han (9.2%) and Tibetan (8.2%) ethnic groups. The participants from the Uygur, Kazakh, Hui and Tibetan ethnic groups consumed more animal-based food (but less salt). The overall prevalences of self-reported diabetes and CVD were low in the baseline survey, and extremely low among several ethnic groups (other than Han). This may be because there is less access to health services and chronic disease management.

Based on the Working Group on Obesity in China (WGOC) criteria (31), the prevalence of overweight (body mass index [BMI], 24 to <28 kg/m2) and obesity (BMI, ≥28 kg/m2) was 37.2% and 16.1%, respectively. Participants from the Uygur, Kazakh and Hui ethnic groups had a higher prevalence of central obesity (WC >85 cm in men or >80 cm in women) compared with those in the Han ethnic group. Even among participants with normal BMI, the prevalence of central obesity, especially in the Uygur ethnic group, was high. The prevalence of hypertension (as defined by the Chinese guidelines for prevention and treatment of hypertension (32)) was higher in the Hui, Tibetan and Han ethnic groups (>30%), and the total prevalence of hypertension was about 30.2% (Figure 2). These results suggest that overweight/obesity and hypertension have become a prominent public health problem in Northwest China.

Current projects include analyzing the predictors of CVD and cost-effectiveness of CVD preventive health policy by using baseline data as well as a study of 450 plasma samples exploring the metabolic profile and possible metabolic path to normal weight obesity. Additionally, another study is underway to construct a haplotype reference sequence set of different ethnicities of northwestern China via high-throughput sequencing based on 1000 ethnic samples of DNA.

**What are the main strengths and weaknesses?**

There are several limitations of this cohort. First, information on personal history of diseases, physical conditions and some lifestyle factors (such as smoking, drinking and dietary food intake) was self-reported, and recall bias could not be avoided. Second, we only included participants in the age range of 35–74 years, which may have excluded information on early-life exposures. Third, due to limited funding, only 40% of participants were tested for biochemical indices. Fourth, no information on non-responders was collected, so it is possible that bias might exist in our sample.

As a unique, large-scale cohort in Northwest China, our study also has some strengths. First, because of the specific features regarding lifestyle, dietary habits, climate, geography and rapid social and economic transitions in Northwest China, this cohort will provide important data and support to estimate the prevalence and risk factors for NCDs in Northwest China, which will allow us to further investigate the aetiology of NCDs. Second, this cohort comprises multi-ethnic individuals, urban and rural inhabitants, plateau and basin residents, populations living in remote mountain areas and those living in highly concentrated and air-polluted areas. These features allow us to evaluate the effects of various environmental exposures on different health-related outcomes. Third, many blood samples (approximately 0.88 million), including whole blood, plasma, serum and buffy coat samples, were collected at baseline. These valuable samples allow us to obtain information on the genetic and epigenetic variants for health outcomes. In addition, genetic information from multi-ethnic individuals, combined with abundant personal and environmental information (including specific lifestyle and dietary habits that were collected at baseline), provides an important opportunity to reveal complex and specific mechanisms of the genetic and environmental factors associated with NCDs. Finally, the information on follow-up outcomes, including the diagnosed diseases that are linked to this cohort database through the NDSS, CDR and NHID systems, enables the reliable and real-time confirmation of participants’ health status and the tracking of medical histories.

**Can I obtain the data? Where can I find out more?**

The RECS welcomes collaboration throughout the world to maximize the use of these data. Currently, the data cannot be downloaded publicly because they contain sensitive information. However, possible collaborators are invited to contact the corresponding author (Shaonong Dang: tjdshn@xjtu.edu.cn).

**Ethics approval**

The study was approved by the Human Research Ethics Committee of the Xi’an Jiaotong University Health Science Center (No: XJTU2016-411).

**Author Contributions**

H Yan and S Dang had full access to all of the data in the study and take responsibility for the integrity of the data and the accuracy of the data analysis.

*Concept and design:* H Yan, S Dang,C Li, L Zeng, Q Li and Y Shen.

*Acquisition, analysis, or interpretation of data:* All authors.

*Drafting of the manuscript:* C Li, Y Liu and S Dang.

*Critical revision of the manuscript for important intellectual content:* All authors.

*Statistical analysis:* C Li, Y Liu, G Shi and S Dang.

*Administrative, technical, or material support:* G Shi, B Mi, Y Zhao, L Pei, Y Kang, S Xiao, X Liu, J Dai, Y Zhang, XH Wang, XJ Wang and D Wang.

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**Conflict of interest**

None declared.

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**Figure 1** Spatial distribution of participants in the baseline survey of the Regional Ethnic Cohort Study in Northwest China



**Figure 2** Prevalence of hypertension, obesity, overweight and central obesity among different regional ethnic groups. Central obesity was defined as waist circumference of more than 85 cm in men or 80 cm in women. Obesity was classified as body mass index (BMI) >28 kg/m2; overweight as BMI of 24.0 to <28 kg/m2; and normal weight as BMI of 18.5 to 23.9 kg/m2.